

Sage Therapeutics, Inc.
Form S-1
April 07, 2015
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As filed with the Securities and Exchange Commission on April 6, 2015.

Registration No. 333-

UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
WASHINGTON, D.C. 20549

FORM S-1
REGISTRATION STATEMENT
UNDER
THE SECURITIES ACT OF 1933

SAGE THERAPEUTICS, INC.

(Exact Name of Registrant as Specified in Its Charter)

Delaware
(State or Other Jurisdiction of

2834
(Primary Standard Industrial

27-4486580
(I.R.S. Employer

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Incorporation or Organization)

Classification Code Number)

Identification Number)

215 First Street

Cambridge, Massachusetts 02142

(617) 299-8380

(Address, Including Zip Code, and Telephone Number, Including Area Code, of Registrant's Principal Executive Offices)

Jeffrey M. Jonas, M.D.

President and Chief Executive Officer

Sage Therapeutics, Inc.

Cambridge, Massachusetts 02142

(617) 299-8380

(Name, Address, Including Zip Code, and Telephone Number, Including Area Code, of Agent For Service)

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Boston, Massachusetts 02109	(617) 299-8380	(617) 951-7000
(617) 570-1000		

Approximate date of commencement of proposed sale to public: As soon as practicable after this Registration Statement becomes effective.

If any of the securities being registered on this Form are to be offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act of 1933, check the following box:

If this Form is filed to register additional securities for an offering pursuant to Rule 462(b) under the Securities Act, please check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering:

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If this Form is a post-effective amendment filed pursuant to Rule 462(c) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering: "

If this Form is a post-effective amendment filed pursuant to Rule 462(d) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering: "

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, or a smaller reporting company. See the definitions of "large accelerated filer," "accelerated filer" and "smaller reporting company" in Rule 12b-2 of the Exchange Act.

Large Accelerated Filer Accelerated Filer
Non-Accelerated Filer (Do not check if a smaller reporting company) Smaller Reporting Company

CALCULATION OF REGISTRATION FEE

Class of Securities to be Registered	Proposed Maximum Aggregate Offering Price(1)	Amount of Registration Fee(2)
Common Stock, par value \$0.0001 per share	\$115,000,000	\$13,363

- (1) Includes shares that the underwriters have the option to purchase.
(2) Calculated pursuant to Rule 457(o) based on an estimate of the proposed maximum aggregate offering price.

The registrant hereby amends this registration statement on such date or dates as may be necessary to delay its effective date until the registrant shall file a further amendment which specifically states that this registration statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933 or until the registration statement shall become effective on such date as the Securities and Exchange Commission, acting pursuant to said Section 8(a), may determine.

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The information in this preliminary prospectus is not complete and may be changed. We may not sell these securities until the registration statement filed with the Securities and Exchange Commission is effective. This preliminary prospectus is not an offer to sell these securities and it is not soliciting an offer to buy these securities in any jurisdiction where the offer or sale is not permitted.

Subject to Completion, dated April 6, 2015.

Preliminary prospectus

Shares

\$100,000,000

Common Stock

We are selling _____ shares of common stock.

Our common stock is listed on The NASDAQ Global Market under the symbol SAGE. The closing price of our common stock on The NASDAQ Global Market on April 2, 2015, was \$46.87 per share.

We are an emerging growth company as that term is used in the Jumpstart Our Business Startups Act of 2012, and, as such, we have elected to take advantage of certain reduced reporting requirements for this prospectus and may elect to comply with certain reduced public company reporting requirements for future filings.

	Per Share	Total
Public offering price	\$	\$
Underwriting discounts and commissions ⁽¹⁾	\$	\$
Proceeds to Sage Therapeutics, Inc. before expenses	\$	\$

(1) See Underwriting beginning on page 159 for additional information regarding underwriting compensation.

We have granted the underwriters an option to purchase up to _____ additional shares of our common stock at the offering price less the underwriting discount. The underwriters can exercise this right at any time within 30 days after the date of this prospectus.

Investing in our common stock involves a high degree of risk. See Risk Factors beginning on page 11.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or determined if this prospectus is truthful or complete. Any representation to the contrary is a criminal offense.

The underwriters expect to deliver the shares of common stock to purchasers on or about _____, 2015.

J.P. Morgan

Goldman, Sachs & Co.

Leerink Partners

Cowen & Company

Prospectus dated _____, 2015

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We and the underwriters have not authorized anyone to provide any information other than that contained in this prospectus or in any free writing prospectus prepared by or on behalf of us or to which we have referred you. We take no responsibility for, and can provide no assurance as to the reliability of, any other information that others may give you. We and the underwriters are not making an offer to sell these securities in any jurisdiction where the offer or sale is not permitted. You should assume that the information appearing in this prospectus is accurate only as of the date on the front cover of this prospectus. Our business, financial condition, results of operations and prospects may have changed since that date.

For investors outside of the United States: We have not, and the underwriters have not, done anything that would permit this offering or possession or distribution of this prospectus in any jurisdiction where action for that purpose is required, other than the United States. Persons outside of the United States who come into possession of this prospectus must inform themselves about, and observe any restrictions relating to, the offering of the shares of common stock and the distribution of this prospectus outside of the United States.

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PROSPECTUS SUMMARY

This summary highlights information contained elsewhere in this prospectus and does not contain all of the information that you should consider in making your investment decision. Before investing in our common stock, you should carefully read this entire prospectus, including our consolidated financial statements and the related notes included elsewhere in this prospectus. You should also consider, among other things, the matters described under Risk Factors and Management's Discussion and Analysis of Financial Condition and Results of Operations, in each case appearing elsewhere in this prospectus. Unless otherwise stated, all references to us, our, SAGE, we, the Company and similar designations refer to Sage Therapeutics, Inc. and its subsidiary.

Overview

We are a biopharmaceutical company committed to developing and commercializing novel medicines to treat life-threatening, rare central nervous system, or CNS, disorders, where there are inadequate or no approved existing therapies. We are targeting CNS indications where patient populations are easily identified, acute treatment is typically initiated in the hospital setting, clinical endpoints are well-defined and development pathways are feasible.

Our initial product candidates, which are summarized in the table below, are aimed at treating different stages of status epilepticus, or SE, a life-threatening condition in which the brain is in a state of persistent seizure, as well as other seizure and non-seizure disorders. The lead product candidate in our SE program, SAGE-547, is an intravenous, or IV, agent entering Phase 3 clinical development as an adjunctive therapy, a therapy combined with current therapeutic approaches, for the treatment of super-refractory SE, or SRSE. The current standard of care for SRSE is empiric, and there are no therapies at present that have been specifically approved for this indication. Over the course of 2014, the U.S. Food and Drug Administration, or FDA, granted us orphan drug designation and Fast Track designation for our investigational new drug application for SAGE-547 as a treatment for SRSE. On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE and we expect to initiate the Phase 3 trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of a New Drug Application, or NDA, submission for SAGE-547.

We continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for essential tremor, a debilitating neurological disorder that causes involuntary, rhythmic shaking with no known cause, and severe post-partum depression, a distinct and readily identified form of major depressive disorder estimated to affect up to 20% of women following childbirth. If these exploratory trials are successful, we plan to use the data from them to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

Our next-generation product candidates, SAGE-689 and SAGE-217, utilize similar mechanistic pathways as SAGE-547 and are designed to have pharmaceutical properties which optimize both their non-clinical profiles and potential clinical profiles for the treatment of different stages of SE, as well as other seizure and non-seizure disorders.

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Status Epilepticus

SE is diagnosed when a patient has a seizure lasting longer than five minutes, and is associated with substantial morbidity and mortality. We estimate that in the United States each year there are up to 150,000 cases of SE, of which 30,000 SE patients die. We estimate that there are 35,000 patients with SE in the United States that are hospitalized in the intensive care unit, or ICU, each year. This results in an overall inpatient cost of \$3.8 billion to \$7.0 billion per year in the United States. An SE patient is first treated with benzodiazepines, or BDZs, and if no response then treated with other, second-line, anti-seizure drugs. If the seizure persists after second-line therapy the patient is diagnosed as having refractory SE, or RSE, admitted to the ICU and placed into a medically induced coma. Currently, there are no therapies that have been specifically approved for RSE; however, physicians typically use anesthetic agents to induce the coma and stop the seizure immediately. After a period of 24 hours, an attempt is made to wean the patient from the anesthetic agents to evaluate whether or not the seizure condition has resolved. Unfortunately, not all patients respond to weaning attempts, in which case the patient must be maintained in the medically induced coma. At this point, the patient is diagnosed as having SRSE.

SAGE-547 Clinical Development Programs

Super Refractory Status Epilepticus (SRSE) Program Summary and Recent Developments

Prior to the start of our Phase 1/2 clinical trial of SAGE-547, we began to collect data in emergency-use cases of SAGE-547 that we believe supports the safety and activity of SAGE-547 for treatment of SRSE. This emergency-use program continues in parallel with our ongoing Phase 1/2 clinical trial. As of January 9, 2015, ten patients were treated with SAGE-547 by independent centers under emergency-use Investigational New Drug applications, or INDs. Each individual case of SRSE arose from a variety of underlying etiologies, the patients were of varying ages, and all patients had been placed in a long-duration medically induced coma prior to the administration of SAGE 547. We

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experienced an overall response rate of 78% in seven of the nine evaluable patients.

In January 2014, we commenced our Phase 1/2 clinical trial to study safety, tolerability and efficacy of SAGE-547 in adult patients with SRSE. This clinical trial is designed as an open-label trial in at least ten patients diagnosed with SRSE. In October 2014, the FDA approved a protocol amendment for our Phase 1/2 trial that enables us to treat pediatric patients as young as two years old, increase the dose of SAGE-547 being administered to patients and increase treatment duration. As of February 28, 2015, there were 17 active trial sites in the United States. We are continuing to enroll patients as an expansion cohort in this trial and we anticipate reporting final clinical data from this Phase 1/2 trial at the Antiepileptic Drug and Device Trials XIII Conference, which is taking place May 13-15, 2015.

On January 9, 2015, we reported results from our Phase 1/2 clinical trial. Consistent with topline data announced in November 2014, the primary endpoint of safety and tolerability, was achieved in all patients. Of the 20 patients enrolled in the Phase 1/2 clinical trial, 17 patients were evaluable for efficacy. 71% of evaluable patients met the key efficacy endpoint of being successfully weaned off their anesthetic agents while SAGE-547 was being administered. In addition, 71% of evaluable patients were successfully weaned off SAGE-547 without recurrence of SRSE. As a group, patients who responded to SAGE-547 generally demonstrated rapid improvement over the first five days following treatment. Patients who responded also continued to improve over the 30-day follow-up period. SAGE-547 was generally well-tolerated and no drug-related serious adverse events, as determined by the Safety Review Committee, were reported in treated patients. In the 20 patients treated with SAGE-547, the mean exposure level of SAGE-547 was approximately 200nM.

On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE. Subject to submission and review by the FDA of a final protocol for the planned Phase 3 clinical trial and updated chemistry, manufacturing and controls information, we expect to initiate the trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547.

Additional SAGE 547 Exploratory Development Programs

We continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for additional indications. In October 2014 we began patient enrollment in an exploratory Phase 2a clinical trial of SAGE-547 in patients with essential tremor. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and activity of SAGE-547 in patients with essential tremor. In January 2015, we initiated a Phase 2a clinical trial of SAGE-547 in women with severe postpartum depression, or PPD. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and efficacy of SAGE-547 for the treatment of severe PPD. We plan to use the data from these exploratory trials to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

Follow-On Product Candidates

SAGE-689 and SAGE-217 are two additional product candidates in our pipeline, which are currently in IND-enabling toxicology and safety pharmacology testing. SAGE-689 is being developed

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as an adjunctive second-line therapy for the treatment of SE. We are currently conducting IND-enabling studies of SAGE-689, with a plan to file an IND in late 2015 and to begin a Phase 1 clinical trial thereafter. SAGE-217 is being developed as an oral monotherapy for orphan epilepsies, such as Dravet syndrome and Rett syndrome. The chemical characteristics of SAGE-217 potentially allow formulation as both an intravenous and oral medication. In addition, we believe related molecules from our portfolio may be useful in the treatment of a variety of neurological and psychiatric disorders, including, for example, fragile X syndrome, anxiety and tremor. We are currently conducting IND-enabling studies of SAGE-217 with a plan to file an IND by late 2015 and to begin a Phase 1 clinical trial thereafter.

Understanding the Foundations of Our Approach

Neurotransmission

The CNS is composed of a vast and complex network of different structures and cell types, most of which serve directly or indirectly to provide a means for the nervous system to signal or communicate with other nerve cells in order to regulate and control all brain function. The cell type responsible for this signaling is called a neuron. Chemical or electrical signals can exert their effects on neurons by traveling across a physical gap located between two neurons, called a synapse. Presynaptic neurons transmit signals, whereas postsynaptic neurons react to the signals.

Neurotransmission is the process by which signaling molecules, called neurotransmitters, are released by a presynaptic neuron, travel over the synaptic space and bind to and interact with receptors on a postsynaptic neuron. Synaptic receptors are primarily located inside the synaptic cleft, or the space where the neurons communicate, and have been historically considered to be the most important part of the neuron. However, recent understanding of neurotransmission and brain function has shown there are many extrasynaptic receptors that also respond to neurotransmitters to exert their effects.

Allosteric modulation

We are focused on developing drugs based on selective allosteric modulation of key CNS synaptic and extrasynaptic receptors. Molecules that function directly on synaptic or extrasynaptic receptors at the site where the native, or natural, molecule binds to inhibit or activate them are known as orthosteric. Alternatively, allosteric modulators are a class of small molecules very different from classical orthosteric drugs, as they interact at a site different from the native site and allow for fine-tuning of neuronal signals. As a result, our drugs under development are capable of varying degrees of desired activity rather than complete activation or inhibition of the receptor as typically observed with orthosteric drugs. We believe this greater selectivity and modulatory control at extrasynaptic GABA_A receptors may allow us to develop CNS drugs that offer significant therapeutic and safety advantages over orthosteric drugs.

Allosteric modulation of extrasynaptic GABA_A receptors to treat SE

Our current near-term product candidates are allosteric modulators of both synaptic and extrasynaptic, or existing outside of the synapse, GABA_A receptors, a characteristic important in distinguishing our approach from current therapies. While altering the level of synaptic GABA_A receptor activity can be beneficial in stopping seizures, this approach has limitations for the treatment of SE. As SE progresses in many patients, select synaptic GABA_A receptors are down-regulated, or removed from the neuronal synaptic surface. As a result, drugs that target down-regulated receptors, such as

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BDZs, often are not effective in stopping SE. In contrast, our product candidates work at both the synaptic and extrasynaptic GABA_A receptors. Non-clinical studies suggest that these extrasynaptic GABA_A receptors remain fully active during SE, offering the potential for drugs that impact GABA via the extrasynaptic GABA_A receptor to alter GABA_A activity and abate seizure. We believe that by creating compounds that target both these receptors, we may be successful in treating seizures that do not respond to BDZ therapy.

Allosteric modulation of GABA_A and NMDA receptors to address other CNS conditions

Now and in the foreseeable future, our product development pipeline will be focused on allosteric modulation of two important receptor systems in the brain GABA_A and NMDA. These receptor systems regulate inhibitory and excitatory neurotransmission, respectively, and are broadly accepted as impacting many psychiatric and neurological disorders. GABA_A and NMDA receptor systems are widely regarded as validated drug targets for a variety of CNS disorders, with decades of research and multiple approved drugs targeting these receptor systems. Drugs approved to modulate these receptor systems have had safety and efficacy limitations related to their poor pharmaceutical properties and adverse side effects. We believe that we will have the opportunity to develop molecules from our internal portfolio to more effectively address many of these disorders in the future.

Our proprietary chemistry platform

Our ability to identify and develop such novel CNS therapies is enabled by our proprietary chemistry platform that is centered on a scaffold of chemically modified endogenous neuroactive steroid compounds. We believe our know-how around the chemistry and activity of allosteric modulators allows us to efficiently design molecules with different characteristics by enabling us to control important properties such as half-life, brain penetration and the types of receptors with which our drugs interact. Therefore, we believe our product candidates will have the potential to bind with targets in the brain with more precision, increased safety and tolerability, and fewer off-target side effects than either current CNS therapies or previous therapies, which have often failed in development.

Our Strategy

Our goal is to become a leading biopharmaceutical company focused on development and commercialization of novel proprietary therapies for the treatment of life-threatening, rare CNS disorders. Key elements of our strategy are to:

Rapidly advance SAGE-547 as a treatment for SRSE.

Utilize SAGE-547 in exploratory trials to help guide the development of second-generation GABA_A receptor modulators for the applicable diseases.

Develop our next generation product candidates, SAGE-689 and SAGE-217, in parallel with SAGE-547.

Enhance the probability of success in treating SE by developing unique assets with differentiated features.

Grow our pipeline more broadly utilizing the strengths of our proprietary chemistry platform and scientific know-how, to lessen our long-term reliance on a single franchise and facilitate long-term growth.

Focus our internal development activities on CNS indications where we can make well-informed, rapid go/no-go decisions.

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Build a commercial capability to bring our CNS therapeutics to physicians and patients for rare target indications.

Selectively partner our programs to enhance our value.

Risk Factors

Our business is subject to many risks and uncertainties of which you should be aware before you decide to invest in our common stock. These risks are discussed more fully under **Risk Factors** in this prospectus. Some of these risks include:

We depend heavily on the success of the product candidates within our seizure programs, of which SAGE-547 is entering Phase 3 clinical development and SAGE-689 and SAGE-217 are in non-clinical development. We cannot be certain that we will be able to obtain regulatory approval for, or successfully commercialize, any of our product candidates.

Prior to commencing enrollment in our planned Phase 3 clinical trial of SAGE-547, we must provide to the FDA additional information. If the additional information we provide is not satisfactory to the FDA, it could delay the start of, or change the design of, our planned Phase 3 clinical trial.

The number of patients suffering from SE, RSE or SRSE is small and has not been established with precision. If the actual number of patients with SE, RSE or SRSE is smaller than we anticipate, we may encounter difficulties in enrolling patients in our clinical trials, thereby delaying or preventing development of our product candidates, and if any of our product candidates are approved, we believe our revenue and ability to achieve profitability would be materially adversely affected.

Positive results from early non-clinical studies and clinical trials of our product candidates are not necessarily predictive of the results of later non-clinical studies and clinical trials of our product candidates. If we cannot replicate the positive results from our earlier non-clinical studies and clinical trials of our product candidates in our later non-clinical studies and clinical trials, we may be unable to successfully develop, obtain regulatory approval for and commercialize our product candidates.

If serious adverse events or other undesirable side effects are identified during the use of SAGE-547 in emergency-use cases, investigator sponsored trials or exploratory clinical trials of SAGE-547, our development of SAGE-547 for SRSE may be adversely effected.

Failures or delays in the commencement or completion of our planned clinical trials of our product candidates could result in increased costs to us and could delay, prevent or limit our ability to generate revenue and continue our business.

Even though we have obtained orphan drug designation for SAGE-547 as a treatment for SE, there may be limitations to the exclusivity afforded by such designation.

We rely, and expect that we will continue to rely, on third parties to conduct any clinical trials for our product candidates. If these third parties do not successfully carry out their contractual duties or meet expected deadlines, we may not be able to obtain regulatory approval for or commercialize our product candidates and our business could be substantially harmed.

We are dependent on licensed intellectual property. If we were to lose our rights to licensed intellectual property, we may not be able to continue developing or commercializing our product candidates, if approved.

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If we are unable to adequately protect our proprietary technology, or to obtain and maintain issued patents that are sufficient to protect our product candidates, others could compete against us more directly, which would have a material adverse impact on our business, results of operations, financial condition and prospects.

Our future success depends on our ability to retain our President and Chief Executive Officer and to attract, retain and motivate qualified personnel.

Implications of being an emerging growth company

We qualify as an emerging growth company as defined in the Jumpstart Our Business Startups Act of 2012, or the JOBS Act. As an emerging growth company, we may take advantage of specified reduced disclosure and other requirements that are otherwise applicable generally to public companies. These provisions include:

reduced disclosure about our executive compensation arrangements;

no non-binding advisory votes on executive compensation or golden parachute arrangements; and

exemption from the auditor attestation requirement in the assessment of our internal control over financial reporting.

We may take advantage of these exemptions for up to five years or such earlier time that we are no longer an emerging growth company. We would cease to be an emerging growth company on the date that is the earliest of (i) the last day of the fiscal year in which we have total annual gross revenues of \$1 billion or more; (ii) the last day of our fiscal year following the fifth anniversary of the date of the completion of our initial public offering; (iii) the date on which we have issued more than \$1 billion in nonconvertible debt during the previous three years; or (iv) the date on which we are deemed to be a large accelerated filer under the rules of the Securities and Exchange Commission, or SEC, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30th. We may choose to take advantage of some but not all of these exemptions. We have taken advantage of reduced reporting requirements in this prospectus. Accordingly, the information contained herein may be different than the information you receive from other public companies in which you hold stock. Also, we have irrevocably elected to opt out of the exemption for the delayed adoption of certain accounting standards and, therefore, will be subject to the same new or revised accounting standards as other public companies that are not emerging growth companies.

Corporate History and Information

We were incorporated under the laws of the state of Delaware in April 2010. Our principal executive office is located at 215 First Avenue, Cambridge, Massachusetts, and our telephone number is (617) 299-8380. Our website address is www.sagerx.com. We do not incorporate the information on or accessible through our website into this prospectus, and you should not consider any information on, or that can be accessed through, our website as part of this prospectus.

We own various U.S. federal trademark registrations and applications and unregistered trademarks, including our corporate logo. All other trademarks or trade names referred to in this prospectus are the property of their respective owners. Solely for convenience, the trademarks and trade names in this prospectus may be referred to without the ® and ™ symbols, but such references should not be construed as any indicator that their respective owners will not assert, to the fullest extent under applicable law, their rights thereto.

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282,000 shares of common stock reserved for future issuance under our 2014 Employee Stock Purchase Plan, as of February 28, 2015.

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Except as otherwise indicated, all information in this prospectus assumes or gives effect to:

no exercise of the outstanding options described above; and

no exercise by the underwriters of their option to purchase up to an additional shares of our common stock in this offering.

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You should read the following summary consolidated financial data together with our consolidated financial statements and the related notes appearing at the end of this prospectus and the Selected Consolidated Financial Data and Management's Discussion and Analysis of Financial Condition and Results of Operations sections of this prospectus. We have derived the consolidated statements of operations data for the years ended December 31, 2014, 2013, and 2012 and the consolidated balance sheet data as of December 31, 2014 from our audited consolidated financial statements included elsewhere in this prospectus. Our historical results are not necessarily indicative of results that should be expected in the future.

	Years Ended December 31,		
	2014	2013	2012
	(in thousands, except for per share amounts)		
Consolidated statements of operations data:			
Operating expenses:			
Research and development	\$ 24,100	\$ 14,357	\$ 7,229
General and administrative	9,710	3,922	2,402
Total operating expenses	33,810	18,279	9,631
Loss from operations	(33,810)	(18,279)	(9,631)
Interest income (expense), net	8	1	
Other income (expense), net	(9)	(3)	(1)
Net loss and comprehensive loss	(33,811)	(18,281)	(9,632)
Accretion of redeemable convertible preferred stock to redemption value	(2,294)	(7)	(4)
Net loss attributable to common stockholders	\$ (36,105)	\$ (18,288)	\$ (9,636)
Net loss per share attributable to common stockholders - basic and diluted ⁽¹⁾	\$ (1.67)	\$ (12.26)	\$ (8.62)
Weighted average common shares outstanding - basic and diluted ⁽¹⁾	21,574	1,492	1,118
		As of December 31, 2014	
		Actual	As Adjusted ⁽³⁾
		(in thousands)	
Consolidated balance sheet data:			
Cash and cash equivalents		\$ 127,766	\$
Working capital ⁽²⁾		121,065	
Total assets		129,665	
Total stockholders' equity (deficit)		121,885	

- (1) See Note 8 to our consolidated financial statements for further details on the calculation of basic and diluted net loss per share attributable to common stockholders.
- (2) We define working capital as current assets less current liabilities.
- (3) As adjusted consolidated balance sheet data gives effect to the sale by us of _____ shares of our common stock in this offering at an assumed public offering price of \$46.87 per share, which was the last reported sales price of our common stock on The NASDAQ Global Market on April 2, 2015 after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

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RISK FACTORS

*Investing in our common stock involves a high degree of risk. You should carefully consider the following risks and uncertainties, together with all other information in this prospectus, including our consolidated financial statements and related notes, before investing in our common stock. Any of the risk factors we describe below could adversely affect our business, financial condition or results of operations. The market price of our common stock could decline if one or more of these risks or uncertainties actually occur, causing you to lose all or part of the money you paid to buy our common stock. Additional risks that we currently do not know about or that we currently believe to be immaterial may also impair our business. Certain statements below are forward-looking statements. See *Cautionary Note Regarding Forward-Looking Statements* in this prospectus.*

Risks Related to Product Development, Regulatory Approval and Commercialization

We depend heavily on the success of the product candidates within our seizure programs, of which SAGE-547 is entering Phase 3 clinical development and SAGE-689 and SAGE-217 are in non-clinical development. We cannot be certain that we will be able to obtain regulatory approval for, or successfully commercialize, any of our product candidates.

We currently have no drug products for sale and may never be able to develop marketable drug products. Our business depends heavily on the successful non-clinical and clinical development, regulatory approval and commercialization of the product candidates in our lead program in status epilepticus, or SE, of which only one product candidate, SAGE-547, is entering Phase 3 clinical development for the treatment of super-refractory SE, or SRSE, and our other product candidates, SAGE-689 and SAGE-217, are in non-clinical development. SAGE-547 will require substantial additional clinical development, testing and regulatory approval before we are permitted to commence its commercialization. The non-clinical studies and clinical trials of our product candidates are, and the manufacturing and marketing of our product candidates will be, subject to extensive and rigorous review and regulation by numerous government authorities in the United States and in other countries where we intend to test and, if approved, market any product candidate. Before obtaining regulatory approvals for the commercial sale of any product candidate, we must demonstrate through non-clinical studies and clinical trials that the product candidate is safe and effective for use in each target indication. Drug development is a long, expensive and uncertain process, and delay or failure can occur at any stage of any of our clinical trials. This process can take many years and may include post-marketing studies and surveillance, which will require the expenditure of substantial resources beyond the proceeds we raise in this offering. Of the large number of drugs in development in the United States, only a small percentage will successfully complete the U.S. Food and Drug Administration, or FDA, regulatory approval process and will be commercialized. Accordingly, even if we are able to obtain the requisite financing to continue to fund our development and non-clinical studies and clinical trials, we cannot assure you that any of our product candidates will be successfully developed or commercialized.

Both SAGE-689 and SAGE-217 are in non-clinical development and have yet to begin the clinical development process. We plan to file Investigational New Drug Applications, or INDs, for both SAGE-689 and SAGE-217 late in 2015 and to begin a Phase 1 clinical trial for each of SAGE-689 and SAGE-217 thereafter.

We are not permitted to market our product candidates in the United States until we receive approval of a New Drug Application, or an NDA, from the FDA, or in any foreign countries until we receive the requisite approval from such countries. Obtaining approval of an NDA is a complex, lengthy, expensive and uncertain process, and the FDA may delay, limit or deny approval of any of our product candidates for many reasons, including, among others:

we may not be able to demonstrate that our product candidates are safe and effective in treating SE, refractory SE, or RSE, or SRSE, as applicable, to the satisfaction of the FDA;

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the results of our non-clinical studies and clinical trials may not meet the level of statistical or clinical significance required by the FDA for marketing approval;

the FDA may disagree with the number, design, size, conduct, implementation of or differing drug formulations used in our non-clinical studies and clinical trials;

the FDA may require that we conduct additional non-clinical studies and clinical trials;

the FDA or the applicable foreign regulatory agency may not approve the formulation, labeling or specifications of any of our product candidates;

the contract research organizations, or CROs, that we retain to conduct our non-clinical studies and clinical trials may take actions outside of our control that materially adversely impact our non-clinical studies and clinical trials;

the FDA may find the data from non-clinical studies and clinical trials insufficient to demonstrate that our product candidates' clinical and other benefits outweigh their safety risks;

the FDA may disagree with our interpretation of data from our non-clinical studies and clinical trials;

the FDA may not accept data generated at our non-clinical studies and clinical trial sites;

if our NDA, if and when submitted, is reviewed by an advisory committee, the FDA may have difficulties scheduling an advisory committee meeting in a timely manner or the advisory committee may recommend against approval of our application or may recommend that the FDA require, as a condition of approval, additional non-clinical studies or clinical trials, limitations on approved labeling or distribution and use restrictions;

the FDA may require development of a Risk Evaluation and Mitigation Strategy, or REMS, as a condition of approval or post-approval;

the FDA or the applicable foreign regulatory agency may determine that the manufacturing processes or facilities of third-party contract manufacturers with which we contract do not conform to applicable requirements, including current Good Manufacturing Practices, or cGMPs; or

the FDA or applicable foreign regulatory agency may change its approval policies or adopt new regulations.

Any of these factors, many of which are beyond our control, could jeopardize our ability to obtain regulatory approval for and successfully market our product candidates. Any such setback in our pursuit of regulatory approval would have a material adverse effect on our business and prospects.

We cannot be certain that our planned Phase 3 clinical trial of SAGE-547 will be sufficient to support the submission of an NDA for this product candidate, and in any event we must obtain additional clinical and non-clinical data before an NDA may be submitted.

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In general, the FDA requires two pivotal trials to support approval of an NDA, but in certain circumstances, will approve an NDA based on only one pivotal trial. If successful, we believe the results from our planned Phase 3 clinical trial of SAGE-547, together with safety and efficacy data from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547. However depending upon the outcome of the current program, the FDA may require that we conduct additional pivotal trials before we can submit an NDA for SAGE-547. To allow dosing in patients below the age of two we would need to either conduct additional clinical trial(s) or amend the protocol for our planned Phase 3 clinical trial.

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Furthermore, we will need to complete several other clinical studies prior to submitting an NDA to the FDA, potentially including an absorption, metabolism, and excretion pharmacokinetics study in healthy volunteers, studies to test the effect of SAGE-547 on exposure to phenytoin and in patients with severe renal impairment and patients with hepatic impairment, as well as a study to test the abuse potential of SAGE-547. If the result of these additional clinical studies are not positive or yield unanticipated results, it may delay or prevent the submission or approval of an NDA for SAGE-547.

While we believe we and the FDA are in general agreement on the design and key elements of our planned Phase 3 clinical trial for SAGE-547, before beginning the trial, the FDA must review the final protocol for the trial. Concurrent with starting the Phase 3 clinical trial, the FDA will review certain updated chemistry, manufacturing and controls, or CMC, information, that we are required to submit. We also plan to share with the FDA the results of our long-term toxicity studies in two animal species, the first segment of which we submitted to the FDA in the second quarter of 2014. Additional long-term toxicity studies, required for an NDA submission, are ongoing. If the FDA does not approve the protocol for the planned trial in the form we submit it, or if the FDA is not satisfied with the additional CMC information we plan to provide, the start or continuation of the planned Phase 3 trial may be delayed or the design of the trial may change. The FDA may require that we conduct additional toxicity studies and other non-clinical studies before submitting an NDA for SAGE-547.

A Fast Track designation by the FDA may not actually lead to a faster development or regulatory review or approval process.

We have received Fast Track designation for our investigational new drug application, or IND, for SAGE-547 for the treatment of SRSE, and in the future we may seek Fast Track designation for other product candidates as well. If a product is intended for the treatment of a serious or life-threatening condition and the product demonstrates the potential to address unmet medical needs for this condition, the sponsor may apply for the FDA Fast Track designation. Fast Track designation does not necessarily lead to a faster development pathway or regulatory review process and does increase the likelihood of regulatory approval. The FDA may withdraw Fast Track designation if it believes that the designation is no longer supported by data from our clinical development programs.

The number of patients suffering from SE, RSE and SRSE is small or has not been established with precision. If the actual number of patients with SE, RSE and SRSE is smaller than we anticipate, we may encounter difficulties in enrolling patients in our clinical trials, thereby delaying or preventing development of our product candidates, and if any of our product candidates are approved, we believe our revenue and ability to achieve profitability would be materially adversely affected.

There is no precise method of establishing actual number of patients with SE, RSE or SRSE in any geography over any time period. Moreover, SE, RSE and SRSE are acute episode conditions. If we are not able to identify patients at the time of SE, RSE or SRSE onset, we will have difficulty completing our clinical trials. We estimate that the annual incidence of SE, RSE and SRSE in the United States is up to 150,000, 35,000 and 25,000 patients, respectively. If the actual number of patients with SE, RSE or SRSE is lower than we believe, we may experience difficulty in enrolling patients in our clinical trials, thereby delaying development of our product candidates. Further, if any of our product candidates are approved, the markets for our product candidates for these indications would be smaller than we anticipate which could limit our ability to achieve profitability.

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Favorable results from the emergency-use cases of SAGE-547 do not ensure that clinical trials will be successful and the results in any future emergency-use cases may not be positive and could adversely impact our clinical development plans.

SAGE-547 has been administered to a small number of patients as part of emergency-use cases, which permitted the administration of SAGE-547 outside of clinical trials. No assurance can be given that positive results observed to date in these emergency-use cases are attributable to SAGE-547, as they were not carried out in the controlled environment of a clinical trial. Further, no assurance can be provided that administration of SAGE-547 to other patients in any future emergency-use cases or otherwise will have positive results. Emergency use is a term that is used to refer to the use of an investigational drug outside of a clinical trial to treat a patient with a serious or immediately life-threatening disease or condition and who has no comparable or satisfactory alternative treatment options. Regulators often allow emergency use on a case-by-case basis for an individual patient or for defined groups of patients with similar treatment needs. In the event there are negative results in future emergency-use cases, it could adversely affect or delay our clinical development of SAGE-547.

If serious adverse events or other undesirable side effects are identified during the use of SAGE-547 in emergency-use cases, investigator sponsored trials or exploratory clinical trials of SAGE-547, it may adversely effect our development of SAGE-547 for SRSE.

In addition to use in emergency cases as described above, SAGE-547 is currently being tested in an investigator sponsored clinical trial for the treatment of traumatic brain injury, or TBI, by one of our collaborators and may be subjected to testing for other indications in additional investigator sponsored trials. Currently, we are also testing SAGE-547 in a proof of concept trial in patients with essential tremor and a proof of concept trial in patients with severe postpartum depression, or PPD. If serious adverse events or other undesirable side effects, or unexpected characteristics of SAGE-547 are observed in emergency-use cases or in investigator sponsored clinical trials of SAGE-547 or our exploratory clinical trials, it may adversely affect or delay our clinical development of SAGE-547, or we may need to abandon its development for SRSE entirely, and the occurrence of these events would have a material adverse effect on our business.

Positive results from early non-clinical studies and clinical trials of our product candidates are not necessarily predictive of the results of later non-clinical studies and clinical trials of our product candidates. If we cannot replicate the positive results from our earlier non-clinical studies and clinical trials of our product candidates in our later non-clinical studies and clinical trials, we may be unable to successfully develop, obtain regulatory approval for and commercialize our product candidates.

Positive results from our non-clinical studies of our product candidates, and any positive results we may obtain from our early clinical trials of our product candidates, may not necessarily be predictive of the results from required later non-clinical studies and clinical trials. Similarly, even if we are able to complete our planned non-clinical studies or clinical trials of our product candidates according to our current development timeline, the positive results from our non-clinical studies and clinical trials of our product candidates may not be replicated in subsequent non-clinical studies or clinical trial results. For example, although 12 of the first 17 patients treated with SAGE-547 and evaluable for efficacy in our Phase 1/2 clinical trial met the key efficacy endpoint and none of the 20 patients enrolled in the study have yet experienced any severe adverse events related to SAGE-547, future patients enrolled and treated with SAGE-547 in later-stage clinical trials may not have the same outcome. Also, our later-stage clinical trials will differ in important ways from our ongoing Phase 1/2 clinical trial of SAGE-547, which could cause the outcome of these later-stage trials to differ from our earlier stage clinical trials. For example, our planned Phase 3 clinical trial of SAGE-547 will be a placebo-controlled trial, while our Phase 1/2 clinical trial was open-label, and intent-to-treat statistical analysis will be employed in our

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planned Phase 3 clinical trial. In addition, the formulation of SAGE-547 we intend to use in our planned Phase 3 trial is somewhat different than the formulation used in the Phase 1/2 trial. We do not believe the change will negatively affect trial results, but we cannot be sure. Many companies in the pharmaceutical and biotechnology industries have suffered significant setbacks in late-stage clinical trials after achieving positive results in early-stage development, and we cannot be certain that we will not face similar setbacks. These setbacks have been caused by, among other things, non-clinical findings made while clinical trials were underway or safety or efficacy observations made in non-clinical studies and clinical trials, including previously unreported adverse events. Moreover, non-clinical and clinical data are often susceptible to varying interpretations and analyses, and many companies that believed their product candidates performed satisfactorily in non-clinical studies and clinical trials nonetheless failed to obtain FDA approval. We have not completed any clinical trials for our product candidates yet, and if we fail to produce positive results in our planned non-clinical studies or clinical trials of any of our product candidates, the development timeline and regulatory approval and commercialization prospects for our product candidates, and, correspondingly, our business and financial prospects, would be materially adversely affected.

Failures or delays in the commencement or completion of our planned clinical trials of our product candidates could result in increased costs to us and could delay, prevent or limit our ability to generate revenue and continue our business.

We have an ongoing Phase 1/2 clinical trial of SAGE-547 as a treatment for SRSE and ongoing proof of concept studies of SAGE-547 for patients with essential tremor and severe PPD. We will need to complete at least one additional trial prior to the submission of an NDA for SAGE-547 as a treatment for SRSE. Successful completion of our clinical trials is a prerequisite to submitting an NDA to the FDA and, consequently, the ultimate approval and commercial marketing of SAGE-547 for SRSE and our other product candidates. We do not know whether any of our clinical trials will begin or be completed on schedule, if at all, as the commencement and completion of clinical trials can be delayed or prevented for a number of reasons, including, among others:

the FDA may deny permission to proceed with our planned clinical trials or any other clinical trials we may initiate, or may place a clinical trial on hold;

delays in filing or receiving approvals of additional INDs that may be required;

negative results from our ongoing non-clinical studies;

delays in reaching or failing to reach agreement on acceptable terms with prospective CROs and clinical trial sites, the terms of which can be subject to extensive negotiation and may vary significantly among different CROs and trial sites;

inadequate quantity or quality of a product candidate or other materials necessary to conduct clinical trials, for example delays in the manufacturing of sufficient supply of finished drug product;

difficulties obtaining Institutional Review Board, or IRB, approval to conduct a clinical trial at a prospective site or sites;

challenges in recruiting and enrolling patients to participate in clinical trials, including the small size of the patient population, acute nature of SRSE, the proximity of patients to trial sites;

eligibility criteria for the clinical trial, the nature of the clinical trial protocol, the availability of approved effective treatments for the relevant disease and competition from other clinical trial programs for similar indications;

severe or unexpected drug-related side effects experienced by patients in a clinical trial;

delays in validating any endpoints utilized in a clinical trial;

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the FDA may disagree with our clinical trial design and our interpretation of data from clinical trials, or may change the requirements for approval even after it has reviewed and commented on the design for our clinical trials;

our inability to satisfy the CMC requirements of the FDA or file amendments to our IND as requested by the FDA prior to the initiation of a clinical trial;

reports from non-clinical or clinical testing of other CNS therapies that raise safety or efficacy concerns; and

difficulties retaining patients who have enrolled in a clinical trial but may be prone to withdraw due to rigors of the clinical trials, lack of efficacy, side effects, personal issues or loss of interest.

Clinical trials may also be delayed or terminated as a result of ambiguous or negative interim results. In addition, a clinical trial may be suspended or terminated by us, the FDA, the IRBs at the sites where the IRBs are overseeing a clinical trial, a data and safety monitoring board, or DSMB, overseeing the clinical trial at issue or other regulatory authorities due to a number of factors, including, among others:

failure to conduct the clinical trial in accordance with regulatory requirements or our clinical protocols;

inspection of the clinical trial operations or trial sites by the FDA or other regulatory authorities that reveals deficiencies or violations that require us to undertake corrective action, including the imposition of a clinical hold;

unforeseen safety issues, including any that could be identified in our ongoing non-clinical carcinogenicity studies, adverse side effects or lack of effectiveness;

changes in government regulations or administrative actions;

problems with clinical supply materials; and

lack of adequate funding to continue clinical trials.

Changes in regulatory requirements, FDA guidance or unanticipated events during our non-clinical studies and clinical trials of our product candidates may occur, which may result in changes to non-clinical studies and clinical trial protocols or additional non-clinical studies and clinical trial requirements, which could result in increased costs to us and could delay our development timeline.

Changes in regulatory requirements, FDA guidance or unanticipated events during our non-clinical studies and clinical trials may force us to amend non-clinical studies and clinical trial protocols or the FDA may impose additional non-clinical studies and clinical trial requirements. Amendments or changes to our clinical trial protocols would require resubmission to the FDA and IRBs for review and approval, which may adversely impact the cost, timing or successful completion of clinical trials. Similarly, amendments to our non-clinical studies may adversely impact the cost, timing, or successful completion of those non-clinical studies. For example, we intend to seek a waiver from the need to perform a study of SAGE-547 on certain cardiac measures. If the FDA does not grant the waiver, we will be required to conduct such a study, the results of which could delay the filing of an NDA for SAGE-547. If we experience delays completing, or if we terminate, any of our non-clinical studies or clinical trials, or if we are required to conduct additional non-clinical studies or clinical trials, the commercial prospects for our product candidates may be harmed and our ability to generate product revenue will be delayed.

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We rely, and expect that we will continue to rely, on third parties to conduct any clinical trials for our product candidates. If these third parties do not successfully carry out their contractual duties or meet expected deadlines, we may not be able to obtain regulatory approval for or commercialize our product candidates and our business could be substantially harmed.

We do not have the ability to independently conduct clinical trials. We rely on medical institutions, clinical investigators, contract laboratories and other third parties, such as CROs, to conduct clinical trials on our product candidates. We enter into agreements with third-party CROs to provide monitors for and to manage data for our ongoing clinical trials. We rely heavily on these parties for execution of clinical trials for our product candidates and control only certain aspects of their activities. As a result, we have less direct control over the conduct, timing and completion of these clinical trials and the management of data developed through clinical trials than would be the case if we were relying entirely upon our own staff. Communicating with outside parties can also be challenging, potentially leading to mistakes as well as difficulties in coordinating activities. Outside parties may:

have staffing difficulties;

fail to comply with contractual obligations;

experience regulatory compliance issues;

undergo changes in priorities or become financially distressed; or

form relationships with other entities, some of which may be our competitors.

These factors may materially adversely affect the willingness or ability of third parties to conduct our clinical trials and may subject us to unexpected cost increases that are beyond our control. Nevertheless, we are responsible for ensuring that each of our clinical trials is conducted in accordance with the applicable protocol, legal, regulatory and scientific requirements and standards, and our reliance on CROs does not relieve us of our regulatory responsibilities. We and our CROs are required to comply with regulations and guidelines, including current Good Clinical Practices, or cGCPs, for conducting, monitoring, recording and reporting the results of clinical trials to ensure that the data and results are scientifically credible and accurate, and that the trial patients are adequately informed of the potential risks of participating in clinical trials. These regulations are enforced by the FDA, the Competent Authorities of the Member States of the European Economic Area and comparable foreign regulatory authorities for any products in clinical development. The FDA enforces cGCP regulations through periodic inspections of clinical trial sponsors, principal investigators and trial sites. If we or our CROs fail to comply with applicable cGCPs, the clinical data generated in our clinical trials may be deemed unreliable and the FDA or comparable foreign regulatory authorities may require us to perform additional clinical trials before approving our marketing applications. We cannot assure you that, upon inspection, the FDA will determine that any of our clinical trials comply with cGCPs. In addition, our clinical trials must be conducted with product candidates produced under cGMPs regulations and will require a large number of test patients. Our failure or the failure of our CROs to comply with these regulations may require us to repeat clinical trials, which would delay the regulatory approval process and could also subject us to enforcement action up to and including civil and criminal penalties.

Although we design our clinical trials for our product candidates, CROs conduct all of the clinical trials. As a result, many important aspects of our drug development programs are outside of our direct control. In addition, the CROs may not perform all of their obligations under arrangements with us or in compliance with regulatory requirements, but we remain responsible and are subject to enforcement action that may include civil penalties up to and including criminal prosecution for any violations of FDA laws and regulations during the conduct of our clinical trials. If the CROs do not perform clinical trials in a satisfactory manner, breach their obligations to us or fail to comply with regulatory requirements, the development and commercialization of our product candidates may be delayed or our development

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program materially and irreversibly harmed. We cannot control the amount and timing of resources these CROs devote to our program or our clinical products. If we are unable to rely on clinical data collected by our CROs, we could be required to repeat, extend the duration of, or increase the size of our clinical trials and this could significantly delay commercialization and require significantly greater expenditures.

If any of our relationships with these third-party CROs terminate, we may not be able to enter into arrangements with alternative CROs. If CROs do not successfully carry out their contractual duties or obligations or meet expected deadlines, if they need to be replaced or if the quality or accuracy of the clinical data they obtain is compromised due to the failure to adhere to our clinical protocols, regulatory requirements or for other reasons, any clinical trials such CROs are associated with may be extended, delayed or terminated, and we may not be able to obtain regulatory approval for or successfully commercialize our product candidates. As a result, we believe that our financial results and the commercial prospects for our product candidates in the subject indication would be harmed, our costs could increase and our ability to generate revenue could be delayed.

We rely completely on third-party suppliers to manufacture our clinical drug supplies for our product candidates, and we intend to rely on third parties to produce non-clinical, clinical and commercial supplies of any future product candidate.

We do not currently have, nor do we plan to acquire, the infrastructure or capability to internally manufacture our clinical drug supply of our product candidates, or any future product candidates, for use in the conduct of our non-clinical studies and clinical trials, and we lack the internal resources and the capability to manufacture any product candidates on a clinical or commercial scale. For example, SAGE-547 used in the emergency-use cases was manufactured at an academic site, the active pharmaceutical ingredient for SAGE-547 for our Phase 1/2 clinical trial was manufactured at an academic site and SAGE-547 as formulated for our Phase 1/2 clinical trial was manufactured at a third-party contract manufacturer's site. The facilities used by our contract manufacturers to manufacture the active pharmaceutical ingredient and final drug product must complete a pre-approval inspection by the FDA and other comparable foreign regulatory agencies to assess compliance with applicable requirements, including cGMPs, after we submit our NDA or relevant foreign regulatory submission to the applicable regulatory agency.

We do not control the manufacturing process of, and are completely dependent on, our contract manufacturers to comply with cGMPs for manufacture of both active drug substances and finished drug products. If our contract manufacturers cannot successfully manufacture material that conforms to our specifications and the strict regulatory requirements of the FDA or applicable foreign regulatory agencies, they will not be able to secure and/or maintain regulatory approval for their manufacturing facilities. In addition, we have no direct control over our contract manufacturers' ability to maintain adequate quality control, quality assurance and qualified personnel. Furthermore, all of our contract manufacturers are engaged with other companies to supply and/or manufacture materials or products for such companies, which exposes our third-party contract manufacturers to regulatory risks for the production of such materials and products. As a result, failure to satisfy the regulatory requirements for the production of those materials and products may affect the regulatory clearance of our contract manufacturers' facilities generally. If the FDA or an applicable foreign regulatory agency determines now or in the future that these facilities for the manufacture of our product candidates are noncompliant, we may need to find alternative manufacturing facilities, which would adversely impact our ability to develop, obtain regulatory approval for or market our product candidates. Our reliance on contract manufacturers also exposes us to the possibility that they, or third parties with access to their facilities, will have access to and may appropriate our trade secrets or other proprietary information.

We do not have long-term supply agreements in place with our contract manufacturers and each batch of our product candidates is individually contracted under a quality and supply agreement. If we

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engage new contract manufacturers, such contractors must complete an inspection by the FDA and other applicable foreign regulatory agencies. We plan to continue to rely upon contract manufacturers and, potentially, collaboration partners to manufacture commercial quantities of our product candidates, if approved. Our current scale of manufacturing is adequate to support all of our needs for non-clinical studies and clinical trial supplies.

Even if we receive marketing approval for our product candidates in the United States, we may never receive regulatory approval to market our product candidates outside of the United States.

We have not yet selected any markets outside of the United States where we intend to seek regulatory approval to market our product candidates. In order to market any product outside of the United States, however, we must establish and comply with the numerous and varying safety, efficacy and other regulatory requirements of other countries. Approval procedures vary among countries and can involve additional product candidate testing and additional administrative review periods. The time required to obtain approvals in other countries might differ from that required to obtain FDA approval. The marketing approval processes in other countries may implicate all of the risks detailed above regarding FDA approval in the United States as well as other risks. In particular, in many countries outside of the United States, products must receive pricing and reimbursement approval before the product can be commercialized. Obtaining this approval can result in substantial delays in bringing products to market in such countries. Marketing approval in one country does not ensure marketing approval in another, but a failure or delay in obtaining marketing approval in one country may have a negative effect on the regulatory process in others. Failure to obtain marketing approval in other countries or any delay or other setback in obtaining such approval would impair our ability to market our product candidates in such foreign markets. Any such impairment would reduce the size of our potential market, which could have a material adverse impact on our business, results of operations and prospects.

If we are unable to establish sales and marketing capabilities or enter into agreements with third parties to market and sell our product candidates, we may not be able to generate any revenue.

We do not currently have an infrastructure for the sales, marketing and distribution of pharmaceutical products. In order to market our product candidates, if approved by the FDA or any other regulatory body, we must build our sales, marketing, managerial and other non-technical capabilities or make arrangements with third parties to perform these services. If we are unable to establish adequate sales, marketing and distribution capabilities, whether independently or with third parties, or if we are unable to do so on commercially reasonable terms, our business, results of operations, financial condition and prospects will be materially adversely affected.

Even if we receive marketing approval for our product candidates, our product candidates may not achieve broad market acceptance, which would limit the revenue that we generate from their sales.

The commercial success of our product candidates, if approved by the FDA or other applicable regulatory authorities, will depend upon the awareness and acceptance of our product candidates among the medical community, including physicians, patients and healthcare payors. Market acceptance of our product candidates, if approved, will depend on a number of factors, including, among others:

the efficacy of our product candidates as demonstrated in clinical trials, and, if required by any applicable regulatory authority in connection with the approval for the applicable indications, to provide patients with incremental health benefits, as compared with other available CNS therapies;

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limitations or warnings contained in the labeling approved for our product candidates by the FDA or other applicable regulatory authorities;

the clinical indications for which our product candidates are approved;

availability of alternative treatments already approved or expected to be commercially launched in the near future;

the potential and perceived advantages of our product candidates over current treatment options or alternative treatments, including future alternative treatments;

the willingness of the target patient population to try new therapies and of physicians to prescribe these therapies;

the strength of marketing and distribution support and timing of market introduction of competitive products;

publicity concerning our products or competing products and treatments;

pricing and cost effectiveness;

the effectiveness of our sales and marketing strategies;

our ability to increase awareness of our product candidates through marketing efforts;

our ability to obtain sufficient third-party coverage or reimbursement; or

the willingness of patients to pay out-of-pocket in the absence of third-party coverage.

If our product candidates are approved but do not achieve an adequate level of acceptance by patients, physicians and payors, we may not generate sufficient revenue from our product candidates to become or remain profitable. Before granting reimbursement approval, healthcare payors may require us to demonstrate that our product candidates, in addition to treating these target indications, also provide incremental health benefits to patients. Our efforts to educate the medical community and third-party payors about the benefits of our product candidates may require significant resources and may never be successful.

Our product candidates may cause undesirable side effects that could delay or prevent their regulatory approval, limit the commercial profile of an approved label, or result in significant negative consequences following marketing approval, if any.

Undesirable side effects caused by our product candidates could cause us or regulatory authorities to interrupt, delay or halt non-clinical studies and clinical trials and could result in a more restrictive label or the delay or denial of regulatory approval by the FDA or other regulatory authorities.

Further, clinical trials by their nature utilize a sample of the potential patient population. With a limited number of patients and limited duration of exposure, rare and severe side effects of our product candidates may only be uncovered with a significantly larger number of patients exposed to the product candidate. If our product candidates receive marketing approval and we or others identify undesirable side effects caused by such product candidates (or any other similar products) after such approval, a number of potentially significant negative consequences could result,

including:

regulatory authorities may withdraw or limit their approval of such product candidates;

regulatory authorities may require the addition of labeling statements, such as a boxed warning or a contraindication;

we may be required to change the way such product candidates are distributed or administered, conduct additional clinical trials or change the labeling of the product candidates;

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we may be subject to regulatory investigations and government enforcement actions;

we may decide to remove such product candidates from the marketplace;

we could be sued and held liable for injury caused to individuals exposed to or taking our product candidates; and

our reputation may suffer.

We believe that any of these events could prevent us from achieving or maintaining market acceptance of the affected product candidates and could substantially increase the costs of commercializing our product candidates and significantly impact our ability to successfully commercialize our product candidates and generate revenues.

Even if we receive marketing approval for our product candidates, we may still face future development and regulatory difficulties.

Even if we receive marketing approval for our product candidates, regulatory authorities may still impose significant restrictions on our product candidates, indicated uses or marketing or impose ongoing requirements for potentially costly post-approval studies. For example, prior to product launch, the U.S. Drug Enforcement Agency, or DEA, needs to determine the controlled substance schedule of SAGE-547, taking into account the recommendation of the FDA. Our product candidates will also be subject to ongoing FDA requirements governing the labeling, packaging, storage and promotion of the product and record keeping and submission of safety and other post-market information. The FDA has significant post-marketing authority, including, for example, the authority to require labeling changes based on new safety information and to require post-marketing studies or clinical trials to evaluate serious safety risks related to the use of a drug. The FDA also has the authority to require, as part of an NDA or post-approval, the submission of a REMS. Any REMS required by the FDA may lead to increased costs to assure compliance with new post-approval regulatory requirements and potential requirements or restrictions on the sale of approved products, all of which could lead to lower sales volume and revenue.

Manufacturers of drug products and their facilities are subject to continual review and periodic inspections by the FDA and other regulatory authorities for compliance with cGMPs and other regulations. If we or a regulatory agency discover problems with our product candidates, such as adverse events of unanticipated severity or frequency, or problems with the facility where our product candidates are manufactured, a regulatory agency may impose restrictions on our product candidates, the manufacturer or us, including requiring withdrawal of our product candidates from the market or suspension of manufacturing. If we, our product candidates or the manufacturing facilities for our product candidates fail to comply with applicable regulatory requirements, a regulatory agency may, among other things:

issue warning letters or untitled letters;

seek an injunction or impose civil or criminal penalties or monetary fines;

suspend or withdraw marketing approval;

suspend any ongoing clinical trials;

refuse to approve pending applications or supplements to applications submitted by us;

suspend or impose restrictions on operations, including costly new manufacturing requirements; or

seize or detain products, refuse to permit the import or export of products, or require that we initiate a product recall.

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Competing therapies could emerge adversely affecting our opportunity to generate revenue from the sale of our product candidates.

The biopharmaceuticals industry is highly competitive. There are many public and private biopharmaceutical companies, universities, governmental agencies and other research organizations actively engaged in the research and development of products that may be similar to our product candidates or address similar markets. It is probable that the number of companies seeking to develop products and therapies similar to our products will increase.

Currently, there are no therapies specifically approved for RSE or SRSE. However, many products approved for other indications, general anesthetics and anti-seizure drugs, are used off-label for various stages of SE therapy. Additionally, though not indicated, acupuncture, hypothermia, and electroconvulsive therapy are sometimes used prior to withdrawal of care for patients with SRSE.

In the field of neuroactive steroids focused on modulation of GABA_A or NMDA receptors, our principal competitor is Marinus Pharmaceuticals, Inc., which is developing a reformulated form of Ganaxolone, a known GABA_A positive allosteric modulator neuroactive steroid, for potential treatment of drug-resistant partial complex seizures and fragile X syndrome.

Many of our potential competitors, alone or with their strategic partners, have substantially greater financial, technical and human resources than we do and significantly greater experience in the discovery and development of product candidates, obtaining FDA and other regulatory approvals of treatments and the commercialization of those treatments. Mergers and acquisitions in the biotechnology and pharmaceutical industries may result in even more resources being concentrated among a smaller number of our competitors. Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize products that are safer, more effective, have fewer or less severe side effects, are more convenient or are less expensive than any products that we may develop. Our competitors also may obtain FDA or other regulatory approval for their products more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market.

We may seek to establish collaborations, and, if we are not able to establish them on commercially reasonable terms, we may have to alter our development and commercialization plans.

Our drug development programs and the potential commercialization of our product candidates will require substantial additional cash to fund expenses. For some of our product candidates, we may decide to collaborate with pharmaceutical and biotechnology companies for the development and potential commercialization of those product candidates.

We face significant competition in seeking appropriate collaborators. Whether we reach a definitive agreement for a collaboration will depend, among other things, upon our assessment of the collaborator's resources and expertise, the terms and conditions of the proposed collaboration and the proposed collaborator's evaluation of a number of factors. Those factors may include the design or results of clinical trials, the likelihood of approval by the FDA or similar regulatory authorities outside the United States, the potential market for the subject product candidate, the costs and complexities of manufacturing and delivering such product candidate to patients, the potential of competing products, the existence of uncertainty with respect to our ownership of technology, which can exist if there is a challenge to such ownership without regard to the merits of the challenge and industry and market conditions generally. The collaborator may also consider alternative product candidates or technologies for similar indications that may be available to collaborate on and whether such collaboration could be more attractive than the one with us for our product candidate. The terms of any collaboration or other arrangements that we may establish may not be favorable to us.

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We may also be restricted under existing collaboration agreements from entering into future agreements on certain terms with potential collaborators. Collaborations are complex and time-consuming to negotiate and document. In addition, there have been a significant number of recent business combinations among large pharmaceutical companies that have resulted in a reduced number of potential future collaborators.

We may not be able to negotiate collaborations on a timely basis, on acceptable terms, or at all. If we are unable to do so, we may have to curtail the development of the product candidate for which we are seeking to collaborate, reduce or delay its development program or one or more of our other development programs, delay its potential commercialization or reduce the scope of any sales or marketing activities, or increase our expenditures and undertake development or commercialization activities at our own expense. If we elect to increase our expenditures to fund development or commercialization activities on our own, we may need to obtain additional capital, which may not be available to us on acceptable terms or at all. If we do not have sufficient funds, we may not be able to further develop our product candidates or bring them to market and generate product revenue.

In addition, any future collaborations that we enter into may not be successful. The success of our collaboration arrangements will depend heavily on the efforts and activities of our collaborators. Collaborators generally have significant discretion in determining the efforts and resources that they will apply to these collaborations. Disagreements between parties to a collaboration arrangement regarding clinical development and commercialization matters can lead to delays in the development process or commercializing the applicable product candidate and, in some cases, termination of the collaboration arrangement. These disagreements can be difficult to resolve if neither of the parties has final decision-making authority. Collaborations with pharmaceutical or biotechnology companies and other third parties often are terminated or allowed to expire by the other party. Any such termination or expiration would adversely affect us financially and could harm our business reputation.

We may not be successful in our efforts to identify or discover additional product candidates or we may expend our limited resources to pursue a particular product candidate or indication and fail to capitalize on product candidates or indications that may be more profitable or for which there is a greater likelihood of success.

The success of our business depends primarily upon our ability to identify, develop and commercialize products based on our proprietary chemistry platform. Although some of our product candidates are in non-clinical and clinical development, our research programs may fail to identify other potential product candidates for clinical development for a number of reasons. Our research methodology may be unsuccessful in identifying potential product candidates or our potential product candidates may be shown to have harmful side effects or may have other characteristics that may make the products unmarketable or unlikely to receive marketing approval.

Because we have limited financial and management resources, we focus on a limited number of research programs and product candidates and are currently focused on our SE program. As a result, we may forego or delay pursuit of opportunities with other product candidates or for other indications that later prove to have greater commercial potential. Our resource allocation decisions may cause us to fail to capitalize on viable commercial drugs or profitable market opportunities. Our spending on current and future research and development programs and product candidates for specific indications may not yield any commercially viable drugs. If we do not accurately evaluate the commercial potential or target market for a particular product candidate, we may relinquish valuable rights to that product candidate through future collaboration, licensing or other royalty arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate.

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If any of these events occur, we may be forced to abandon our development efforts for a program or programs, which would have a material adverse effect on our business and could potentially cause us to cease operations. Research programs to identify new product candidates require substantial technical, financial and human resources. We may focus our efforts and resources on potential programs or product candidates that ultimately prove to be unsuccessful.

We are subject to healthcare laws and regulations, which could expose us to criminal sanctions, civil penalties, contractual damages, reputational harm and diminished profits and future earnings.

Although we do not currently have any products on the market, once we begin commercializing our products, we may be subject to additional healthcare statutory and regulatory requirements and enforcement by the federal government and the states and foreign governments in which we conduct our business. Healthcare providers, physicians and others will play a primary role in the recommendation and prescription of our product candidates, if approved. Our future arrangements with third-party payors will expose us to broadly applicable fraud and abuse and other healthcare laws and regulations that may constrain the business or financial arrangements and relationships through which we market, sell and distribute our product candidates, if we obtain marketing approval. Restrictions under applicable federal and state healthcare laws and regulations include the following:

The federal anti-kickback statute prohibits, among other things, persons from knowingly and willfully soliciting, offering, receiving or providing remuneration, directly or indirectly, in cash or in kind, to induce or reward either the referral of an individual for, or the purchase, order or recommendation of, any good or service, for which payment may be made under federal healthcare programs such as Medicare and Medicaid.

The federal False Claims Act imposes criminal and civil penalties, including those from civil whistleblower or qui tam actions, against individuals or entities for knowingly presenting, or causing to be presented, to the federal government, claims for payment that are false or fraudulent or making a false statement to avoid, decrease, or conceal an obligation to pay money to the federal government.

The federal Health Insurance Portability and Accountability Act of 1996, as amended by the Health Information Technology for Economic and Clinical Health Act, imposes criminal and civil liability for executing a scheme to defraud any healthcare benefit program and also imposes obligations, including mandatory contractual terms, with respect to safeguarding the privacy, security and transmission of individually identifiable health information.

The federal false statements statute prohibits knowingly and willfully falsifying, concealing or covering up a material fact or making any materially false statement in connection with the delivery of or payment for healthcare benefits, items or services.

The federal transparency requirements, sometimes referred to as the Sunshine Act, under the Patient Protection and Affordable Care Act, require manufacturers of drugs, devices, biologics and medical supplies that are reimbursable under Medicare, Medicaid, or the Children's Health Insurance Program to report to the Department of Health and Human Services information related to physician payments and other transfers of value and physician ownership and investment interests.

Analogous state laws and regulations, such as state anti-kickback and false claims laws and transparency laws, may apply to sales or marketing arrangements and claims involving healthcare items or services reimbursed by non-governmental third-party payors, including private insurers, and some state laws require pharmaceutical companies to comply with the pharmaceutical industry's voluntary compliance guidelines and the relevant compliance.

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Guidance promulgated by the federal government in addition to requiring drug manufacturers to report information related to payments to physicians and other healthcare providers or marketing expenditures and drug pricing.

Ensuring that our future business arrangements with third parties comply with applicable healthcare laws and regulations could be costly. It is possible that governmental authorities will conclude that our business practices do not comply with current or future statutes, regulations or case law involving applicable fraud and abuse or other healthcare laws and regulations. If our operations, including anticipated activities to be conducted by our sales team, were found to be in violation of any of these laws or any other governmental regulations that may apply to us, we may be subject to significant civil, criminal and administrative penalties, damages, fines and exclusion from government funded healthcare programs, such as Medicare and Medicaid, any of which could substantially disrupt our operations. If any of the physicians or other providers or entities with whom we expect to do business is found not to be in compliance with applicable laws, they may be subject to criminal, civil or administrative sanctions, including exclusions from government funded healthcare programs.

The FDA and other regulatory agencies actively enforce the laws and regulations prohibiting the promotion of off-label uses. If we are found to have improperly promoted off-label uses, we may become subject to significant liability.

The FDA and other regulatory agencies strictly regulate the promotional claims that may be made about prescription products, such as SAGE-547, SAGE-689, and SAGE-217, if approved. In particular, a product may not be promoted for uses that are not approved by the FDA or such other regulatory agencies as reflected in the product's approved labeling. For example, if we receive marketing approval for SAGE-547 as a treatment for SRSE, physicians may nevertheless prescribe SAGE-547 to their patients in a manner that is inconsistent with the approved label. If we are found to have promoted such off-label uses, we may become subject to significant liability. The federal government has levied large civil and criminal fines against companies for alleged improper promotion and has enjoined several companies from engaging in off-label promotion. The FDA has also requested that companies enter into consent decrees or permanent injunctions under which specified promotional conduct is changed or curtailed. If we cannot successfully manage the promotion of our product candidates, if approved, we could become subject to significant liability, which would materially adversely affect our business and financial condition.

SAGE-547 will, and our other product candidates may, contain controlled substances, the manufacture, use, sale, importation, exportation, prescribing and distribution of which are subject to regulation by the DEA.

Before we can commercialize SAGE-547, and potentially our other product candidates, the DEA will need to determine the controlled substance schedule, taking into account the recommendation of the FDA. This may be a lengthy process that could delay our marketing of a product candidate and could potentially diminish any regulatory exclusivity periods for which we may be eligible. SAGE-547 will, and our other product candidates may, if approved, be regulated as controlled substances as defined in the Controlled Substances Act of 1970, or CSA, and the implementing regulations of the DEA, which establish registration, security, recordkeeping, reporting, storage, distribution, importation, exportation, inventory, quota and other requirements administered by the DEA. These requirements are applicable to us, to our third-party manufacturers and to distributors, prescribers and dispensers of our product candidates. The DEA regulates the handling of controlled substances through a closed chain of distribution. This control extends to the equipment and raw materials used in their manufacture and packaging, in order to prevent loss and diversion into illicit channels of commerce. A number of states and foreign countries also independently regulate these drugs as controlled substances.

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The DEA regulates controlled substances as Schedule I, II, III, IV or V substances. Schedule I substances by definition have no established medicinal use, and may not be marketed or sold in the United States. A pharmaceutical product may be listed as Schedule II, III, IV or V, with Schedule II substances considered to present the highest risk of abuse and Schedule V substances the lowest relative risk of abuse among such substances.

We expect that SAGE-547 will, and our other product candidates may, be listed by the DEA as Schedule IV controlled substances under the CSA. Consequently, the manufacturing, shipping, storing, selling and using of the products will be subject to a high degree of regulation. Also, distribution, prescribing and dispensing of these drugs are highly regulated.

Annual registration is required for any facility that manufactures, distributes, dispenses, imports or exports any controlled substance. The registration is specific to the particular location, activity and controlled substance schedule.

Because of their restrictive nature, these laws and regulations could limit commercialization of our product candidates containing controlled substances. Failure to comply with these laws and regulations could also result in withdrawal of our DEA registrations, disruption in manufacturing and distribution activities, consent decrees, criminal and civil penalties and state actions, among other consequences.

Even if approved, reimbursement policies could limit our ability to sell our product candidates.

Market acceptance and sales of our product candidates will depend on reimbursement policies and may be affected by healthcare reform measures. Government authorities and third-party payors, such as private health insurers and health maintenance organizations, decide which medications they will pay for and establish reimbursement levels for those medications. Cost containment is a primary concern in the U.S. healthcare industry and elsewhere. Government authorities and these third-party payors have attempted to control costs by limiting coverage and the amount of reimbursement for particular medications. We cannot be sure that reimbursement will be available for our product candidates and, if reimbursement is available, the level of such reimbursement. Reimbursement may impact the demand for, or the price of, our product candidates. If reimbursement is not available or is available only at limited levels, we may not be able to successfully commercialize our product candidates.

In some foreign countries, particularly in Canada and European countries, the pricing of prescription pharmaceuticals is subject to strict governmental control. In these countries, pricing negotiations with governmental authorities can take six months or longer after the receipt of regulatory approval and product launch. To obtain favorable reimbursement for the indications sought or pricing approval in some countries, we may be required to conduct a clinical trial that compares the cost-effectiveness of our product candidates with other available therapies. If reimbursement for our product candidates is unavailable in any country in which we seek reimbursement, if it is limited in scope or amount, if it is conditioned upon our completion of additional clinical trials, or if pricing is set at unsatisfactory levels, our operating results could be materially adversely affected.

Even though we have obtained orphan drug designation for SAGE-547 as a treatment for SE, there may be limits to the regulatory exclusivity afforded by such designation.

Even though we have obtained orphan drug designation for SAGE-547 for treatment of SE from the FDA, there are limitations to exclusivity afforded by such designation. In the United States, the company that first obtains FDA approval for a designated orphan drug for the specified rare disease or condition receives orphan drug marketing exclusivity for that drug for a period of seven years. This orphan drug exclusivity prevents the FDA from approving another application, including a full NDA to

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market the same drug for the same orphan indication, except in very limited circumstances, including when the FDA concludes that the later drug is safer, more effective or makes a major contribution to patient care. For purposes of small molecule drugs, the FDA defines "same drug" as a drug that contains the same active moiety and is intended for the same use as the drug in question. To obtain orphan drug exclusivity for a drug that shares the same active moiety as an already approved drug, it must be demonstrated to the FDA that the drug is safer or more effective than the approved orphan designated drug, or that it makes a major contribution to patient care. In addition, a designated orphan drug may not receive orphan drug exclusivity if it is approved for a use that is broader than the indication for which it received orphan designation. In addition, orphan drug exclusive marketing rights in the United States may be lost if the FDA later determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantity of the drug to meet the needs of patients with the rare disease or condition or if another drug with the same active moiety is determined to be safer, more effective, or represents a major contribution to patient care.

Our future growth may depend, in part, on our ability to penetrate foreign markets, where we would be subject to additional regulatory burdens and other risks and uncertainties.

Our future profitability may depend, in part, on our ability to commercialize our product candidates in foreign markets for which we may rely on collaboration with third parties. If we commercialize our product candidates in foreign markets, we would be subject to additional risks and uncertainties, including:

our customers' ability to obtain reimbursement for our product candidates in foreign markets;

our inability to directly control commercial activities because we are relying on third parties;

the burden of complying with complex and changing foreign regulatory, tax, accounting and legal requirements;

different medical practices and customs in foreign countries affecting acceptance in the marketplace;

import or export licensing requirements;

longer accounts receivable collection times;

longer lead times for shipping;

language barriers for technical training;

reduced protection of intellectual property rights in some foreign countries;

the existence of additional potentially relevant third party intellectual property rights;

foreign currency exchange rate fluctuations; and

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the interpretation of contractual provisions governed by foreign laws in the event of a contract dispute. Foreign sales of our product candidates could also be adversely affected by the imposition of governmental controls, political and economic instability, trade restrictions and changes in tariffs.

Risks Related to Our Intellectual Property Rights

If we are unable to adequately protect our proprietary technology, or obtain and maintain issued patents that are sufficient to protect our product candidates, others could compete against us more directly, which would have a material adverse impact on our business, results of operations, financial condition and prospects.

We strive to protect and enhance the proprietary technologies that we believe are important to our business, including seeking patents intended to cover our products and compositions, their

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methods of use and any other inventions that are important to the development of our business. We also rely on trade secrets to protect aspects of our business that are not amenable to, or that we do not consider appropriate for, patent protection.

Our success will depend significantly on our ability to obtain and maintain patent and other proprietary protection for commercially important technology, inventions and know-how related to our business, defend and enforce our patents, should they issue, preserve the confidentiality of our trade secrets and operate without infringing the valid and enforceable patents and proprietary rights of third parties. We also rely on know-how, continuing technological innovation and in-licensing opportunities to develop, strengthen and maintain the proprietary position of our product candidates. Our owned and licensed patent applications relate to SAGE-547, GABA_A receptor modulators, including genus and species claims to SAGE-689 and NMDA receptor modulators.

We currently have no issued patents covering any of our lead product candidates, SAGE-547, SAGE-689, or SAGE-217. We cannot provide any assurances that any of our pending patent applications will mature into issued patents and, if they do, that such patents will include, claims with a scope sufficient to protect our product candidates or otherwise provide any competitive advantage. For example, the patent applications that may provide coverage for SAGE-547, only cover particular formulations and particular methods of using such formulations to treat seizure conditions, such as SE. As a result, if a patent issues from such patent applications, it would not prevent third-party competitors from creating, making and marketing alternative formulations, that fall outside the scope of our patent claims or practicing alternative methods. There can be no assurance that any such alternative formulations will not be equally effective as our formulation of SAGE-547. Moreover, other parties have developed technologies that may be related or competitive to our approach, and may have filed or may file patent applications and may have received or may receive patents that may overlap or conflict with our patent applications, either by claiming the same methods or formulations or by claiming subject matter that could dominate our patent position. Such third-party patent positions may limit or even eliminate our ability to obtain patent protection for certain inventions.

The patent positions of biotechnology and pharmaceutical companies, including our patent position, involve complex legal and factual questions, and, therefore, the issuance, scope, validity and enforceability of any patent claims that we may obtain cannot be predicted with certainty. Patents, if issued, may be challenged, deemed unenforceable, invalidated, or circumvented. U.S. patents and patent applications may also be subject to interference proceedings, ex parte reexamination, or inter partes review proceedings, supplemental examination and challenges in district court. Patents may be subjected to opposition, post-grant review, or comparable proceedings lodged in various foreign, both national and regional, patent offices. These proceedings could result in either loss of the patent or denial of the patent application or loss or reduction in the scope of one or more of the claims of the patent or patent application. In addition, such proceedings may be costly. Thus, any patents, should they issue, that we may own or exclusively license may not provide any protection against competitors. Furthermore, an adverse decision in an interference proceeding can result in a third party receiving the patent right sought by us, which in turn could affect our ability to develop, market or otherwise commercialize our product candidates.

Furthermore, though a patent, if it were to issue, is presumed valid and enforceable, its issuance is not conclusive as to its validity or its enforceability and it may not provide us with adequate proprietary protection or competitive advantages against competitors with similar products. Even if a patent issues and is held to be valid and enforceable, competitors may be able to design around our patents, such as using pre-existing or newly developed technology. Other parties may develop and obtain patent protection for more effective technologies, designs or methods. We may not be able to prevent the unauthorized disclosure or use of our technical knowledge or trade secrets by consultants, vendors, former employees and current employees. The laws of some foreign countries do not protect

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our proprietary rights to the same extent as the laws of the United States, and we may encounter significant problems in protecting our proprietary rights in these countries. If these developments were to occur, they could have a material adverse effect on our sales.

Our ability to enforce our patent rights depends on our ability to detect infringement. It is difficult to detect infringers who do not advertise the components that are used in their products. Moreover, it may be difficult or impossible to obtain evidence of infringement in a competitor's or potential competitor's product. Any litigation to enforce or defend our patent rights, even if we were to prevail, could be costly and time-consuming and would divert the attention of our management and key personnel from our business operations. We may not prevail in any lawsuits that we initiate and the damages or other remedies awarded if we were to prevail may not be commercially meaningful.

In addition, proceedings to enforce or defend our patents, if and when issued, could put our patents at risk of being invalidated, held unenforceable, or interpreted narrowly. Such proceedings could also provoke third parties to assert claims against us, including that some or all of the claims in one or more of our patents are invalid or otherwise unenforceable. If any of our patents, if and when issued, covering our product candidates are invalidated or found unenforceable, our financial position and results of operations would be materially and adversely impacted. In addition, if a court found that valid, enforceable patents held by third parties covered our product candidates, our financial position and results of operations would also be materially and adversely impacted.

The degree of future protection for our proprietary rights is uncertain, and we cannot ensure that:

any of our pending patent applications, if issued, will include claims having a scope sufficient to protect our product candidates or any other products or product candidates;

any of our pending patent applications will issue as patents at all;

we will be able to successfully commercialize our product candidates, if approved, before our relevant patents expire;

we were the first to make the inventions covered by each of our patents and pending patent applications;

we were the first to file patent applications for these inventions;

others will not develop similar or alternative technologies that do not infringe our patents;

others will not use pre-existing technology to effectively compete against us;

any of our patents, if issued, will be found to ultimately be valid and enforceable;

any patents issued to us will provide a basis for an exclusive market for our commercially viable products, will provide us with any competitive advantages or will not be challenged by third parties;

we will develop additional proprietary technologies or product candidates that are separately patentable; or

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that our commercial activities or products will not infringe upon the patents or proprietary rights of others.

We rely upon unpatented trade secrets, unpatented know-how and continuing technological innovation to develop and maintain our competitive position, which we seek to protect, in part, by confidentiality agreements with our employees and our collaborators and consultants. It is possible that technology relevant to our business will be independently developed by a person that is not a party to such an agreement. Furthermore, if the employees and consultants who are parties to these

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agreements breach or violate the terms of these agreements, we may not have adequate remedies for any such breach or violation, and we could lose our trade secrets through such breaches or violations. Further, our trade secrets could otherwise become known or be independently discovered by our competitors.

We may infringe the intellectual property rights of others, which may prevent or delay our product development efforts and stop us from commercializing or increase the costs of commercializing our product candidates, if approved.

Our success will depend in part on our ability to operate without infringing the intellectual property and proprietary rights of third parties. We cannot assure you that our business, products and methods do not or will not infringe the patents or other intellectual property rights of third parties.

The pharmaceutical industry is characterized by extensive litigation regarding patents and other intellectual property rights. Other parties may allege that our product candidates or the use of our technologies infringes patent claims or other intellectual property rights held by them or that we are employing their proprietary technology without authorization. As we continue to develop and, if approved, commercialize our current product candidates and future product candidates, competitors may claim that our technology infringes their intellectual property rights as part of business strategies designed to impede our successful commercialization. There may be third-party patents or patent applications with claims to materials, formulations, methods of manufacture or methods for treatment related to the use or manufacture of our product candidates. Because patent applications can take many years to issue, third parties may have currently pending patent applications which may later result in issued patents that our product candidates may infringe, or which such third parties claim are infringed by our technologies. The outcome of intellectual property litigation is subject to uncertainties that cannot be adequately quantified in advance. The coverage of patents is subject to interpretation by the courts, and the interpretation is not always uniform. If we are sued for patent infringement, we would need to demonstrate that our product candidates, products or methods either do not infringe the patent claims of the relevant patent or that the patent claims are invalid, and we may not be able to do this. Even if we are successful in these proceedings, we may incur substantial costs and the time and attention of our management and scientific personnel could be diverted in pursuing these proceedings, which could have a material adverse effect on us. In addition, we may not have sufficient resources to bring these actions to a successful conclusion.

Patent and other types of intellectual property litigation can involve complex factual and legal questions, and their outcome is uncertain. Any claim relating to intellectual property infringement that is successfully asserted against us may require us to pay substantial damages, including treble damages and attorney's fees if we are found to be willfully infringing another party's patents, for past use of the asserted intellectual property and royalties and other consideration going forward if we are forced to take a license. In addition, if any such claim were successfully asserted against us and we could not obtain such a license, we may be forced to stop or delay developing, manufacturing, selling or otherwise commercializing our product candidates.

Even if we are successful in these proceedings, we may incur substantial costs and divert management time and attention in pursuing these proceedings, which could have a material adverse effect on us. If we are unable to avoid infringing the patent rights of others, we may be required to seek a license, defend an infringement action or challenge the validity of the patents in court, or redesign our products. Patent litigation is costly and time-consuming. We may not have sufficient resources to bring these actions to a successful conclusion. In addition, intellectual property litigation or claims could force us to do one or more of the following:

cease developing, selling or otherwise commercializing our product candidates;

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pay substantial damages for past use of the asserted intellectual property;

obtain a license from the holder of the asserted intellectual property, which license may not be available on reasonable terms, if at all;
and

in the case of trademark claims, redesign, or rename, some or all of our product candidates to avoid infringing the intellectual property rights of third parties, which may not be possible and, even if possible, could be costly and time-consuming.

Any of these risks coming to fruition could have a material adverse effect on our business, results of operations, financial condition and prospects.

We may be subject to claims challenging the inventorship or ownership of our patents and other intellectual property.

We enter into confidentiality and intellectual property assignment agreements with our employees, consultants, outside scientific collaborators, sponsored researchers and other advisors. These agreements generally provide that inventions conceived by the party in the course of rendering services to us will be our exclusive property. However, these agreements may not be honored and may not effectively assign intellectual property rights to us. For example, even if we have a consulting agreement in place with an academic advisor pursuant to which such academic advisor is required to assign any inventions developed in connection with providing services to us, such academic advisor may not have the right to assign such inventions to us, as it may conflict with his or her obligations to assign all such intellectual property to his or her employing institution.

Litigation may be necessary to defend against these and other claims challenging inventorship or ownership. If we fail in defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights, such as exclusive ownership of, or right to use, valuable intellectual property. Such an outcome could have a material adverse effect on our business. Even if we are successful in defending against such claims, litigation could result in substantial costs and be a distraction to management and other employees.

Obtaining and maintaining our patent protection depends on compliance with various procedural, document submission, fee payment and other requirements imposed by governmental patent agencies, and our patent protection could be reduced or eliminated for non-compliance with these requirements.

The U.S. Patent and Trademark Office, or U.S. PTO, and various foreign governmental patent agencies require compliance with a number of procedural, documentary, fee payment and other provisions during the patent process. There are situations in which noncompliance can result in abandonment or lapse of a patent or patent application, resulting in partial or complete loss of patent rights in the relevant jurisdiction. In such an event, competitors might be able to enter the market earlier than would otherwise have been the case.

We may be involved in lawsuits to protect or enforce our patents or the patents of our licensors, which could be expensive, time-consuming and unsuccessful.

Even if the patent applications we own or license are issued, competitors may infringe these patents. To counter infringement or unauthorized use, we may be required to file infringement claims, which can be expensive and time-consuming. In addition, in an infringement proceeding, a court may decide that a patent of ours or our licensors is not valid, is unenforceable and/or is not infringed, or may refuse to stop the other party from using the technology at issue on the grounds that our patents do not cover the technology in question. An adverse result in any litigation or defense proceedings could put one or more of our patents at risk of being invalidated or interpreted narrowly and could put our patent applications at risk of not issuing.

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Interference proceedings provoked by third parties or brought by us may be necessary to determine the priority of inventions with respect to our patents or patent applications or those of our licensors. An unfavorable outcome could require us to cease using the related technology or to attempt to license rights to it from the prevailing party. Our business could be harmed if the prevailing party does not offer us a license on commercially reasonable terms. Our defense of litigation or interference proceedings may fail and, even if successful, may result in substantial costs and distract our management and other employees. We may not be able to prevent, alone or with our licensors, misappropriation of our intellectual property rights, particularly in countries where the laws may not protect those rights as fully as in the United States.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation. There could also be public announcements of the results of hearings, motions or other interim proceedings or developments. If securities analysts or investors perceive these results to be negative, it could have a material adverse effect on the price of our common stock.

Issued patents covering our product candidates could be found invalid or unenforceable if challenged in court.

If we or one of our licensing partners initiated legal proceedings against a third party to enforce a patent, if and when issued, covering one of our product candidates, the defendant could counterclaim that the patent covering our product candidate is invalid and/or unenforceable. In patent litigation in the United States, defendant counterclaims alleging invalidity and/or unenforceability are commonplace. Grounds for a validity challenge include alleged failures to meet any of several statutory requirements, including lack of novelty, obviousness or non-enablement. Grounds for unenforceability assertions include allegations that someone connected with prosecution of the patent withheld relevant information from the U.S. PTO, or made a misleading statement, during prosecution. Third parties may also raise similar claims before administrative bodies in the United States or abroad, even outside the context of litigation. Such mechanisms include re-examination, post grant review and equivalent proceedings in foreign jurisdictions, e.g., opposition proceedings. Such proceedings could result in revocation or amendment of our patents in such a way that they no longer cover our product candidates or competitive products. The outcome following legal assertions of invalidity and unenforceability is unpredictable. With respect to validity, for example, we cannot be certain that there is no invalidating prior art, of which we and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity and/or unenforceability, we would lose at least part, and perhaps all, of the patent protection on our product candidates. Such a loss of patent protection would have a material adverse impact on our business.

We will not seek to protect our intellectual property rights in all jurisdictions throughout the world and we may not be able to adequately enforce our intellectual property rights even in the jurisdictions where we seek protection.

Filing, prosecuting and defending patents on product candidates in all countries and jurisdictions throughout the world would be prohibitively expensive, and our intellectual property rights in some countries outside the United States could be less extensive than those in the United States, assuming that rights are obtained in the United States. In addition, the laws of some foreign countries do not protect intellectual property rights to the same extent as federal and state laws in the United States. Consequently, we may not be able to prevent third parties from practicing our inventions in all countries outside the United States, or from selling or importing products made using our inventions in and into the United States or other jurisdictions. The statutory deadlines for pursuing patent protection in individual foreign jurisdictions are based on the priority date of each of our patent applications. For the

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patent families related to SAGE-689 and SAGE-217, and many of the other patent families that we own or license, the relevant statutory deadlines have not yet expired. For each of the patent families that we believe provide coverage for our lead product candidates, we will need to decide whether and where to pursue protection outside the United States.

Competitors may use our technologies in jurisdictions where we do not pursue and obtain patent protection to develop their own products and further, may export otherwise infringing products to territories where we have patent protection, but enforcement is not as strong as that in the United States. These products may compete with our products and our patents or other intellectual property rights may not be effective or sufficient to prevent them from competing. Even if we pursue and obtain issued patents in particular jurisdictions, our patent claims or other intellectual property rights may not be effective or sufficient to prevent third parties from so competing.

The laws of some foreign countries do not protect intellectual property rights to the same extent as the laws of the United States. Many companies have encountered significant problems in protecting and defending intellectual property rights in certain foreign jurisdictions. The legal systems of some countries, particularly developing countries, do not favor the enforcement of patents and other intellectual property protection, especially those relating to biotechnology. This could make it difficult for us to stop the infringement of our patents, if obtained, or the misappropriation of our other intellectual property rights. For example, many foreign countries have compulsory licensing laws under which a patent owner must grant licenses to third parties. In addition, many countries limit the enforceability of patents against third parties, including government agencies or government contractors. In these countries, patents may provide limited or no benefit. Patent protection must ultimately be sought on a country-by-country basis, which is an expensive and time-consuming process with uncertain outcomes. Accordingly, we may choose not to seek patent protection in certain countries, and we will not have the benefit of patent protection in such countries.

Furthermore, proceedings to enforce our patent rights in foreign jurisdictions could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly, could put our patent applications at risk of not issuing and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate and the damages or other remedies awarded, if any, may not be commercially meaningful. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license.

We are dependent on licensed intellectual property. If we were to lose our rights to licensed intellectual property, we may not be able to continue developing or commercializing our product candidates, if approved. If we breach any of the agreements under which we license the use, development and commercialization rights to our product candidates or technology from third parties or, in certain cases, we fail to meet certain development deadlines, we could lose license rights that are important to our business.

We are a party to a number of license agreements under which we are granted rights to intellectual property that are important to our business and we expect that we may need to enter into additional license agreements in the future. Our existing license agreements impose, and we expect that future license agreements will impose on us, various development, regulatory and/or commercial diligence obligations, payment of milestones and/or royalties and other obligations. If we fail to comply with our obligations under these agreements, or we are subject to a bankruptcy, the licensor may have the right to terminate the license, in which event we would not be able to market products covered by the license. Our business could suffer, for example, if any current or future licenses terminate, if the licensors fail to abide by the terms of the license, if the licensed patents or other rights are found to be

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invalid or unenforceable, or if we are unable to enter into necessary licenses on acceptable terms. See **Business Licenses** for a description of our license agreements, which includes a description of the termination provisions of these agreements.

As we have done previously, we may need to obtain licenses from third parties to advance our research or allow commercialization of our product candidates, and we cannot provide any assurances that third-party patents do not exist that might be enforced against our current product candidates or future products in the absence of such a license. We may fail to obtain any of these licenses on commercially reasonable terms, if at all. Even if we are able to obtain a license, it may be non-exclusive, thereby giving our competitors access to the same technologies licensed to us. In that event, we may be required to expend significant time and resources to develop or license replacement technology. If we are unable to do so, we may be unable to develop or commercialize the affected product candidates, which could materially harm our business and the third parties owning such intellectual property rights could seek either an injunction prohibiting our sales, or, with respect to our sales, an obligation on our part to pay royalties and/or other forms of compensation.

Licensing of intellectual property is of critical importance to our business and involves complex legal, business and scientific issues. Disputes may arise between us and our licensors regarding intellectual property subject to a license agreement, including:

the scope of rights granted under the license agreement and other interpretation-related issues;

whether and the extent to which our technology and processes infringe on intellectual property of the licensor that is not subject to the licensing agreement;

our right to sublicense patent and other rights to third parties under collaborative development relationships;

our diligence obligations with respect to the use of the licensed technology in relation to our development and commercialization of our product candidates, and what activities satisfy those diligence obligations; and

the ownership of inventions and know-how resulting from the joint creation or use of intellectual property by our licensors and us and our partners.

If disputes over intellectual property that we have licensed prevent or impair our ability to maintain our current licensing arrangements on acceptable terms, we may be unable to successfully develop and commercialize the affected product candidates.

We have entered into several licenses to support our various programs. We completed an exclusive license agreement with Washington University, or WU, under certain patent families that comprise a variety of small molecule allosteric modulators of GABA_A receptors and for which we have the worldwide right to develop and commercialize. A patent family that discloses and claims SAGE-689 is licensed to us under this agreement. We are obligated to pay WU certain clinical/regulatory milestones and single-digit royalties on products developed from this technology. Termination of our license agreement with WU would have a material adverse impact on our ability to develop and commercialize SAGE-689.

We have also entered into an exclusive license agreement with CyDex Pharmaceuticals, Inc., or CyDex, a wholly owned subsidiary of Ligand Pharmaceuticals, Inc., to use its Captisol technology to develop SAGE-547 for the field of use, which includes all fields for the treatment, prevention or diagnosis of any disease or symptom in humans or animals. We are obligated to pay CyDex certain clinical/regulatory milestones and single-digit royalties on SAGE-547. In addition, we entered into a supply agreement with CyDex, pursuant to which they supply us with Captisol to formulate SAGE-547.

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Absent an alternative agreement by the parties, our rights under our exclusive license agreement terminate in the event that the supply agreement terminates. Currently, our SAGE-547 product candidate in clinical development is formulated in Captisol. Termination of our license agreement with CyDex would have a material adverse impact on our ability to develop and commercialize SAGE-547 in its current formulation.

We also entered into a non-exclusive license with The Regents of the University of California, or the Regents. Pursuant to this agreement the Regents granted us a non-exclusive, non-transferable license under all personal property rights of the Regents covering the tangible personal property in an IND application package owned by the Regents, or the Data, and a specified quantity of cGMP grade allopregnanolone, or the Material, to (i) use the Data for reference or incorporation in an IND for use of the Material as a treatment of SE, essential tremor and/or postpartum depression and (ii) use the Material or modifications of the Material to develop a pharmaceutical formulation for clinical trials for SE, essential tremor and/or postpartum depression. This agreement requires us to pay milestone payments in connection with the first derived product, which would include SAGE-547, that meets the relevant milestones and we must also pay single-digit royalties for each derived product for a period of 15 years following the first commercial sale of such derived product. Termination of our license agreement with the Regents would have a material adverse impact on our ability to develop and commercialize derived products, which would include SAGE-547.

We may enter into additional license(s) to third-party intellectual property that are necessary or useful to our business. Our current licenses and any future licenses that we may enter into impose various royalty payment, milestone, and other obligations on us. For example, as is the case for the Washington University license, the licensor may retain control over patent prosecution and maintenance under a license agreement, in which case, we may not be able to adequately influence patent prosecution or prevent inadvertent lapses of coverage due to failure to pay maintenance fees. If we fail to comply with any of our obligations under a current or future license agreement, our licensor(s) may allege that we have breached our license agreement and may accordingly seek to terminate our license with them. In addition, future licensor(s) may decide to terminate our license at will. Termination of any of our current or future licenses could result in our loss of the right to use the licensed intellectual property, which could materially adversely affect our ability to develop and commercialize a product candidate or product, if approved, as well as harm our competitive business position and our business prospects.

In addition, if our licensors fail to abide by the terms of the license, if the licensors fail to prevent infringement by third parties, if the licensed patents or other rights are found to be invalid or unenforceable, or if we are unable to enter into necessary licenses on acceptable terms our business could suffer.

Some intellectual property which we have licensed may have been discovered through government funded programs and thus may be subject to federal regulations such as march-in rights, certain reporting requirements, and a preference for U.S. industry. Compliance with such regulations may limit our exclusive rights, subject us to expenditure of resources with respect to reporting requirements, and limit our ability to contract with non-U.S. manufacturers.

Some of the intellectual property rights we have licensed may have been generated through the use of U.S. government funding and may therefore be subject to certain federal regulations. For example, some of the intellectual property rights licensed to us under the license agreements with WU and the Regents may have been generated using U.S. government funds. As a result, the U.S. government may have certain rights to intellectual property embodied in our current or future product candidates pursuant to the Bayh-Dole Act of 1980, or Bayh-Dole Act. These U.S. government rights in certain inventions developed under a government-funded program include a non-exclusive, non-transferable, irrevocable worldwide license to use inventions for any governmental purpose. In

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addition, the U.S. government has the right to require us to grant exclusive, partially exclusive, or non-exclusive licenses to any of these inventions to a third party if it determines that: (i) adequate steps have not been taken to commercialize the invention; (ii) government action is necessary to meet public health or safety needs; or (iii) government action is necessary to meet requirements for public use under federal regulations (also referred to as "march-in rights"). The U.S. government also has the right to take title to these inventions if we fail, or the applicable licensor fails, to disclose the invention to the government and fail to file an application to register the intellectual property within specified time limits. In addition, the U.S. government may acquire title to these inventions in any country in which a patent application is not filed within specified time limits. Intellectual property generated under a government funded program is also subject to certain reporting requirements, compliance with which may require us, or the applicable licensor, to expend substantial resources. In addition, the U.S. government requires that any products embodying the subject invention or produced through the use of the subject invention be manufactured substantially in the United States. The manufacturing preference requirement can be waived if the owner of the intellectual property can show that reasonable but unsuccessful efforts have been made to grant licenses on similar terms to potential licensees that would be likely to manufacture substantially in the United States or that under the circumstances domestic manufacture is not commercially feasible. This preference for U.S. manufacturers may limit our ability to contract with non-U.S. product manufacturers for products covered by such intellectual property.

We currently do not plan to apply for additional U.S. government funding, but if we do, and we discover compounds or drug candidates as a result of such funding, intellectual property rights to such discoveries may be subject to the applicable provisions of the Bayh-Dole Act.

If we do not obtain additional protection under the Hatch-Waxman Amendments and similar foreign legislation by extending the patent terms and obtaining data exclusivity for our product candidates, our business may be materially harmed.

Depending upon the timing, duration and specifics of FDA marketing approval of our product candidates, one or more of the U.S. patents we own or license may be eligible for limited patent term restoration under the Drug Price Competition and Patent Term Restoration Act of 1984, referred to as the Hatch-Waxman Amendments. The Hatch-Waxman Amendments permit a patent restoration term of up to five years as compensation for patent term lost during product development and the FDA regulatory review process. However, we may not be granted an extension because of, for example, failing to apply within applicable deadlines, failing to apply prior to expiration of relevant patents or otherwise failing to satisfy applicable requirements. For example, we may not be granted an extension if the active ingredient of SAGE-547, allopregnanolone, is used in another drug company's product candidate and that product candidate is the first to obtain FDA approval. Moreover, the applicable time period or the scope of patent protection afforded could be less than we request. If we are unable to obtain patent term extension or restoration or the term of any such extension is less than we request, our competitors may obtain approval of competing products following our patent expiration, and our ability to generate revenues could be materially adversely affected.

Changes in U.S. patent law could diminish the value of patents in general, thereby impairing our ability to protect our products.

As is the case with other biotechnology companies, our success is heavily dependent on intellectual property, particularly patents. Obtaining and enforcing patents in the biotechnology industry involve both technological and legal complexity, and is therefore costly, time-consuming and inherently uncertain. In addition, the United States has recently enacted and is currently implementing wide-ranging patent reform legislation: the Leahy-Smith America Invents Act, referred to as the America Invents Act. The America Invents Act includes a number of significant changes to U.S. patent law.

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These include provisions that affect the way patent applications will be prosecuted and may also affect patent litigation. It is not yet clear what, if any, impact the America Invents Act will have on the operation of our business. However, the America Invents Act and its implementation could increase the uncertainties and costs surrounding the prosecution of our patent applications and the enforcement or defense of any patents that may issue from our patent applications, all of which could have a material adverse effect on our business and financial condition.

In addition, recent U.S. Supreme Court rulings have narrowed the scope of patent protection available in certain circumstances and weakened the rights of patent owners in certain situations. The full impact of these decisions is not yet known. For example, on March 20, 2012 in *Mayo Collaborative Services, DBA Mayo Medical Laboratories, et al. v. Prometheus Laboratories, Inc.*, the Court held that several claims drawn to measuring drug metabolite levels from patient samples and correlating them to drug doses were not patentable subject matter. The decision appears to impact diagnostics patents that merely apply a law of nature via a series of routine steps and it has created uncertainty around the ability to obtain patent protection for certain inventions. Additionally, on June 13, 2013 in *Association for Molecular Pathology v. Myriad Genetics, Inc.*, the Court held that claims to isolated genomic DNA are not patentable, but claims to complementary DNA molecules are patent eligible because they are not a natural product. The effect of the decision on patents for other isolated natural products is uncertain. In *Alice Corporation Pty. Ltd. v. CLS Bank International, et al.*, a case involving patent claims directed to a method for mitigating settlement risk, the Court held that the patent eligibility of claims directed to abstract ideas, products of nature, and laws of nature should be determined using the same framework set forth in *Prometheus*. The U.S. PTO recently issued a set of guidelines setting forth procedures for determining subject matter eligibility of claims directed to abstract ideas, products of nature, and laws of nature in line with the *Prometheus*, *Myriad*, and *Alice* decisions. The guidance does not limit the application of *Myriad* to DNA but, rather, applies the decision to other natural products.

In addition to increasing uncertainty with regard to our ability to obtain future patents, this combination of events has created uncertainty with respect to the value of patents, once obtained. Depending on these and other decisions by the U.S. Congress, the federal courts and the U.S. PTO, the laws and regulations governing patents could change in unpredictable ways that would weaken our ability to obtain new patents or to enforce any patents that may issue in the future.

We may be subject to damages resulting from claims that we or our employees have wrongfully used or disclosed alleged trade secrets of their former employers.

Our employees have been previously employed at other biotechnology or pharmaceutical companies, including our competitors or potential competitors. We also engage advisors and consultants who are concurrently employed at universities or who perform services for other entities.

Although we are not aware of any claims currently pending against us, we may be subject to claims that we or our employees, advisors or consultants have inadvertently or otherwise used or disclosed intellectual property, including trade secrets or other proprietary information, of a former employer or other third party. We have and may in the future also be subject to claims that an employee, advisor or consultant performed work for us that conflicts with that person's obligations to a third party, such as an employer, and thus, that the third party has an ownership interest in the intellectual property arising out of work performed for us. Litigation may be necessary to defend against these claims. Even if we are successful in defending against these claims, litigation could result in substantial costs and be a distraction to management. If we fail in defending such claims, in addition to paying monetary claims, we may lose valuable intellectual property rights or personnel. A loss of key personnel or their work product could hamper or prevent our ability to commercialize our product candidates, which would materially adversely affect our commercial development efforts.

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Numerous factors may limit any potential competitive advantage provided by our intellectual property rights.

The degree of future protection afforded by our intellectual property rights is uncertain because intellectual property rights have limitations, and may not adequately protect our business, provide a barrier to entry against our competitors or potential competitors, or permit us to maintain our competitive advantage. Moreover, if a third party has intellectual property rights that cover the practice of our technology, we may not be able to fully exercise or extract value from our intellectual property rights. The following examples are illustrative:

others may be able to develop and/or practice technology that is similar to our technology or aspects of our technology but that is not covered by the claims of patents, should such patents issue from our patent applications;

we might not have been the first to make the inventions covered by a pending patent application that we own;

we might not have been the first to file patent applications covering an invention;

others may independently develop similar or alternative technologies without infringing our intellectual property rights;

pending patent applications that we own or license may not lead to issued patents;

patents, if issued, that we own or license may not provide us with any competitive advantages, or may be held invalid or unenforceable, as a result of legal challenges by our competitors;

third parties may compete with us in jurisdictions where we do not pursue and obtain patent protection;

we may not be able to obtain and/or maintain necessary or useful licenses on reasonable terms or at all;

third parties may assert an ownership interest in our intellectual property and, if successful, such disputes may preclude us from exercising exclusive rights over that intellectual property;

we may not develop or in-license additional proprietary technologies that are patentable; and

the patents of others may have an adverse effect on our business.

Should any of these events occur, they could significantly harm our business and results of operations.

General Company-Related Risks

As our product candidates reach later stage clinical development, we will need to develop and expand our company, and we may encounter difficulties in managing this development and expansion, which could disrupt our operations.

As of February 28, 2015, we had 30 full-time employees and no part-time employees, and as our product candidates reach later stage clinical development, we expect to increase our number of employees and the scope of our operations. To manage our anticipated development and

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expansion, we must continue to implement and improve our managerial, operational and financial systems, expand our facilities and continue to recruit and train additional qualified personnel. Also, our management may need to divert a disproportionate amount of its attention away from its day-to-day activities and devote a substantial amount of time to managing these development activities. Due to our limited resources, we may not be able to effectively manage the expansion of our operations or recruit and train additional qualified personnel. This may result in weaknesses in our infrastructure;

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give rise to operational mistakes, loss of business opportunities, loss of employees and reduced productivity among remaining employees. The physical expansion of our operations may lead to significant costs and may divert financial resources from other projects, such as the development of our product candidates. If our management is unable to effectively manage our expected development and expansion, our expenses may increase more than expected, our ability to generate or increase our revenue could be reduced and we may not be able to implement our business strategy. Our future financial performance and our ability to commercialize our product candidates, if approved, and compete effectively will depend, in part, on our ability to effectively manage the future development and expansion of our company.

Our future success depends on our ability to retain our President and Chief Executive Officer and to attract, retain and motivate qualified personnel.

We are highly dependent on Dr. Jeffrey M. Jonas, our President and Chief Executive Officer. We have entered into an employment agreement with Dr. Jonas, but he may terminate his employment with us at any time. Although we do not have any reason to believe that we will lose the services of Dr. Jonas in the foreseeable future, the loss of his services might impede the achievement of our research, development and commercialization objectives. We also do not have any key-man life insurance on Dr. Jonas. We rely on consultants and advisors, including scientific and clinical advisors, to assist us in formulating our development and commercialization strategy. Our consultants and advisors may be employed by employers other than us and may have commitments under consulting or advisory contracts with other entities that may limit their availability to us and may not be subject to our standard non-compete agreements. Recruiting and retaining qualified scientific personnel and sales and marketing personnel will also be critical to our success. We may not be able to attract and retain these personnel on acceptable terms given the competition among numerous pharmaceutical and biotechnology companies for similar personnel. We also experience competition for the hiring of scientific personnel from universities and research institutions. Failure to succeed in clinical trials may make it more challenging to recruit and retain qualified scientific personnel.

Our employees may engage in misconduct or other improper activities, including violating applicable regulatory standards and requirements or engaging in insider trading, which could significantly harm our business.

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees could include intentional failures to comply with the regulations of the FDA and applicable non-U.S. regulators, provide accurate information to the FDA and applicable non-U.S. regulators, comply with healthcare fraud and abuse laws and regulations in the United States and abroad, report financial information or data accurately or disclose unauthorized activities to us. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, misconduct, kickbacks, self-dealing and other abusive practices. These laws and regulations restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Employee misconduct could also involve the improper use of, including trading on, information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. We have adopted a code of conduct, but it is not always possible to identify and deter employee misconduct, and the precautions we take to detect and prevent this activity may be ineffective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to comply with these laws or regulations. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could have a significant impact on our business, including the imposition of significant fines or other sanctions.

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We face potential product liability exposure, and, if claims are brought against us, we may incur substantial liability.

The use of our product candidates in clinical trials and the sale of our product candidates, if approved, exposes us to the risk of product liability claims. Product liability claims might be brought against us by patients, healthcare providers or others selling or otherwise coming into contact with our product candidates. For example, we may be sued if any product we develop allegedly causes injury or is found to be otherwise unsuitable during product testing, manufacturing, marketing or sale. Any such product liability claims may include allegations of defects in manufacturing, defects in design, a failure to warn of dangers inherent in the product, including as a result of interactions with alcohol or other drugs, negligence, strict liability and a breach of warranties. Claims could also be asserted under state consumer protection acts. If we become subject to product liability claims and cannot successfully defend ourselves against them, we could incur substantial liabilities. In addition, regardless of merit or eventual outcome, product liability claims may result in, among other things:

withdrawal of patients from our clinical trials;

substantial monetary awards to patients or other claimants;

decreased demand for our product candidates or any future product candidates following marketing approval, if obtained;

damage to our reputation and exposure to adverse publicity;

increased FDA warnings on product labels;

litigation costs;

distraction of management's attention from our primary business;

loss of revenue; and

the inability to successfully commercialize our product candidates or any future product candidates, if approved.

We maintain product liability insurance coverage for our clinical trials with a \$10 million annual aggregate coverage limit. Nevertheless, our insurance coverage may be insufficient to reimburse us for any expenses or losses we may suffer. Moreover, in the future, we may not be able to maintain insurance coverage at a reasonable cost or in sufficient amounts to protect us against losses, including if insurance coverage becomes increasingly expensive. If and when we obtain marketing approval for our product candidates, we intend to expand our insurance coverage to include the sale of commercial products; however, we may not be able to obtain this product liability insurance on commercially reasonable terms. Large judgments have been awarded in class action lawsuits based on drugs that had unanticipated side effects. The cost of any product liability litigation or other proceedings, even if resolved in our favor, could be substantial, particularly in light of the size of our business and financial resources. A product liability claim or series of claims brought against us could cause our stock price to decline and, if we are unsuccessful in defending such a claim or claims and the resulting judgments exceed our insurance coverage, our financial condition, business and prospects could be materially adversely affected.

We will incur increased costs as a result of operating as a public company, and our management team will be required to devote substantial time to new compliance initiatives.

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Now that we are a public company, and particularly after we are no longer considered an emerging growth company, we will incur significant legal, accounting and other expenses that we did not incur as a private company. In addition, the Sarbanes-Oxley Act of 2002 and rules subsequently implemented by the Securities and Exchange Commission and The NASDAQ Stock Market have

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imposed various requirements on public companies, including establishment and maintenance of effective disclosure and financial controls and corporate governance practices. Our management and other personnel will need to devote a substantial amount of time to these compliance initiatives. Moreover, these rules and regulations will increase our legal and financial compliance costs and will make some activities more time-consuming and costly.

Pursuant to Section 404 of the Sarbanes-Oxley Act of 2002, or Section 404, we will be required to furnish a report by our management on our internal control over financial reporting, including an attestation report on internal control over financial reporting issued by our independent registered public accounting firm. However, while we remain an emerging growth company, we will not be required to include an attestation report on internal control over financial reporting issued by our independent registered public accounting firm. To achieve compliance with Section 404 within the prescribed period, we will be engaged in a process to document and evaluate our internal control over financial reporting, which is both costly and challenging. In this regard, we will need to continue to dedicate internal resources, potentially engage outside consultants and adopt a detailed work plan to assess and document the adequacy of internal control over financial reporting, continue steps to improve control processes as appropriate, validate through testing that controls are functioning as documented and implement a continuous reporting and improvement process for internal control over financial reporting. Despite our efforts, there is a risk that neither we nor our independent registered public accounting firm will be able to conclude that our internal control over financial reporting is effective as required by Section 404. This could result in an adverse reaction in the financial markets due to a loss of confidence in the reliability of our consolidated financial statements.

As we continue to grow, we will need to hire additional qualified accounting and financial personnel with appropriate public company experience.

As we continue to grow our organization, we will need to establish and maintain effective disclosure and financial controls and make changes in our corporate governance practices. We will need to hire additional accounting and financial personnel with appropriate public company experience and technical accounting knowledge, and it may be difficult to recruit and maintain such personnel. Even if we are able to hire appropriate personnel, our existing operating expenses and operations will be impacted by the direct costs of their employment and the indirect consequences related to the diversion of management resources from product development efforts.

Our ability to use our net operating loss carryforwards and certain tax credit carryforwards may be subject to limitation.

As of December 31, 2014, we had federal and state net operating loss carryforwards of \$55.8 million and \$55.4 million, respectively, which begin to expire in 2031. As of December 31, 2014, we also had federal and state research and development tax credit carryforwards of \$0.7 million and \$0.3 million, respectively, which begin to expire in 2031 and 2027, respectively. As of December 31, 2014, we had federal orphan drug tax credit carryforwards of \$3.6 million, which begin to expire in 2034. Under Section 382 of the Internal Revenue Code of 1986, as amended, or the Code, changes in our ownership may limit the amount of our net operating loss carryforwards and tax credit carryforwards that could be utilized annually to offset our future taxable income, if any. This limitation would generally apply in the event of a cumulative change in ownership of our company of more than 50% within a three-year period. Any such limitation may significantly reduce our ability to utilize our net operating loss carryforwards and tax credit carryforwards before they expire. The completion of this offering and our initial public offering, or IPO, together with private placements and other transactions that have occurred since our inception, may trigger such an ownership change pursuant to Section 382. Any such limitation, whether as the result of this offering, our IPO, prior private placements, sales of our common stock by our existing stockholders or additional sales of our common stock by us after this

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offering, could have a material adverse effect on our results of operations in future years. We have not completed a study to assess whether an ownership change for purposes of Section 382 has occurred, or whether there have been multiple ownership changes since our inception, due to the significant costs and complexities associated with such study.

Unfavorable global economic conditions could adversely affect our business, financial condition or results of operations.

Our results of operations could be adversely affected by general conditions in the global economy and in the global financial markets. The recent global financial crisis caused extreme volatility and disruptions in the capital and credit markets. A severe or prolonged economic downturn, such as the recent global financial crisis, could result in a variety of risks to our business, including, weakened demand for our product candidates and our ability to raise additional capital when needed on acceptable terms, if at all. A weak or declining economy could also strain our suppliers, possibly resulting in supply disruption, or cause our customers to delay making payments for our services. Any of the foregoing could harm our business and we cannot anticipate all of the ways in which the current economic climate and financial market conditions could adversely impact our business.

We or the third parties upon whom we depend may be adversely affected by natural disasters and our business continuity and disaster recovery plans may not adequately protect us from a serious disaster.

Natural disasters could severely disrupt our operations, and have a material adverse effect on our business, results of operations, financial condition and prospects. If a natural disaster, power outage or other event occurred that prevented us from using all or a significant portion of our headquarters, that damaged critical infrastructure, such as the manufacturing facilities of our third-party CMOs, or that otherwise disrupted operations, it may be difficult or, in certain cases, impossible for us to continue our business for a substantial period of time. The disaster recovery and business continuity plans we have in place may prove inadequate in the event of a serious disaster or similar event. We may incur substantial expenses as a result of the limited nature of our disaster recovery and business continuity plans, which could have a material adverse effect on our business.

Our internal computer systems, or those of our third-party CROs or other contractors or consultants, may fail or suffer security breaches, which could result in a material disruption of our product candidates development programs.

Despite the implementation of security measures, our internal computer systems and those of our third-party CROs and other contractors and consultants are vulnerable to damage from computer viruses, unauthorized access, natural disasters, terrorism, war and telecommunication and electrical failures. While we have not experienced any such system failure, accident, or security breach to date, if such an event were to occur and cause interruptions in our operations, it could result in a material disruption of our programs. For example, the loss of clinical trial data for our product candidates could result in delays in our regulatory approval efforts and significantly increase our costs to recover or reproduce the data. To the extent that any disruption or security breach results in a loss of or damage to our data or applications or other data or applications relating to our technology or product candidates, or inappropriate disclosure of confidential or proprietary information, we could incur liabilities and the further development of our product candidates could be delayed.

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We may acquire businesses or products, or form strategic alliances, in the future, and we may not realize the benefits of such acquisitions.

We may acquire additional businesses or products, form strategic alliances or create joint ventures with third parties that we believe will complement or augment our existing business. If we acquire businesses with promising markets or technologies, we may not be able to realize the benefit of acquiring such businesses if we are unable to successfully integrate them with our existing operations and company culture. We may encounter numerous difficulties in developing, manufacturing and marketing any new products resulting from a strategic alliance or acquisition that delay or prevent us from realizing their expected benefits or enhancing our business. We cannot assure you that, following any such acquisition, we will achieve the expected synergies to justify the transaction.

Risks Related to Our Financial Position and Need for Capital

We are a biopharmaceutical company with a limited operating history and have not generated any revenue from product sales. We have incurred significant operating losses since our inception and anticipate that we will incur continued losses for the foreseeable future.

We are a biopharmaceutical company with a limited operating history on which to base your investment decision. Biopharmaceutical product development is a highly speculative undertaking and involves a substantial degree of risk. We were incorporated in April 2010. Our operations to date have been limited primarily to organizing and staffing our company, raising capital and conducting research and development activities for our product candidates. We have never generated any revenue from product sales. We have not obtained regulatory approvals for any of our product candidates.

We have funded our operations to date through proceeds from sales of redeemable convertible preferred stock, the issuance of convertible notes and the sale of common stock in our IPO. As of December 31, 2014, our cash and cash equivalents were \$127.8 million. We have incurred net losses in each year since our inception, and we have an accumulated deficit of \$66.8 million as of December 31, 2014. Our net losses were \$36.1 million, \$18.3 million and \$9.6 million for the years ended December 31, 2014, 2013 and 2012, respectively. Substantially all of our operating losses have resulted from costs incurred in connection with our research and development programs and from general and administrative costs associated with our operations. We expect to incur increasing levels of operating losses over the next several years and for the foreseeable future. Our prior losses, combined with expected future losses, have had and will continue to have an adverse effect on our stockholders' deficit and working capital. We expect our research and development expenses to significantly increase in connection with our clinical trials of our product candidates. In addition, if we obtain marketing approval for our product candidates, we will incur significant sales, marketing and outsourced-manufacturing expenses. As a public company, we incur additional costs associated with operating as a public company. As a result, we expect to continue to incur significant and increasing operating losses for the foreseeable future. Because of the numerous risks and uncertainties associated with developing pharmaceutical products, we are unable to predict the extent of any future losses or when we will become profitable, if at all. Even if we do become profitable, we may not be able to sustain or increase our profitability on a quarterly or annual basis.

Our ability to become profitable depends upon our ability to generate revenue. To date, we have not generated any revenue from our lead product candidates, SAGE-547, SAGE-689 and SAGE-217, and we do not know when, or if, we will generate any revenue. We do not expect to generate significant revenue unless and until we obtain marketing approval of, and begin to sell, SAGE-547, SAGE-689 or SAGE-217. Our ability to generate revenue depends on a number of factors, including, but not limited to, our ability to:

initiate and successfully complete clinical trials that meet their clinical endpoints;

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initiate and successfully complete all safety studies required to obtain U.S. and foreign marketing approval for our product candidates;

commercialize our product candidates, if approved, by developing a sales force or entering into collaborations with third parties; and

achieve market acceptance of our product candidates in the medical community and with third-party payors.

Absent our entering into a collaboration or partnership agreement, we expect to incur significant sales and marketing costs as we prepare to commercialize our product candidates. Even if we initiate and successfully complete pivotal clinical trials of our product candidates, and our product candidates are approved for commercial sale, and despite expending these costs, our product candidates may not be a commercially successful drug. We may not achieve profitability soon after generating product sales, if ever. If we are unable to generate product revenue, we will not become profitable and may be unable to continue operations without continued funding.

Failure to obtain capital when needed may force us to delay, limit or terminate our product development efforts or other operations.

We are currently advancing our product candidates through non-clinical and clinical development. Developing small molecule products is expensive, and we expect our research and development expenses to increase substantially in connection with our ongoing activities, particularly as we advance our product candidate in clinical trials. Depending on the status of regulatory approval or, if approved, commercialization of our product candidates, as well as the progress we make in selling our product candidates, we may require additional capital to fund operating needs thereafter. We may also need to raise additional funds sooner if we choose to pursue additional indications and/or geographies for our product candidates or otherwise expand more rapidly than we presently anticipate.

As of December 31, 2014, our cash and cash equivalents were \$127.8 million. We estimate that the net proceeds from this offering will be approximately \$ million, assuming a public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, after deducting estimated underwriting discounts and commissions and offering expenses payable by us. We expect that the net proceeds from this offering and our existing cash and cash equivalents will be sufficient to fund our current operations for at least the next 12 months. However, our operating plan may change as a result of many factors currently unknown to us, and we may need to seek additional funds sooner than planned, through public or private equity or debt financings, government or other third-party funding, marketing and distribution arrangements and other collaborations, strategic alliances and licensing arrangements or a combination of these approaches. In any event, we will require additional capital to obtain regulatory approval for, and to commercialize, our product candidates. Raising funds in the current economic environment may present additional challenges. Even if we believe we have sufficient funds for our current or future operating plans, we may seek additional capital if market conditions are favorable or if we have specific strategic considerations.

Any additional fundraising efforts may divert our management from their day-to-day activities, which may adversely affect our ability to develop and commercialize our product candidates. In addition, we cannot guarantee that future financing will be available in sufficient amounts or on terms acceptable to us, if at all. Moreover, the terms of any financing may adversely affect the holdings or the rights of our stockholders and the issuance of additional securities, whether equity or debt, by us, or the possibility of such issuance, may cause the market price of our shares to decline. The sale of additional equity or convertible securities would dilute all of our stockholders. The incurrence of indebtedness would result in increased fixed payment obligations and we may be required to agree to

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certain restrictive covenants, such as limitations on our ability to incur additional debt, limitations on our ability to acquire, sell or license intellectual property rights and other operating restrictions that could adversely impact our ability to conduct our business. We could also be required to seek funds through arrangements with collaborative partners or otherwise at an earlier stage than otherwise would be desirable and we may be required to relinquish rights to some of our technologies or product candidate or otherwise agree to terms unfavorable to us, any of which may have a material adverse effect on our business, operating results and prospects.

If we are unable to obtain funding on a timely basis, we may be required to significantly curtail, delay or discontinue one or more of our research or development programs or the commercialization of any product candidate or be unable to expand our operations or otherwise capitalize on our business opportunities, as desired, which could materially affect our business, financial condition and results of operations.

Raising additional capital may cause dilution to our existing stockholders, restrict our operations or require us to relinquish rights.

We may seek additional capital through a combination of private and public equity offerings, debt financings, collaborations and strategic and licensing arrangements. To the extent that we raise additional capital through the sale of common stock or securities convertible or exchangeable into common stock, your ownership interest in our company will be diluted. In addition, the terms of any such securities may include liquidation or other preferences that materially adversely affect your rights as a stockholder. Debt financing, if available, would increase our fixed payment obligations and may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends. If we raise additional funds through collaboration, strategic partnerships and licensing arrangements with third parties, we may have to relinquish valuable rights to our product candidates, our intellectual property, future revenue streams or grant licenses on terms that are not favorable to us.

If you purchase our common stock in this offering, you will incur immediate and substantial dilution in the book value of your shares.

You will suffer immediate and substantial dilution in the net tangible book value of the common stock you purchase in this offering. Assuming a public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, purchasers of common stock in this offering will experience immediate dilution of \$ _____ per share in net tangible book value of the common stock. In addition, investors purchasing common stock in this offering will contribute _____ % of the total amount invested by stockholders since inception but will only own _____ % of the shares of common stock outstanding. In the past, we issued securities to acquire common stock at prices significantly below the offering price. To the extent these outstanding securities are ultimately exercised, investors purchasing common stock in this offering will sustain further dilution. See Dilution for a more detailed description of the dilution to new investors in the offering.

Risks Related to Our Common Stock

Market volatility may affect our stock price and the value of your investment.

Our IPO was completed on July 23, 2014. Therefore, there has only been a public market for our common stock for a short period of time.

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The market price for our common stock, similar to other biopharmaceutical companies, is likely to be volatile. The market price of our common stock may fluctuate significantly in response to a number of factors, most of which we cannot control, including, among others:

plans for, progress of or results from non-clinical studies and clinical trials of our product candidates;

the failure of the FDA to approve our product candidates;

announcements of new products, technologies, commercial relationships, acquisitions or other events by us or our competitors;

the success or failure of other CNS therapies;

regulatory or legal developments in the United States and other countries;

failure of our product candidates, if approved, to achieve commercial success;

fluctuations in stock market prices and trading volumes of similar companies;

general market conditions and overall fluctuations in U.S. equity markets;

variations in our quarterly operating results;

changes in our financial guidance or securities analysts' estimates of our financial performance;

changes in accounting principles;

our ability to raise additional capital and the terms on which we can raise it;

sales of large blocks of our common stock, including sales by our executive officers, directors and significant stockholders;

additions or departures of key personnel;

discussion of us or our stock price by the press and by online investor communities; and

other risks and uncertainties described in these risk factors.

We have a significant stockholder, which will limit your ability to influence corporate matters and may give rise to conflicts of interest.

A fund affiliated with Third Rock Ventures, or TRV, is our largest stockholder. As of February 28, 2015, TRV beneficially owned approximately 37.9% of our common stock. Accordingly, TRV exerts and will continue to exert significant influence over us and any action requiring the approval of the holders of our common stock, including the election of directors and amendments to our organizational documents, such as increases in our authorized shares of common stock and approval of significant corporate transactions. Furthermore, the interests of TRV may not always coincide with your interests or the interests of other stockholders and TRV may act in a manner that advances its best interests and not necessarily those of other stockholders, including seeking a premium value for its common stock, which might affect the prevailing market price for our common stock.

Our executive officers, directors, principal stockholders and their affiliates will continue to exercise significant control over our company after this offering, which will limit your ability to influence corporate matters and could delay or prevent a change in corporate control.

Immediately following the completion of this offering, and disregarding any shares of common stock that they purchase in this offering, the existing holdings of our executive officers, directors, principal stockholders and their affiliates, including investment funds affiliated with ARCH Venture

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Fund VII, L.P., or ARCH, TRV, and entities affiliated with Fidelity Investment, or Fidelity, will represent beneficial ownership, in the aggregate, of approximately % of our outstanding common stock, assuming no exercise of the underwriters' option to acquire additional common stock in this offering and assuming we issue the number of shares of common stock as set forth on the cover page of this prospectus. As a result, these stockholders, if they act together, will be able to influence our management and affairs and control the outcome of matters submitted to our stockholders for approval, including the election of directors and any sale, merger, consolidation, or sale of all or substantially all of our assets. These stockholders acquired their shares of common stock for substantially less than the price of the shares of common stock being acquired in this offering, and these stockholders may have interests, with respect to their common stock, that are different from those of investors in this offering and the concentration of voting power among these stockholders may have an adverse effect on the price of our common stock. In addition, this concentration of ownership might adversely affect the market price of our common stock by:

delaying, deferring or preventing a change of control of us;

impeding a merger, consolidation, takeover or other business combination involving us; or

discouraging a potential acquirer from making a tender offer or otherwise attempting to obtain control of us.

See Principal Stockholders in this prospectus for more information regarding the ownership of our outstanding common stock by our executive officers, directors, principal stockholders and their affiliates.

Sales of a substantial number of shares of our common stock in the public market could cause our stock price to fall.

If our existing stockholders sell, or indicate an intention to sell, substantial amounts of our common stock in the public market after the lock-up and other legal restrictions on resale discussed in this prospectus lapse, the market price of our common stock could decline. Based upon the number of shares of common stock, on an as-converted basis, outstanding as of February 28, 2015, upon the completion of this offering, we will have outstanding a total of shares of common stock, assuming no exercise of the underwriters' option to purchase additional shares. Of these shares, as of the date of this prospectus, approximately shares of our common stock, plus any shares sold upon exercise of the underwriters' option to purchase additional shares, will be freely tradable, without restriction, in the public market immediately following this offering, assuming that current stockholders do not purchase shares in this offering. The representatives of the underwriters, however, may, in their sole discretion, permit our officers, directors and other stockholders who are subject to these lock-up agreements to sell shares prior to the expiration of the lock-up agreements.

The lock-up agreements pertaining to this offering will expire 60 days from the date of this prospectus. After the lock-up agreements expire, based upon the number of shares of common stock, on an as-converted basis, outstanding as of February 28, 2015, up to an additional 14,157,400 shares of common stock will be eligible for sale in the public market, 14,157,400 of which shares are held by directors, executive officers and other affiliates and will be subject to Rule 144 under the Securities Act of 1933, as amended, or the Securities Act.

In addition, as of February 28, 2015, 4,545,582 shares of common stock that are either subject to outstanding options or reserved for future issuance under our equity incentive plans will become eligible for sale in the public market to the extent permitted by the provisions of various vesting schedules, the lock-up agreements and Rule 144 and Rule 701 under the Securities Act. If these additional shares of common stock are sold, or if it is perceived that they will be sold, in the public market, the market price of our common stock could decline.

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After this offering, the holders of approximately 14,084,664 shares of our common stock will be entitled to rights with respect to the registration of their shares under the Securities Act, subject to the lock-up agreements described above. Registration of these shares under the Securities Act would result in the shares becoming freely tradable without restriction under the Securities Act, except for shares purchased by affiliates. Any sales of securities by these stockholders could have a material adverse effect on the market our common stock.

We have broad discretion in how we use the proceeds of this offering and may not use these proceeds effectively, which could affect our results of operations and cause our stock price to decline.

We will have considerable discretion in the application of the net proceeds of this offering. We intend to use the net proceeds from this offering to fund the planned Phase 3 development of SAGE-547 and costs associated with initial NDA preparatory work, to fund Phase 1 development activities for SAGE-217 and the remaining proceeds to fund new and ongoing research and development activities and early planning and pre-launch investments in commercial infrastructure for working capital and other general corporate purposes, which may include funding for the hiring of additional personnel, capital expenditures and for working capital and other general corporate purposes. As a result, investors will be relying upon management's judgment with only limited information about our specific intentions for the use of the balance of the net proceeds of this offering. We may use the net proceeds for purposes that do not yield a significant return or any return at all for our stockholders. In addition, pending their use, we may invest the net proceeds from this offering in a manner that does not produce income or that loses value.

After the completion of this offering, we may be at an increased risk of securities class action litigation.

Historically, securities class action litigation has often been brought against a company following a decline in the market price of its securities. This risk is especially relevant for us because biotechnology and pharmaceutical companies have experienced significant stock price volatility in recent years. If we were to be sued, it could result in substantial costs and a diversion of management's attention and resources, which could harm our business.

Anti-takeover provisions in our charter documents and under Delaware law could make an acquisition of us, even one that may be beneficial to our stockholders, more difficult and may prevent attempts by our stockholders to replace or remove our current management.

Provisions in our amended and restated certificate of incorporation and amended and restated bylaws may delay or prevent an acquisition of us or a change in our management. These provisions include a classified board of directors, a prohibition on actions by written consent of our stockholders and the ability of our board of directors to issue preferred stock without stockholder approval. In addition, because we are incorporated in Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporation Law, which limits the ability of stockholders owning in excess of 15% of our outstanding voting stock to merge or combine with us. Although we believe these provisions collectively provide for an opportunity to obtain greater value for stockholders by requiring potential acquirers to negotiate with our board of directors, they would apply even if an offer rejected by our board were considered beneficial by some stockholders. In addition, these provisions may frustrate or prevent any attempts by our stockholders to replace or remove our current management by making it more difficult for stockholders to replace members of our board of directors, which is responsible for appointing the members of our management.

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We are an emerging growth company, and as a result of the reduced disclosure and governance requirements applicable to emerging growth companies, our common stock may be less attractive to investors.

We are an emerging growth company, as defined in the JOBS Act, and we have taken advantage of certain exemptions from various reporting requirements that are applicable to other public companies that are not emerging growth companies including, but not limited to, not being required to comply with the auditor attestation requirements of Section 404, reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements, and exemptions from the requirements of holding a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved. Since we have chosen not to comply with the auditor attestation requirements of Section 404 of the Sarbanes-Oxley Act, our auditors are not required to attest to the effectiveness of our internal control over financial reporting. As a result, investors may become less comfortable with the effectiveness of our internal controls and the risk that material weaknesses or other deficiencies in our internal controls go undetected may increase. Since we have chosen to provide reduced disclosures in our periodic reports and proxy statements while we are an emerging growth company, investors have access to less information and analysis about our executive compensation, which may make it difficult for investors to evaluate our executive compensation practices. We cannot predict if investors will find our common stock less attractive because we have chosen to rely on these exemptions. If some investors find our common stock less attractive as a result, there may be a less active trading market for our common stock and our stock price may be more volatile. We may take advantage of these reporting exemptions until we are no longer an emerging growth company. We will remain an emerging growth company until the earlier of (a) the last day of the fiscal year following the fifth anniversary of the completion of our IPO, (b) the last day of the fiscal year in which we have total annual gross revenue of at least \$1.0 billion, (c) the date on which we are deemed to be a large accelerated filer, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30th, and (d) the date on which we have issued more than \$1.0 billion in non-convertible debt during the prior three-year period.

We do not intend to pay dividends on our common stock and, consequently, your ability to achieve a return on your investment will depend on appreciation in the price of our common stock.

We have never declared or paid any cash dividend on our common stock and do not currently intend to do so in the foreseeable future. We currently anticipate that we will retain future earnings for the development, operation and expansion of our business and do not anticipate declaring or paying any cash dividends in the foreseeable future. Therefore, the success of an investment in shares of our common stock will depend upon any future appreciation in their value. There is no guarantee that shares of our common stock will appreciate in value or even maintain the price at which you purchased them.

If equity research analysts do not publish research or reports about our business or if they issue unfavorable commentary or downgrade our common stock, the price of our common stock could decline.

The trading market for our common stock relies in part on the research and reports that equity research analysts publish about us and our business. We do not control these analysts. The price of our common stock could decline if one or more equity research analysts downgrade our common stock or if analysts issue other unfavorable commentary or cease publishing reports about us or our business.

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CAUTIONARY NOTE REGARDING FORWARD-LOOKING STATEMENTS

This prospectus, including the sections entitled Prospectus Summary, Risk Factors, Management's Discussion and Analysis of Financial Condition and Results of Operations and Business, contains forward-looking statements that are based on our management's belief and assumptions and on information currently available to our management. Although we believe that the expectations reflected in these forward-looking statements are reasonable, these statements relate to our strategy, future operations, future financial position, future revenue, projected costs, prospects, plans, objectives of management and expected market growth, and involve known and unknown risks, uncertainties and other factors that may cause our actual results, levels of activity, performance or achievements to be materially different from any future results, levels of activity, performance or achievements expressed or implied by these forward-looking statements. Forward-looking statements in this prospectus include, but are not limited to, statements about:

our use of the net proceeds from this offering;

the accuracy of our estimates regarding expenses, future revenues and capital requirements;

our ability to achieve orphan drug designation for our product candidates, or to take advantage of receipt of such designation;

our plans to develop and commercialize our product candidates, initially as treatments for SE, RSE, SRSE and orphan epilepsies such as Dravet syndrome and Rett syndrome;

our ability to advance our product candidates into and through clinical trials, including pivotal clinical trials, and successfully complete such clinical trials;

regulatory developments in the United States and foreign countries;

the need to obtain additional regulatory approvals following receipt of marketing authorization, such as from the DEA;

the performance of our third-party manufacturers and CROs;

our ability to obtain and maintain intellectual property protection for our proprietary assets;

the size of the potential markets for our product candidates and our ability to serve those markets;

the rate and degree of market acceptance of our product candidates for any indication once approved;

our ability to obtain additional financing when needed;

the success of competing products that are or become available for the indications that we are pursuing; and

the loss of key scientific or management personnel.

In some cases, you can identify forward-looking statements by terminology such as may, will, should, expects, intends, plans, anticipate, believes, estimates, predicts, potential, continue or the negative of these terms or other comparable terminology. These statements are only predictions. You should not place undue reliance on forward-looking statements because they involve known and unknown risks, uncertainties and other factors, which are, in some cases, beyond our control and which could materially affect results. Factors that may cause actual results to differ materially from current expectations include, among other things, those listed under Risk Factors and elsewhere in this prospectus. If one or more of these risks or uncertainties occur, or if our underlying assumptions prove to be incorrect, actual events or results may vary significantly from those implied or projected by the forward-looking statements. No forward-looking statement is a guarantee of future

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performance. You should read this prospectus and the documents that we reference in this prospectus and have filed with the SEC as exhibits to the registration statement, of which this prospectus is a part, completely and with the understanding that our actual future results may be materially different from any future results expressed or implied by these forward-looking statements.

The forward-looking statements in this prospectus represent our views as of the date of this prospectus. We anticipate that subsequent events and developments will cause our views to change. However, while we may elect to update these forward-looking statements at some point in the future, we have no current intention of doing so except to the extent required by applicable law. You should, therefore, not rely on these forward-looking statements as representing our views as of any date subsequent to the date of this prospectus.

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USE OF PROCEEDS

We estimate that our net proceeds from the sale of shares of our common stock in this offering will be approximately \$ million, or \$ million if the underwriters exercise in full their option to purchase additional shares, assuming a public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

A \$1.00 increase (decrease) in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase (decrease) the net proceeds to us from this offering by \$ million, assuming the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. An increase (decrease) of shares in the number of shares offered by us, as set forth on the cover page of this prospectus, would increase (decrease) our net proceeds from this offering by \$ million, assuming no change in the assumed public offering price per share and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. A share increase in the number of shares offered by us together with a concomitant \$1.00 increase in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase the net proceeds to us from this offering by \$ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. Conversely, a share decrease in the number of shares offered by us together with a concomitant \$1.00 decrease in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would decrease the net proceeds to us from this offering by \$ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

We intend to use the net proceeds from this offering, plus, if needed, cash on hand, as follows:

approximately \$60.0 to 65.0 million to fund the Phase 3 development of SAGE-547 for SRSE, including costs related to the preparation of a potential NDA submission pending a successful outcome of our planned Phase 3 clinical trial;

approximately \$10.0 million to fund Phase 1 development activities for SAGE-217; and

the remaining proceeds to fund new and ongoing research and development activities, early planning and pre-launch investments in commercial infrastructure, working capital and other general corporate purposes, which may include funding for the hiring of additional personnel, capital expenditures and for working capital and other general corporate purposes.

Based on our current plans, we believe our cash, cash equivalents and short-term investments, together with the net proceeds to us from this offering, will be sufficient to fund our operations for at least the next 12 months.

This expected use of the net proceeds from this offering represents our intentions based upon our current plans and business conditions. As of the date of this prospectus, we cannot predict with certainty all of the particular uses for the net proceeds to be received upon the completion of this offering or the amounts that we will actually spend on the uses set forth above. The amounts and timing of our actual expenditures and the extent of clinical development may vary significantly depending on numerous factors, including the progress of our development efforts, the status of and results from non-clinical studies and ongoing clinical trials or any clinical trials we may commence in the future, as well as any collaborations that we may enter into with third parties for our product

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candidates and any unforeseen cash needs. As a result, our management will retain broad discretion over the allocation of the net proceeds from this offering.

Pending our use of the net proceeds from this offering, we intend to invest the net proceeds in a variety of capital preservation investments, including short-term, investment-grade, interest-bearing instruments and U.S. government securities.

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Our common stock began trading on The NASDAQ Global Market under the symbol SAGE on July 18, 2014. Prior to that time, there was no public market for our common stock. The following table sets forth the high and low sale prices per share of our common stock, as reported on The NASDAQ Global Market, for the periods indicated.

	High	Low
Year Ended December 31, 2014		
Third quarter (from July 18, 2014)	\$ 33.40	\$ 25.86
Fourth quarter	\$ 43.75	\$ 30.50
Year Ending December 31, 2015		
First quarter	\$ 53.38	\$ 36.99
Second quarter (through April 2, 2015)	\$ 50.86	\$ 46.87

On April 2, 2015, the last reported sale price of our common stock as reported on The NASDAQ Global Market was \$46.87 per share. As of February 28, 2015, we had approximately 38 holders of record of our common stock. The actual number of stockholders is greater than this number of record holders and includes stockholders who are beneficial owners but whose shares are held in street name by brokers and other nominees. This number of holders of record also does not include stockholders whose shares may be held in trust by other entities.

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DIVIDEND POLICY

We have never declared or paid dividends on our capital stock. We do not anticipate paying any dividends on our capital stock in the foreseeable future. We currently intend to retain all available funds and any future earnings to fund the development and growth of our business. Any future determination to declare dividends will be subject to the discretion of our board of directors and will depend on various factors, including applicable laws, our results of operations, financial condition, future prospects and any other factors deemed relevant by our board of directors. Investors should not purchase our common stock with the expectation of receiving cash dividends.

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The following table sets forth our cash and cash equivalents and our capitalization as of December 31, 2014:

on an actual basis;

on a pro forma basis, to give effect to the issuance and sale by us of _____ shares of common stock in this offering at the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us

The pro forma information below is illustrative only, and our capitalization following the completion of this offering will be adjusted based on the actual public offering price and other terms of this offering determined at pricing. You should read this table in conjunction with the sections of this prospectus entitled Selected Consolidated Financial Data and Management's Discussion and Analysis of Financial Condition and Results of Operations and with our consolidated financial statements and related notes included elsewhere in this prospectus.

	As of December 31, 2014,	
	Actual	Pro Forma
Cash and cash equivalents	\$ 127,766	\$
Stockholders' equity:		
Preferred stock, \$0.0001 par value; 5,000,000 shares authorized, no shares issued and outstanding, actual and pro forma		
Common stock, \$0.0001 par value; 120,000,000 shares authorized, 25,792,623 shares issued and outstanding, actual; 120,000,000 shares authorized, _____ shares issued and outstanding, pro forma		3
Additional paid-in capital	188,727	
Accumulated deficit	(66,845)	
Total stockholders' equity	121,885	
Total capitalization	\$ 121,885	\$

A \$1.00 increase (decrease) in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase (decrease) each of cash and cash equivalents, total stockholders' equity and total capitalization on a pro forma as adjusted basis by \$ _____ million, assuming the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. An increase (decrease) of _____ shares in the number of shares offered by us, as set forth on the cover page of this prospectus, would increase (decrease) our net proceeds from this offering by \$ _____ million, assuming the assumed public offering price remains the same and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. A _____ share increase in the number of shares offered by us together with a concomitant \$1.00 increase in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase each of cash and cash equivalents, total stockholders' equity and total capitalization by \$ _____ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. Conversely, a _____ share decrease in the number of shares offered by us together with

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a concomitant \$1.00 decrease in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would decrease each of cash and cash equivalents, total stockholders' equity and total capitalization by \$ _____ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

The number of common shares shown as outstanding on an actual and on a pro forma basis in the table above is based on 25,792,623 shares of common stock outstanding as of December 31, 2014, which includes 170,832 shares that are subject to repurchase by us and are not considered outstanding for accounting purposes until vested, and excludes:

1,996,615 shares of common stock issuable upon the exercise of outstanding options as of December 31, 2014 at a weighted average exercise price of \$7.01 per share;

1,509,253 shares of our common stock reserved for future issuance under our 2014 Stock Option Plan as of December 31, 2014; and

282,000 shares of common stock reserved for future issuance under our 2014 Employee Stock Purchase Plan as of December 31, 2014.

Table of Contents**DILUTION**

If you invest in our common stock in this offering, your ownership interest will be diluted immediately to the extent of the difference between the public offering price per share you will pay in this offering and the pro forma as adjusted net tangible book value per share of our common stock immediately after this offering.

Our historical net tangible book value (deficit) as of December 31, 2014 was \$121.9 million, or \$4.73 per share of common stock. Our historical net tangible book value (deficit) is the amount of our total tangible assets less our total liabilities. Historical net tangible book value per share is our historical net tangible book value (deficit) divided by the number of shares of common stock outstanding as of December 31, 2014.

Our pro forma net tangible book value as of December 31, 2014 was \$ million, or \$ per share of common stock. Pro forma net tangible book value represents total tangible assets less total liabilities. Pro forma net tangible book value per share represents pro forma net tangible book value divided by the pro forma number of shares of our common stock outstanding as of December 31, 2014.

After giving effect to the sale by us of shares of common stock in this offering at an assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us, our pro forma as adjusted net tangible book value as of December 31, 2014 would have been \$ million, or \$ per share. This represents an immediate increase in pro forma as adjusted net tangible book value of \$ per share to existing stockholders and an immediate dilution of \$ per share to new investors purchasing common stock in this offering at the public offering price. Dilution per share to new investors is determined by subtracting pro forma as adjusted net tangible book value per share after this offering from the assumed public offering price per share paid by new investors. The following table illustrates this dilution on a per share basis:

Assumed public offering price per share	\$
Historical net tangible book value (deficit) per share as of December 31, 2014	\$ 4.73
Pro forma net tangible book value per share as of December 31, 2014	
Increase in pro forma as adjusted net tangible book value per share attributable to new investors purchasing shares in this offering	
Pro forma as adjusted net tangible book value per share after this offering	
Dilution per share to new investors participating in this offering	\$

A \$1.00 increase (decrease) in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase (decrease) the pro forma as adjusted net tangible book value per share after this offering by \$ per share and would increase (decrease) the dilution per share to new investors participating in this offering by \$ per share, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. An increase of shares in the number of shares offered by us, as set forth on the cover page of this prospectus, would increase the pro forma as adjusted net tangible book value per share after this offering by \$ and decrease the dilution per share to new investors participating in this offering by \$, assuming the assumed public offering price remains the same and after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. A decrease

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of _____ shares in the number of shares offered by us, as set forth on the cover page of this prospectus, would decrease the pro forma as adjusted net tangible book value per share after this offering by \$ _____ and increase the dilution per share to new investors participating in this offering by \$ _____, assuming the assumed public offering price remains the same and after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us. A _____ share increase in the number of shares offered by us together with a concomitant \$1.00 increase in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would increase the pro forma as adjusted net tangible book value per share after this offering by \$ _____ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. Conversely, a _____ share decrease in the number of shares offered by us together with a concomitant \$1.00 decrease in the assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, would decrease the pro forma as adjusted net tangible book value per share after this offering by \$ _____ million after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. The information discussed above is illustrative only and will be adjusted based on the actual public offering price and other terms of this offering determined at pricing.

If the underwriters exercise their option to purchase additional shares in full, the pro forma as adjusted net tangible book value per share after this offering would be \$ _____ per share, representing an increase in pro forma as adjusted net tangible book value per share to existing stockholders of \$ _____ per share and immediate dilution in pro forma as adjusted net tangible book value per share to new investors purchasing shares in this offering of \$ _____ per share, assuming an public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015.

The following table summarizes, as of December 31, 2014, on a pro forma as adjusted basis described above, the number of shares of our common stock, the total consideration and the average price per share (i) paid to us by existing stockholders and (ii) to be paid by new investors purchasing shares of common stock in this offering at an assumed public offering price of \$46.87 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on April 2, 2015, before deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

	Shares Purchased		Total Consideration		Average Price Per Share
	Number	Percent	Amount	Percent	Share
Existing stockholders		%	\$	%	\$
New investors					\$
Total		%	\$	%	

The table above assumes no exercise of the underwriters' option to purchase additional shares in this offering. If the underwriters' option to purchase additional shares is exercised in full, the number of shares of our common stock held by existing stockholders would be reduced to _____ % of the total number of shares of our common stock outstanding after this offering, and the number of shares of common stock held by new investors participating in the offering would be increased to _____ % of the total number of shares of our common stock outstanding after this offering.

The above discussion and tables are based on 25,792,623 shares of common stock outstanding as of December 31, 2014 (which includes 170,832 shares that are subject to repurchase by us and are not considered outstanding for accounting purposes until vested). The discussion and tables above

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assume no exercise of any outstanding stock options. As of December 31, 2014, there were 1,996,615 shares of common stock issuable upon exercise of outstanding options at a weighted average exercise price of \$7.01 per share. The tables above also exclude 1,509,253 shares of common stock reserved for future issuance under our 2014 Stock Option Plan as of December 31, 2014, and 282,000 shares of common stock reserved for future issuance under our 2014 Employee Stock Purchase Plan as of December 31, 2014.

To the extent that outstanding options are exercised, you will experience further dilution. In addition, we may choose to raise additional capital due to market conditions or strategic considerations even if we believe we have sufficient funds for our current or future operating plans. To the extent that additional capital is raised through the sale of equity or convertible debt securities, the issuance of these securities may result in further dilution to our stockholders.

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You should read the following selected consolidated financial data together with our consolidated financial statements and the related notes appearing at the end of this prospectus and the Management's Discussion and Analysis of Financial Condition and Results of Operations section of this prospectus. We have derived the consolidated statements of operations data for the years ended December 31, 2014, 2013, and 2012 and the consolidated balance sheet data as of December 31, 2014 and 2013 from our audited consolidated financial statements included elsewhere in this prospectus. Our historical results are not necessarily indicative of the results that should be expected in the future.

	Years Ended December 31,		
	2014	2013	2012
	(in thousands, except for per share amounts)		
Consolidated statements of operations data:			
Operating expenses:			
Research and development	\$ 24,100	\$ 14,357	\$ 7,229
General and administrative	9,710	3,922	2,402
Total operating expenses	33,810	18,279	9,631
Loss from operations	(33,810)	(18,279)	(9,631)
Interest income (expense), net	8	1	
Other income (expense), net	(9)	(3)	(1)
Net loss and comprehensive loss	(33,811)	(18,281)	(9,632)
Accretion of redeemable convertible preferred stock to redemption value	(2,294)	(7)	(4)
Net loss attributable to common stockholders	\$ (36,105)	\$ (18,288)	\$ (9,636)
Net loss per share attributable to common stockholders - basic and diluted ⁽¹⁾	\$ (1.67)	\$ (12.26)	\$ (8.62)
Weighted average common shares outstanding - basic and diluted ⁽¹⁾	21,574	1,492	1,118

	Years Ended December 31,	
	2014	2013
	(in thousands)	
Consolidated balance sheet data:		
Cash and cash equivalents	\$ 127,766	\$ 8,066
Working capital ⁽²⁾	121,065	6,092
Total assets	129,665	8,532
Redeemable convertible preferred stock		37,709
Common stock and additional paid-in capital	188,730	139
Total stockholders' equity (deficit)	121,885	(31,536)

- (1) See Note 8 to our consolidated financial statements for further details on the calculation of basic and diluted net loss per share attributable to common stockholders.
- (2) We define working capital as current assets less current liabilities.

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MANAGEMENT'S DISCUSSION AND ANALYSIS OF FINANCIAL CONDITION AND RESULTS OF OPERATIONS

You should read the following discussion and analysis of our financial condition and results of operations together with the section entitled "Selected Consolidated Financial Data" and our consolidated financial statements and related notes included elsewhere in this prospectus. This discussion and other parts of this prospectus contain forward-looking statements that involve risks and uncertainties, such as statements regarding our plans, objectives, expectations, intentions and projections. Our actual results could differ materially from those discussed in these forward-looking statements. Factors that could cause or contribute to such differences include, but are not limited to, those discussed in the "Risk Factors" section of this prospectus.

Overview

We are a biopharmaceutical company committed to developing and commercializing novel medicines to treat life-threatening, rare central nervous system, or CNS, disorders, where there are inadequate or no approved existing therapies. We are targeting CNS indications where patient populations are easily identified, acute treatment is typically initiated in the hospital setting, clinical endpoints are well-defined and development pathways are feasible.

Our initial product candidates, which are summarized in the table below, are aimed at treating different stages of status epilepticus, or SE, a life-threatening condition in which the brain is in a state of persistent seizure, as well as other seizure and non-seizure disorders. The lead product candidate in our SE program, SAGE-547, is an intravenous, or IV, agent entering Phase 3 clinical development as an adjunctive therapy, a therapy combined with current therapeutic approaches, for the treatment of super-refractory SE, or SRSE. The current standard of care for SRSE is empiric, and there are no therapies at present that have been specifically approved for this indication. Over the course of 2014, the U.S. Food and Drug Administration, or FDA, granted us orphan drug designation and Fast Track designation for our investigational new drug application for SAGE-547 as a treatment for SRSE. On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE and we expect to initiate the Phase 3 trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of a New Drug Application, or NDA, submission for SAGE-547.

We continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for essential tremor, a debilitating neurological disorder that causes involuntary, rhythmic shaking with no known cause, and severe post-partum depression, a distinct and readily identified form of major depressive disorder estimated to affect up to 20% of women following childbirth. If these exploratory trials are successful, we plan to use the data from them to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

Our next-generation product candidates, SAGE-689 and SAGE-217, utilize similar mechanistic pathways as SAGE-547 and are designed to have pharmaceutical properties which optimize both their non-clinical profiles and potential clinical profiles for the treatment of different stages of SE, as well as other seizure and non-seizure disorders.

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Since our inception in April 2010, we have devoted substantially all of our resources to organizing and staffing our company, business planning, raising capital, identifying and developing our product candidates, preparing to conduct and conducting non-clinical and clinical trials of our product candidates, providing general and administrative support for these operations and protecting our intellectual property. We have funded our operations to date through sales of our common stock and redeemable convertible preferred stock; the issuance of convertible notes and through proceeds from our initial public offering, or IPO.

We have not generated any revenue to date. We have incurred net losses in each year since our inception, and we have an accumulated deficit of \$66.8 million as of December 31, 2014. Our net losses were \$36.1 million, \$18.3 million, and \$9.6 million for the years ended December 31, 2014, 2013, and 2012, respectively. These losses have resulted principally from costs incurred in connection with research and development activities and general and administrative costs associated with our operations. We expect to incur significant expenses and increasing operating losses for the foreseeable future.

We expect that our expenses will increase substantially in connection with our ongoing activities, as we:

advance clinical development of SAGE-547, our lead product candidate in our SE program, including completing our planned Phase 3 clinical trial for SAGE-547 in SRSE, late stage non-clinical studies of SAGE-547 and initial preparations for a potential commercial launch;

advance our clinical trials to establish proof of principle in additional indications including severe PPD and essential tremor;

advance development of SAGE-689 as an adjunctive second-line therapy for the treatment of SE, including conducting a Phase 1 clinical trial;

advance development of SAGE-217 as an oral monotherapy for orphan epilepsies such as Dravet syndrome and Rett syndrome, including conducting a Phase 1 clinical trial;

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continue our research and development efforts for other drug candidates in the treatment of CNS disorders including on our early-stage novel allosteric modulators for NMDA;

seek regulatory approvals for our product candidates;

add personnel, including personnel to support our product development and future commercialization;

add operational, financial and management information systems;

maintain, leverage and expand our intellectual property portfolio; and

operate as a public company.

As a result, we will need additional financing to support our continuing operations. Until such time that we can generate significant revenue from product sales, if ever, we expect to finance our operations through a combination of public or private equity or debt financings or other sources, which may include collaborations with third parties. Arrangements with collaborators or others may require us to relinquish rights to certain of our technologies or product candidates. In addition, we may never successfully complete development of any of our product candidates, obtain adequate patent protection for our technology, obtain necessary regulatory approval for our product candidates or achieve commercial viability for any approved product candidates. Adequate additional financing may not be available to us on acceptable terms, or at all. Our inability to raise capital as and when needed would have a negative impact on our financial condition and our ability to pursue our business strategy. We will need to generate significant revenue to achieve profitability, and we may never do so.

Financial Operations Overview

Revenue

We have not generated any revenue from product sales since our inception and do not expect to generate any revenue from the sale of products in the near future. If our development efforts result in clinical success and regulatory approval or collaboration agreements with third parties for our product candidates, we may generate revenue from those product candidates.

Operating Expenses

Our operating expenses since inception have consisted primarily of research and development activities and general and administrative costs.

Research and Development Expenses

Research and development expenses, which consist primarily of costs associated with our product research and development efforts, are expensed as incurred. Research and development expenses consist primarily of:

personnel costs, including salaries, related benefits, stock-based compensation and related travel expenses for employees engaged in scientific research and development functions;

expenses incurred under agreements with contract research organizations, or CROs, and investigative sites that conduct our non-clinical studies and clinical trials;

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expenses associated with manufacturing clinical study materials and developing external manufacturing capabilities;

costs of outside consultants, including their fees, stock-based compensation and related travel expenses;

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other expenses related to our non-clinical studies and expenses related to our regulatory activities; and

payments made under our third-party licensing agreements.

Costs for certain development activities are recognized based on an evaluation of the progress to completion of specific tasks using information and data provided to us by our vendors and our clinical sites.

We have been developing SAGE-547, SAGE-689 and SAGE-217 and focusing on other research and development programs related to exploratory efforts, target validation and lead optimization for our earlier-validated programs. Our direct research and development expenses are tracked on a program-by-program basis and consist primarily of external costs, such as fees paid to investigators, central laboratories, CROs and contract manufacturing organizations, or CMOs in connection with our non-clinical studies and clinical trials; third-party license fees related to our product candidates; fees paid to outside consultants who perform work on our programs; and costs related to manufacturing or purchasing clinical trial materials. We do not allocate employee-related costs and other indirect costs to specific research and development programs because these costs are deployed across multiple product programs under research and development and, as such, are separately classified as unallocated research and development expenses.

The following table summarizes our research and development expenses by program:

	2014	2013	2012
	(in thousands)		
SAGE-547	\$ 9,137	\$ 3,918	\$ 125
SAGE-689	3,058	2,772	1,047
SAGE-217	2,764	1,129	
Other research and development programs	3,088	3,388	3,495
Unallocated expenses	6,053	3,150	2,562
Total research and development programs	\$ 24,100	\$ 14,357	\$ 7,229

Research and development activities are central to our business. Product candidates in later stages of clinical development generally have higher development costs than those in earlier stages of clinical development, primarily due to the increased size and duration of later-stage clinical trials. We expect that our research and development expenses will continue to increase in the foreseeable future as we initiate clinical trials for certain product candidates and pursue later stages of clinical development of our product candidates.

We cannot determine with certainty the duration and completion costs of the current or future clinical trials of our product candidates or if, when, or to what extent we will generate revenue from the commercialization and sale of any of our product candidates that obtain regulatory approval. We may never succeed in achieving regulatory approval for any of our product candidates. The duration, costs, and timing of clinical trials and development of our product candidates will depend on a variety of factors, including:

the scope, rate of progress, and expense of our ongoing as well as any additional non-clinical studies, clinical trials and other research and development activities;

future clinical trial results;

uncertainties in clinical trial enrollment rate or design;

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significant and changing government regulation; and

the timing and receipt of any regulatory approvals.

A change in the outcome of any of these variables with respect to the development of a product candidate could mean a significant change in the costs and timing associated with the development of that product candidate. For example, if the FDA or another regulatory authority were to require us to conduct clinical trials beyond those that we currently anticipate will be required for the completion of clinical development of a product candidate, or if we experience significant delays in enrollment in any of our clinical trials, we could be required to expend significant additional financial resources and time on the completion of clinical development.

General and Administrative Expenses

General and administrative expenses consist primarily of personnel costs, consisting of salaries, related benefits, stock-based compensation and related travel expenses of our executive, finance, business and corporate development and other administrative functions. General and administrative expenses also include expenses incurred under agreements with third parties relating to initial commercial evaluation and planning; facilities and other expenses, including rent, depreciation, maintenance of facilities, insurance and supplies; and professional fees for audit, tax and legal services, including legal expenses to pursue patent protection of our intellectual property.

We anticipate that our general and administrative expenses will increase in the future as we increase our headcount to support the expected growth in our research and development activities and the potential commercialization of our product candidates. We also anticipate increased expenses associated with being a public company, including costs related to audit, legal, regulatory and tax-related services associated with maintaining compliance with exchange listing and SEC requirements, director and officer insurance premiums, and investor relations costs. Additionally, if and when we believe that a regulatory approval of the first product candidate appears likely, we anticipate an increase in payroll and related expenses as a result of our preparation for commercial operations, especially as it relates to the sales and marketing of our product candidates.

Other Income (Expense)

Interest income (expense), net. Interest income (expense), net consists of interest earned on our cash and cash equivalents and interest expense on prior debt. Our interest income has not been significant due to low interest earned on invested balances. We anticipate that our interest income will increase in the future due to increased balances from the net proceeds of \$146.8 million we received from our Series B and Series C preferred stock financings in the first quarter of 2014 and our IPO on July 23, 2014.

Other income (expense), net. Other income (expense), net consists of the realized and unrealized net gains and losses from foreign currency-denominated vendor payables.

Critical Accounting Policies and Significant Judgments and Estimates

Our consolidated financial statements are prepared in accordance with generally accepted accounting principles in the United States. The preparation of our consolidated financial statements and related disclosures requires us to make estimates and assumptions that affect the reported amount of assets, liabilities, revenue, costs and expenses, and related disclosures. We believe that the estimates and assumptions involved in the accounting policies described below may have the greatest potential impact on our consolidated financial statements and, therefore, consider these to be our

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critical accounting policies. We evaluate our estimates and assumptions on an ongoing basis. Our actual results may differ from these estimates under different assumptions and conditions.

While our significant accounting policies are described in more detail in the notes to our consolidated financial statements appearing elsewhere in this prospectus, we believe that the following accounting policies are those most critical to the judgments and estimates used in the preparation of our consolidated financial statements.

Accrued Research and Development Expenses

As part of the process of preparing our consolidated financial statements, we are required to estimate our accrued research and development expenses. This process involves reviewing open contracts and purchase orders, communicating with our personnel to identify services that have been performed on our behalf and estimating the level of service performed and the associated costs incurred for the services when we have not yet been invoiced or otherwise notified of the actual costs. The majority of our service providers invoice us in arrears for services performed, on a pre-determined schedule or when contractual milestones are met; however, some require advanced payments. We make estimates of our accrued expenses as of each balance sheet date in our consolidated financial statements based on facts and circumstances known to us at that time. Examples of estimated accrued research and development expenses include fees paid to:

CROs in connection with performing research and development services on our behalf;

investigative sites or other providers in connection with clinical trials;

vendors in connection with non-clinical development activities; and

vendors related to product manufacturing, development and distribution of clinical supplies.

We base our expenses related to clinical trials on our estimates of the services received and efforts expended pursuant to contracts with multiple CROs that conduct and manage clinical trials on our behalf. The financial terms of these agreements are subject to negotiation, vary from contract to contract and may result in uneven payment flows. There may be instances in which payments made to our vendors will exceed the level of services provided and result in a prepayment of the clinical expense. Payments under some of these contracts depend on factors such as the successful enrollment of patients and the completion of clinical trial milestones. In accruing service fees, we estimate the time period over which services will be performed, enrollment of patients, number of sites activated and level of effort to be expended in each period. If the actual timing of the performance of services or the level of effort varies from our estimate, we adjust the accrual or prepaid accordingly. Although we do not expect our estimates to be materially different from amounts actually incurred, our understanding of the status and timing of services performed relative to the actual status and timing of services performed may vary and may result in us reporting expenses that are too high or too low in any particular period. To date, we have not made any material adjustments to our prior estimates of accrued research and development expenses.

Stock-Based Compensation

We measure stock-based awards granted to our employees and nonemployee directors at fair value on the date of grant and recognize the corresponding compensation expense of those awards, net of estimated forfeitures, over the requisite service period, which is generally the vesting period of the respective award. Generally, we issue stock options and restricted stock with only service-based vesting conditions and record the expense for these awards using the straight-line method. We have historically granted stock options with exercise prices equivalent to the fair value of our common stock as of the date of grant.

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We measure stock-based awards granted to nonemployee consultants at the fair value of the award on the date at which the related service is complete. Compensation expense is recognized over the period during which services are rendered by such nonemployee consultants until completed. At the end of each financial reporting period prior to the completion of the service, the fair value of these awards is re-measured using, for options, the then-current fair value of our common stock and updated assumptions in the Black-Scholes option-pricing model and using, for restricted stock, the then-current fair value of our common stock.

The fair value of each stock option grant is estimated using the Black-Scholes option-pricing model. Until July 18, 2014, we were a private company and we lacked company-specific historical and implied volatility information. Considering this and the short history of being a public company, we estimate our expected volatility based on the historical volatility of our publicly traded peer companies and expect to continue to do so until such time as we have adequate historical data regarding the volatility of our traded stock price. The expected term of our options has been determined utilizing the simplified method for awards that qualify as plain-vanilla options, while the expected term of our options granted to consultants and nonemployees has been determined based on the contractual term of the options. The risk-free interest rate is determined by reference to the U.S. Treasury yield curve in effect at the time of grant of the award for time periods approximately equal to the expected term of the award. Expected dividend yield is based on the fact that we have never paid cash dividends and do not expect to pay any cash dividends in the foreseeable future.

The assumptions we used to determine the fair value of stock options granted to employees and directors are as follows, presented on a weighted average basis:

	Year Ended December 31,		
	2014	2013	2012
Expected dividend yield	0.00%	0.00%	0.00%
Expected volatility	98.86%	99.89%	0.00%
Risk free interest rate	1.95%	1.66%	0.00%
Expected life of option	6.38 years	6.04 years	

These assumptions represented our best estimates, but the estimates involve inherent uncertainties and the application of our judgment. As a result, if factors change and we use significantly different assumptions or estimates when valuing our stock options, our stock-based compensation expense could be materially different. We recognize compensation expense for only the portion of awards that are expected to vest. In developing a forfeiture rate estimate for pre-vesting forfeitures, we have considered our historical experience of actual forfeitures. If our future actual forfeiture rate is materially different from our estimate, our stock-based compensation expense could be significantly different from what we have recorded in the current period.

During the years ended December 31, 2014 and 2013, we recognized stock-based compensation expense of \$2,512 and \$61, respectively, of which \$1,093 and \$38, respectively, was recorded as research and development expense and \$1,419 and \$23, respectively, was recorded as general and administrative expense in our statement of operations. During the year ended December 31, 2012, we did not record any stock-based compensation expense, as the amounts were inconsequential.

JOBS Act

On April 5, 2012, the Jumpstart Our Business Startups Act, or the JOBS Act, was signed into law. The JOBS Act contains provisions that, among other things, reduce certain reporting requirements for an emerging growth company. As an emerging growth company, we are electing not to take advantage of the extended transition period afforded by the JOBS Act for the implementation of new or

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revised accounting standards and, as a result, we will comply with new or revised accounting standards on the relevant dates on which adoption of such standards is required for non-emerging growth companies. Section 107 of the JOBS Act provides that our decision not to take advantage of the extended transition period is irrevocable.

In addition, we intend to rely on the other exemptions and reduced reporting requirements provided by the JOBS Act. Subject to certain conditions set forth in the JOBS Act, if as an emerging growth company we intend to rely on such exemptions, we are not required to, among other things, (i) provide an auditor's attestation report on our system of internal controls over financial reporting pursuant to Section 404, (ii) provide all of the compensation disclosure that may be required of non-emerging growth public companies under the Dodd-Frank Wall Street Reform and Consumer Protection Act, (iii) comply with any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation or a supplement to the auditor's report providing additional information about the audit and the consolidated financial statements (auditor discussion and analysis), and (iv) disclose certain executive compensation-related items such as the correlation between executive compensation and performance and comparisons of the Chief Executive Officer's compensation to median employee compensation. These exemptions will apply for a period of five years following the date of our IPO or until we no longer meet the requirements of being an emerging growth company, whichever is earlier.

Our significant accounting policies are described in more detail in Note 2 to our consolidated financial statements appearing elsewhere in this prospectus.

Results of Operations***Comparison of the Years Ended December 31, 2014 and 2013***

The following table summarizes our results of operations for the years ended December 31, 2014 and 2013:

	Year Ended December 31,		Increase (Decrease)
	2014	2013	
	(in thousands)		
Operating expenses:			
Research and development	\$ 24,100	\$ 14,357	\$ 9,743
General and administrative	9,710	3,922	5,788
Total operating expenses	33,810	18,279	15,531
Loss from operations	(33,810)	(18,279)	(15,531)
Interest income (expense), net	8	1	7
Other income (expense), net	(9)	(3)	(6)
Net loss	\$ (33,811)	\$ (18,281)	\$ (15,530)

Table of Contents**Research and development expenses**

	Year Ended December 31, (in thousands)		Increase (Decrease)
	2014	2013	
SAGE-547	\$ 9,137	\$ 3,918	\$ 5,219
SAGE-689	3,058	2,772	286
SAGE-217	2,764	1,129	1,635
Other research and development programs	3,088	3,388	(300)
Unallocated expenses	6,053	3,150	2,903
Total research and development programs	\$ 24,100	\$ 14,357	\$ 9,743

Research and development expenses for the fiscal year ended December 31, 2014 were \$24.1 million, compared to \$14.4 million for the year ended December 31, 2013. The increase of \$9.7 million period over period was primarily due to the following:

an increase of \$5.2 million in expenses of our SAGE-547 program. We initiated the Phase 1/2 clinical trial of SAGE-547 in SRSE in early 2014;

an increase of \$0.3 million in expenses of our SAGE-689 program with advancement of the lead optimization program into IND-enabling non-clinical development (e.g. toxicology studies, process development and drug substance manufacturing);

an increase of \$1.6 million in expenses of our SAGE-217 program with advancement of the lead optimization program into IND-enabling non-clinical development (e.g. toxicology studies, process development and drug substance manufacturing);

a net decrease of \$0.3 million in expenses of our other research and development programs reflecting a focus on advancing SAGE-689 and SAGE-217 into IND-enabling non-clinical development, portfolio priorities and timing of investment in certain research programs; and

an increase of \$2.9 million in employee-related spending to support the growth in our research and development activities, reflecting the effects of hiring additional, full-time employees during 2014.

General and administrative expenses

	Year Ended December 31, (in thousands)		Increase (Decrease)
	2014	2013	
Personnel related	\$ 4,337	\$ 1,764	\$ 2,573
Professional fees	3,788	1,253	2,535
Facilities	370	364	6
Other	1,215	541	674
Total general and administrative expenses	\$ 9,710	\$ 3,922	\$ 5,788

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General and administrative expenses for the year ended December 31, 2014 were \$9.7 million, compared to \$3.9 million for the year ended December 31, 2013. The increase of \$5.8 million in general and administrative expenses was primarily due to the \$2.6 million increase in personnel-related costs due to the effects of hiring additional, full-time employees during 2014 to support operations, finance, human resources and early commercial planning activities as well as an increase

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in stock compensation expense costs and \$2.5 million increase in professional fees associated with being a public company, including costs related to audit, legal, regulatory and tax-related services associated with maintaining compliance with exchange listing and SEC requirements, and investor relations costs.

Other income (expense), net

Interest income (expense), net and other income (expense), net were insignificant for the years ended December 31, 2014 and 2013.

Comparison of the Years Ended December 31, 2013 and 2012

The following table summarizes our results of operations for the years ended December 31, 2013 and 2012:

	Year Ended December 31, 2013 2012 (in thousands)		Increase (Decrease)
Operating expenses:			
Research and development	\$ 14,357	\$ 7,229	\$ 7,128
General and administrative	3,922	2,402	1,520
Total operating expenses	18,279	9,631	8,648
Loss from operations	(18,279)	(9,631)	(8,648)
Interest income (expense), net	1		1
Other income (expense), net	(3)	(1)	(2)
Net loss	\$ (18,281)	\$ (9,632)	\$ (8,649)

Research and development expenses

	Year Ended December 31, 2012 2013 (in thousands)		Increase (Decrease)
SAGE-547	\$ 3,918	\$ 125	\$ 3,793
SAGE-689	2,772	1,047	1,725
SAGE-217	1,129		1,129
Other research and development programs	3,388	3,495	(107)
Unallocated expenses	3,150	2,562	588
Total research and development programs	\$ 14,357	\$ 7,229	\$ 7,128

Research and development expenses for the year ended December 31, 2013 were \$14.4 million, compared to \$7.2 million for the year ended December 31, 2012. The increase of \$7.1 million year over year was primarily due to the following:

an increase of \$3.8 million in expenses of our SAGE-547 program, consisting primarily of external clinical and drug manufacturing costs associated with the preparation of our Phase 1/2 clinical trial of SAGE-547, as compared to only \$0.1 million being spent on the program in 2012;

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an increase of \$1.7 million in expenses of our SAGE-689 program, consisting primarily of external costs related to IND-enabling toxicology and safety pharmacology testing and manufacturing activities that were incurred as that program progressed into non-clinical studies during the second half of 2013;

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	First Quarter	Second Quarter	2013 Third Quarter	Fourth Quarter	Total
	(in thousands, except per share data)				
Total operating expense	\$ 3,389	\$ 4,655	\$ 4,519	\$ 5,716	\$ 18,279
Loss from operations	(3,389)	(4,655)	(4,519)	(5,716)	(18,279)
Net loss and comprehensive loss	(3,389)	(4,655)	(4,519)	(5,718)	(18,281)
Net loss attributable to common stockholders	(3,389)	(4,655)	(4,519)	(5,725)	(18,288)
Net loss per share attributable to common stockholders basic and diluted	\$ (2.39)	\$ (3.18)	\$ (2.98)	\$ (3.65)	\$ (12.26)

Liquidity and Capital Resources

Since our inception in April 2010, we have not generated any revenue and have incurred recurring net losses. As of December 31, 2014, we had an accumulated deficit of \$66.8 million. From our inception through December 31, 2014, we have received net proceeds of \$184.6 million from the sales of redeemable convertible preferred stock, the issuance of convertible notes and the proceeds from our IPO.

In July 2014, we completed our IPO pursuant to a registration statement on Form S-1. In the IPO, we issued and sold an aggregate of 5,750,000 shares of common stock under the registration statement, including the underwriters' exercise in full of their over-allotment option at a public offering price of \$18.00 per share. Net proceeds were approximately \$94.0 million, after deducting underwriting discounts and commissions and offering expenses.

As of December 31, 2014, our primary sources of liquidity were our cash and cash equivalents, which totaled \$127.8 million. We invest our cash equivalents and investments in highly liquid, interest-bearing investment-grade and government securities to preserve principal.

The following table summarizes the primary sources and uses of cash for the periods presented below:

	Years Ended December 31,		
	2014	2013	2012
	(in thousands)		
Net cash provided by (used in):			
Operating activities	\$ (27,042)	\$ (17,516)	\$ (8,926)
Investing activities	(128)	(3)	(111)
Financing activities	146,870	22,783	8,997
Net increase (decrease) in cash and cash equivalents	\$ 119,700	\$ 5,264	\$ (40)

Operating Activities

Operating activities used \$27.0 million of cash during the year ended December 31, 2014. The cash flow used in operating activities resulted primarily from our net loss of \$33.8 million for the period and cash used for changes in our operating assets and liabilities of \$4.1 million and by non-cash charges of \$2.7 million. Our net loss was primarily attributable to research and development activities related to our lead programs in development and our general and administrative expenses, as we had no revenue in the period. Our net non-cash charges during the year ended December 31, 2014 primarily consisted of stock-based compensation expenses of \$2.5 million and non-cash licensing fees

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paid in shares of our common stock of \$0.1 million. Net cash used in changes in our operating assets and liabilities consisted primarily of an increase in accrued expenses and other liabilities of \$4.4 million offset by an increase in prepaid expenses and other current assets of \$0.7 million. Our prepaid expenses and other current assets, accounts payable and accrued expense balances were affected by the timing of vendor invoicing and payments.

During the year ended December 31, 2013, operating activities used \$17.5 million of cash, primarily resulting from our net loss of \$18.3 million, partially offset by cash provided by changes in our operating assets and liabilities and non-cash charges totaling \$0.8 million. Our net loss was primarily attributed to research and development activities related to our lead programs in development and our general and administrative expenses. Net cash provided by changes in our operating assets and liabilities for the year ended December 31, 2013 consisted primarily of an increase in accrued expenses and accounts payable of \$0.9 million partially offset by an increase in prepaid expenses and other current assets of \$0.3 million. Our prepaid expenses and other current assets, accounts payable and accrued expense balances were affected by the timing of vendor invoicing and payments.

During the year ended December 31, 2012, operating activities used \$8.9 million of cash, primarily resulting from our net loss of \$9.6 million, partially offset by cash provided from changes in our operating assets and liabilities of \$0.7 million. Our net loss was primarily attributed to research and development activities related to our lead programs in development and our general and administrative expenses, as we had no revenue in the period. Net cash provided by changes in our operating assets and liabilities during the year ended December 31, 2012 consisted primarily of increases in accounts payable and accrued expenses totaling \$0.6 million. Our accounts payable and accrued expense balances were affected by the timing of vendor invoicing and payments.

Investing Activities

During the year ended December 31, 2014, we used \$0.1 million of cash for purchases of property and equipment. During the year ended December 31, 2013, we had no significant purchases of property and equipment. During the year ended December 31, 2012, we used \$0.1 million for purchases of property and equipment.

Financing Activities

During the years ended December 31, 2014 and 2013, net cash provided by financing activities was \$146.9 million and \$22.8 million, respectively. Net cash provided by financing activities in the year ended December 31, 2014 consisted of \$94.0 million in net proceeds from our IPO on July 23, 2014 and \$52.9 million from the issuance of Series B and Series C redeemable preferred stock and from the exercise of stock options. Net cash provided by financing activities in the year ended December 31, 2013 consisted of \$22.8 million from the issuance of Series A redeemable convertible preferred stock and from the exercise of stock options. During the year ended December 31, 2012, net cash provided by financing activities was \$9.0 million, primarily resulting from the net proceeds we received from the issuance of Series A redeemable convertible preferred stock.

Operating Capital Requirements

To date, we have not generated any revenue from product sales. We do not know when, or if, we will generate any revenue from product sales. We do not expect to generate significant revenue from product sales unless and until we obtain regulatory approval of and commercialize one of our current or future product candidates. We anticipate that we will continue to generate losses for the foreseeable future, and we expect the losses to increase as we continue the development of, and seek regulatory approvals for, our product candidates and begin to commercialize any approved products. We expect

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to incur additional costs associated with operating as a public company. In addition, subject to obtaining regulatory approval of any of our product candidates, we expect to incur significant commercialization expenses for product sales, marketing and manufacturing. Accordingly, we anticipate that we will need substantial additional funding in connection with our continuing operations.

Based on our current operating plan, we expect that our existing cash and cash equivalents as of December 31, 2014, including the net proceeds from our IPO which closed on July 23, 2014, will enable us to fund our operating expenses and capital expenditure requirements for at least the next 12 months. In that time, we expect that our expenses will increase substantially as we advance clinical development of SAGE-547 including completing our Phase 3 clinical trial, fund IND-enabling activities and Phase 1 clinical development for SAGE-689, fund IND-enabling activities for SAGE-217, fund new and ongoing research and development activities and working capital, and fund other general corporate purposes. We have based our estimates on assumptions that may prove to be wrong, and we may use our available capital resources sooner than we currently expect. Because of the numerous risks and uncertainties associated with the development and commercialization of our product candidates, we are unable to estimate the amounts of increased capital outlays and operating expenditures necessary to complete the development and commercialization of our product candidates.

Our future capital requirements will depend on many factors, including:

the costs, timing, and outcome of regulatory reviews and approvals;

the ability of our product candidates to progress through clinical development successfully;

the initiation, progress, timings, costs, and results of non-clinical studies and clinical trials for our other programs and potential product candidates;

the number and characteristics of the product candidates we pursue;

the costs of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending intellectual property-related claims;

the extent to which we acquire or in-license other products and technologies; and

our ability to establish any future collaboration arrangements on favorable terms, if at all.

Until such time, if ever, as we can generate substantial product revenue, we expect to finance our cash needs through a combination of equity offerings, debt financings, collaborations, strategic alliances and licensing arrangements. To the extent that we raise additional capital through the sale of equity or convertible debt securities, the ownership interest of our stockholders will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Debt financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends and may require the issuance of warrants, which could potentially dilute your ownership interest. If we raise additional funds through collaborations, strategic alliances or licensing arrangements with third parties, we may have to relinquish valuable rights to our technologies, future revenue streams or research programs or to grant licenses on terms that may not be favorable to us. If we are unable to raise additional funds through equity or debt financings when needed, we may be required to delay, limit, reduce or terminate our product development or future commercialization efforts or grant rights to develop and market products or product candidates that we would otherwise prefer to develop and market ourselves.

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The following table summarizes our contractual obligations at December 31, 2014 and the effect such obligations are expected to have on our liquidity and cash flow in future periods:

	Total	Payments Due by Period			More Than 5 years
		Less Than 1 year	1-3 Years (in thousands)	3-5 Years	
Operating lease commitments ⁽¹⁾	\$ 575	\$ 262	\$ 313	\$	\$
Total ⁽²⁾⁽³⁾⁽⁴⁾	\$ 575	\$ 262	\$ 313	\$	\$

- (1) We lease office space in Cambridge, Massachusetts under an operating lease agreement that initially expires on February 28, 2017. The minimum lease payments in the table do not include related common area maintenance charges or real estate taxes, which costs are variable.
- (2) We have acquired exclusive and non-exclusive rights to use, research, develop and offer for sale certain products and patents under three separate licensing agreements, including amendments entered into in April and May 2014, with Washington University, CyDex Pharmaceuticals, Inc. and The Regents of the University of California. The licensing rights obligate us to make payments to the licensors for license fees, milestones, license maintenance fees and royalties. We are obligated to make future milestone payments under these agreements of up to \$29.8 million upon achieving certain pre-commercialization milestones, such as clinical trials and regulatory approvals. We reasonably anticipate that we may be required to pay \$0.5 million of milestone payments in 2015, provided various development milestones are achieved. Amounts related to contingent milestone payments are not considered contractual obligations as they are contingent on the successful achievement of certain milestones. These milestones may not be achieved. Because the achievement of these milestones had not occurred as of December 31, 2014, no liabilities for such contingencies have been recorded in our consolidated financial statements. In addition, under the licensing agreements, we will owe single-digit royalties on sales of commercial products, if any, developed using the licensed technologies. Under two of these license agreements, we are obligated to pay to the licensors a percentage of fees received if and when we sublicense the technologies. As of December 31, 2014, we had not developed a commercial product using the licensed technologies and we had not entered into any sublicense agreements for the technologies. We have not included any of these amounts in the table as we cannot estimate or predict when, or if, these amounts will become due.
- (3) We enter into contracts in the normal course of business with CROs for clinical trials, non-clinical research studies and testing, manufacturing and other services and products for operating purposes. These contracts generally provide for termination upon notice, and therefore we believe that our non-cancelable obligations under these agreements are not material.
- (4) Under a January 2014 consulting agreement, we are obligated to make milestone payments of up to \$2.0 million and to issue up to 126,984 shares of our common stock to a nonemployee consultant upon achieving certain clinical trial milestones and regulatory approval milestones. As of December 31, 2014, we paid \$50,000 and issued 15,872 shares of common stock relating to this consulting agreement. We have not included remaining amounts in the table as we cannot estimate or predict when, or if, these amounts will become due.

Off-Balance Sheet Arrangements

We do not currently have, nor did we have during the periods presented, any off-balance sheet arrangements as defined by SEC rules.

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Quantitative and Qualitative Disclosures about Market Risk

We had cash and cash equivalents of approximately \$127.8 million at December 31, 2014. The primary objectives of our investment activities are to preserve principal, provide liquidity and maximize income without significantly increasing risk. Our primary exposure to market risk relates to fluctuations in interest rates, which are affected by changes in the general level of U.S. interest rates. Given the short-term nature of our cash and cash equivalents, we believe that a sudden change in market interest rates would not be expected to have a material impact on our financial condition and/or results of operation. We do not have any foreign currency or other derivatives financial instruments.

We do not believe that our cash and cash equivalents have significant risk of default or illiquidity. While we believe our cash and cash equivalents do not contain excessive risk, we cannot provide absolute assurance that in the future our investments will not be subject to adverse changes in market value. In addition, we maintain significant amounts of cash and cash equivalents at one or more financial institutions that are in excess of federally insured limits.

Inflation generally affects us by increasing our cost of labor and clinical trial costs. We do not believe that inflation had a material effect on our results of operations during the year ended December 31, 2014.

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BUSINESS

Overview

We are a biopharmaceutical company committed to developing and commercializing novel medicines to treat life-threatening, rare central nervous system, or CNS, disorders, where there are inadequate or no approved existing therapies. We are targeting CNS indications where patient populations are easily identified, acute treatment is typically initiated in the hospital setting, clinical endpoints are well-defined, and development pathways are feasible.

Our initial product candidates, which are summarized in the table below, are aimed at treating different stages of status epilepticus, or SE, a life-threatening condition in which the brain is in a state of persistent seizure, as well as other seizure and non-seizure disorders. The lead product candidate in our SE program, SAGE-547, is an intravenous, or IV, agent entering Phase 3 clinical development as an adjunctive therapy, a therapy combined with current therapeutic approaches, for the treatment of super-refractory SE, or SRSE. The current standard of care for SRSE is empiric, and there are no therapies at present that have been specifically approved for this indication. Over the course of 2014, the U.S. Food and Drug Administration, or FDA, granted us orphan drug designation and Fast Track designation for our investigational new drug application for SAGE-547 as a treatment for SRSE. On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE and we expect to initiate the Phase 3 trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of a New Drug Application, or NDA, submission for SAGE-547.

We continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for essential tremor, a debilitating neurological disorder that causes involuntary, rhythmic shaking with no known cause, and severe post-partum depression, a distinct and readily identified form of major depressive disorder estimated to affect up to 20% of women following childbirth. If these exploratory trials are successful, we plan to use the data from them to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

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Our next-generation product candidates, SAGE-689 and SAGE-217, utilize similar mechanistic pathways as SAGE-547 and are designed to have pharmaceutical properties which optimize both their non-clinical profiles and potential clinical profiles for the treatment of different stages of SE, as well as other seizure and non-seizure disorders.

SE is diagnosed when a patient has a seizure lasting longer than five minutes, and is associated with substantial morbidity and mortality. We estimate that in the United States each year there are up to 150,000 cases of SE, of which 30,000 SE patients die. We estimate that there are 35,000 patients with SE in the United States that are hospitalized in the intensive care unit, or ICU, each year. An SE patient is first treated with benzodiazepines, or BDZs, and if no response then treated with other, second-line, anti-seizure drugs. If the seizure persists after second-line therapy the patient is diagnosed as having refractory SE, or RSE, admitted to the ICU and placed into a medically induced coma. Currently, there are no therapies that have been specifically approved for refractory SE, or RSE; however, physicians typically use anesthetic agents to induce the coma and stop the seizure immediately. After a period of 24 hours, an attempt is made to wean the patient from the anesthetic agents to evaluate whether or not the seizure condition has resolved. Unfortunately, not all patients respond to weaning attempts, in which case the patient must be maintained in the medically induced coma. At this point, the patient is diagnosed as having SRSE.

Prior to the start of our Phase 1/2 clinical trial of SAGE-547, we began to collect data in emergency-use of SAGE-547 that we believe supports the safety and activity of SAGE-547 for treatment of SRSE. As of January 9, 2015, ten patients were treated with SAGE-547 by independent centers under emergency-use Investigational New Drug Applications, or INDs. Each individual case of SRSE arose from a variety of underlying etiologies, the patients were of varying ages, and all patients had been placed in a long-duration medically induced coma prior to the administration of SAGE-547. We experienced an overall response rate of 78% in seven of the nine evaluable patients.

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In January 2014, we commenced our Phase 1/2 clinical trial to study safety, tolerability and efficacy of SAGE-547 in adult patients with SRSE. This clinical trial is designed as an open-label trial in at least ten patients diagnosed with SRSE. In October 2014, the FDA approved a protocol amendment for our Phase 1/2 trial that enables us to treat pediatric patients as young as two years old, to increase the dose of SAGE-547 being administered to patients and to increase the duration of treatment to up to two weeks. As of February 28, 2015, there were 17 active trial sites in the United States. We are continuing to enroll patients as an expansion cohort in this trial and we anticipate reporting final clinical data from this Phase 1/2 clinical trial of SAGE-547 at the Antiepileptic Drug and Device Trials XIII Conference, which is taking place May 13-15, 2015.

On January 9, 2015, we reported results from our Phase 1/2 clinical trial. Consistent with topline data announced in November 2014, the primary endpoint of safety and tolerability, was achieved in all patients. Of the 20 patients enrolled in the Phase 1/2 clinical trial, 17 patients were evaluable for efficacy. 71% of evaluable patients met the key efficacy endpoint of being successfully weaned off their anesthetic agents while SAGE-547 was being administered. In addition, 71% of evaluable patients were successfully weaned off SAGE-547 without recurrence of SRSE. As a group, patients who responded to SAGE-547 generally demonstrated rapid improvement over the first five days following treatment. Patients who responded also continued to improve over the 30-day follow-up period as assessed by several measures. SAGE-547 was generally well-tolerated and no drug-related serious adverse events, as determined by the Safety Review Committee, were reported in treated patients. In the 20 patients treated with SAGE-547, the mean exposure level of SAGE-547 was approximately 200nM.

On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE. Subject to submission and review by the FDA of a final protocol for the planned Phase 3 clinical trial and updated chemistry, manufacturing and controls information, we expect to initiate the trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547.

We continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for additional indications. In October 2014, we began patient enrollment in an exploratory Phase 2a clinical trial of SAGE-547 in patients with essential tremor, a debilitating neurological disorder that causes involuntary, rhythmic shaking with no known cause. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and activity of SAGE-547 in patients with essential tremor. In January 2015, we initiated a Phase 2a clinical trial of SAGE-547 in women with severe postpartum depression, or severe PPD, a distinct and readily identified form of major depressive disorder estimated to affect up to 20% of women following childbirth. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and efficacy of SAGE-547 for the treatment of severe PPD. We plan to use the data from these exploratory trials to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

SAGE-689 and SAGE-217 are two additional product candidates in our pipeline, which are currently in IND-enabling toxicology and safety pharmacology testing. SAGE-689 is being developed as an adjunctive second-line therapy for the treatment of SE. We are currently conducting IND-enabling studies of SAGE-689, with a plan to file an IND in late 2015 and to begin a Phase 1 clinical trial thereafter. SAGE-217 is being developed as an oral monotherapy for orphan epilepsies, such as Dravet syndrome and Rett syndrome. The chemical characteristics of SAGE-217 potentially allow formulation as both an IV and oral medication. In addition, we believe related molecules from our portfolio may be useful in the treatment of a variety of neurological and psychiatric disorders, including, for example, Fragile X syndrome, anxiety and tremor. We are currently conducting IND-enabling studies of SAGE-217 with a plan to file an IND by late 2015 and to begin a Phase 1 clinical trial thereafter.

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Our current near-term product candidates are allosteric modulators of both synaptic and extrasynaptic, or existing outside of the synapse, GABA_A receptors, a characteristic important in distinguishing our approach from current therapies. While altering the level of synaptic GABA_A receptor activity can be beneficial in stopping seizures, this approach has limitations for the treatment of SE. As SE progresses in many patients, select synaptic GABA_A receptors are down-regulated, or removed from the neuronal synaptic surface. As a result, drugs that target down-regulated receptors, such as benzodiazepines, or BDZs, often are not effective in stopping SE. In contrast, our product candidates work at both the synaptic and extrasynaptic GABA_A receptors. Non-clinical studies suggest that these extrasynaptic GABA_A receptors remain fully active during SE, offering the potential for drugs that impact GABA via the extrasynaptic GABA_A receptor to alter GABA_A activity and abate seizure. We believe that by creating compounds that target both these receptors, we may be successful in treating seizures that do not respond to BDZ therapy.

Now and in the foreseeable future, our product development pipeline will be focused on allosteric modulation of two important receptor systems in the brain GABA_A and NMDA. These receptor systems regulate inhibitory and excitatory neurotransmission, respectively. The GABA_A and NMDA receptor systems are broadly accepted as impacting many psychiatric and neurological disorders, spanning disorders of mood, seizure, cognition, anxiety, sleep, pain, epilepsy, and movement disorders among others. Thus these receptor systems are widely regarded as validated drug targets for a variety of CNS disorders, with decades of research and multiple approved drugs targeting these receptor systems. Drugs approved to modulate these receptor systems have had safety and efficacy limitations related to their poor pharmaceutical properties and adverse side effects. We believe that we will have the opportunity to develop molecules from our internal portfolio to more effectively address many of these disorders in the future.

Our ability to identify and develop such novel CNS therapies is enabled by our proprietary chemistry platform that is centered on a scaffold of chemically modified endogenous neuroactive steroid compounds. We believe our know-how around the chemistry and activity of allosteric modulators allows us to efficiently design molecules with different characteristics by enabling us to control important properties such as half-life, brain penetration and the types of receptors with which our drugs interact. Therefore, we believe our product candidates will have the potential to bind with targets in the brain with more precision, increased safety and tolerability, and fewer off-target side effects than either current CNS therapies or previous therapies, which have often failed in development.

We were founded in 2010, based on leading research in the areas of brain function and neuroactive steroids, to explore novel approaches to CNS therapeutics. Since our inception, we have continued to expand our know-how of CNS therapeutics through our research and development programs and to pursue intellectual property protection for our proprietary chemistry platform. In addition, we have assembled a world-class management team that together has been a part of the successful discovery, development and commercialization of more than 20 marketed CNS therapies.

Our Strategy

Our goal is to become a leading biopharmaceutical company focused on development and commercialization of novel proprietary therapies for the treatment of life-threatening, rare CNS disorders.

Key elements of our strategy are to:

Rapidly advance SAGE-547 as a treatment for SRSE. We are developing SAGE-547 as an adjunctive therapy for the treatment of SRSE. We expect to initiate our planned Phase 3 clinical

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trial for SAGE-547 in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547.

Utilize SAGE-547 in exploratory trials to help guide the development of second generation GABA_A receptor modulators for the applicable diseases. We are using SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for additional indications. The first two indications are essential tremor and severe PPD. The trials are designed to evaluate safety, tolerability, pharmacokinetics, and efficacy. We will use the data from the exploratory trials to help guide the design of second-generation GABA_A receptor modulators for the chronic treatment of these diseases.

Develop our next generation product candidates SAGE-689 and SAGE-217, in parallel with SAGE-547. Our follow-on product candidates, SAGE-689 and SAGE-217, will utilize similar mechanistic pathways as SAGE-547 and are designed to have pharmaceutical properties that optimize their non-clinical profiles and potential clinical profiles for the treatment of RSE and orphan epilepsies, such as Dravet syndrome and Rett syndrome.

Enhance the probability of success in treating SE by developing unique assets with differentiated features. Our initial product candidates are all positive allosteric modulators of the synaptic and extrasynaptic GABA_A receptor. GABA is the major inhibitory neurotransmitter in the CNS and mediates downstream neurologic and bodily function via activation of GABA_A receptors. However, while their mechanisms are similar, our newer compounds are differentiated from SAGE-547, in terms of their activity and pharmacokinetic profiles affording compounds with superior sedation properties or the opportunity to be dosed orally. Thus, while success with SAGE-547 would augur well for our earlier-stage compounds, the profiles of our new GABA_A agents may allow better risk-benefit. All of our initial SE product candidates represent a class of selective agents that target both GABA_A synaptic and extrasynaptic receptors that we believe can overcome the tolerability and sedation limitations of existing GABA_A targeted agents for the treatment of SE, including BDZs.

Grow our pipeline more broadly utilizing the strengths of our proprietary chemistry platform and scientific know-how, to lessen our long-term reliance on a single franchise and facilitate long-term growth. The potential of our GABA_A platform goes beyond treatment of SE, our initial focus. We will have the potential to discover and develop GABA_A receptor agents with differentiated selectivity for GABA_A synaptic and extrasynaptic receptors, as well as having differing half-lives and routes of administration. These new molecules may have the potential to treat a wide range of psychiatric and neurological disorders, such as fragile X syndrome, anxiety, depression, sleep disorders, mania, tremor, tinnitus and post-traumatic stress disorder. Similarly, our NMDA platform will be explored to develop positive and negative allosteric modulators of NMDA receptors. These molecules may find use in the treatment of depression, Alzheimer's disease, attention deficit hyperactivity disorder and schizophrenia, as well as certain aspects of Huntington's disease and neuropathic pain. We believe our capacity to develop unique molecules creates important optionality for us; in the event any particular program should not meet its desired endpoint.

Focus our internal development activities on CNS indications where we can make well-informed, rapid go/no-go decisions. We believe our ability to design molecules that target CNS indications where patient populations are easily identified and, where well-defined objective endpoints and development pathways exist, allows us to make highly informed decisions when advancing our product candidates. For example, the information we learned with respect to SAGE-547 through our emergency-use cases in SRSE, provided us with a core understanding of the potential utility of this compound in our ongoing and planned clinical trials in the SRSE patient population.

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Build a commercial capability to bring our CNS therapeutics to physicians and patients for rare target indications. We are concentrating our internal efforts on CNS disorders where we believe the target commercial audience can be reached utilizing a highly specialized sales force similar to those of other rare-disease companies. As a result, we believe we can successfully launch and commercialize our initial product candidates on our own. In addition, SAGE-547, if approved, will reach the market in advance of our next product candidate, and therefore will allow physicians and hospitals sufficient time to use SAGE-547 and gain familiarity with its mechanism of action, which we believe has the potential to accelerate adoption of our follow-on products, SAGE-689 and SAGE-217.

Selectively partner our programs to enhance our value. We believe that we are differentiated in our ability to create or develop proprietary novel molecules that impact validated targets such as GABA_A and NMDA receptors for a wide variety of CNS indications. As a result we have identified potential drug candidates that may have advantageous profiles compared to existing and development-stage therapies for large underserved CNS indications, such as depression and cognition. Given the large number of potential patients and physicians for these indications, we may enter into selective partnerships with companies who have clinical expertise and pre-existing commercial infrastructure in areas such as depression and cognition, in order to accelerate development or maximize our return on investment.

Understanding the Foundations of Our Approach

Neurotransmission

The CNS is composed of a vast and complex network of different structures and cell types, most of which serve directly or indirectly to provide a means for the nervous system to signal or communicate with other nerve cells in order to regulate and control all brain function. The cell type responsible for this signaling is called a neuron. Chemical or electrical signals can exert their effects on neurons by traveling across a physical gap located between two neurons, called a synapse. Presynaptic neurons transmit signals whereas postsynaptic neurons react to the signals. The human brain contains approximately 86 billion neurons, each having hundreds to tens of thousands of synapses to allow for this communication. This process is essential to all things, from organ function, to movement, to memory and all behavioral processes.

Neurotransmission is the process by which signaling molecules, called neurotransmitters, are released by a presynaptic neuron, travel over the synaptic space and bind to and interact with receptors on a postsynaptic neuron. Depending on the nature of the neurotransmitter and receptor, this interaction results in excitation, inhibition or modulation of the receiving neuron's behavior.

Synaptic receptors are primarily located inside the synaptic cleft, or the space where the neurons communicate, and have been historically considered to be the most important part of the neuron. However, recent understanding of neurotransmission and brain function has shown there are many extrasynaptic receptors that also respond to neurotransmitters to exert their effects. For example, it is becoming increasingly understood that extrasynaptic GABA_A receptor-mediated neurotransmission is critical to generalized neurological function and has demonstrated influence over general physiological states such as sleep, hunger, anxiety and seizure among other things.

Allosteric modulation

We are focused on developing drugs based on selective allosteric modulation of key CNS synaptic and extrasynaptic receptors. Molecules that function directly on synaptic or extrasynaptic receptors at the site where the native, or natural, molecule binds to inhibit or activate them are known

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as orthosteric. Alternatively, allosteric modulators are a class of small molecules very different from classical orthosteric drugs, as they interact at a site different from the native site and allow for fine-tuning of neuronal signals.

Orthosteric drugs aimed at these key synaptic receptors are inherently limited due to their targeted effects of complete activation or complete inhibition of the neuron, with little subtlety in how they exert their effect. As a result, neurons are unable to respond to normal stimuli and can become over-stimulated by a neurotransmitter or be unable to respond to normal neurotransmission, thus negatively impacting both the efficacy and safety profile of a potential CNS drug. We believe that nowhere in the body is it more important to maintain normal rhythms than in the brain, and accordingly we believe that allosteric modulation approaches are better suited for the treatment of seizure and other CNS diseases.

We utilize our proprietary chemistry capabilities to design and identify drugs that are allosteric modulators that bind to either or both synaptic and extrasynaptic receptors. As a result, our drugs under development are capable of varying degrees of desired activity rather than complete activation or inhibition of the receptor as typically observed with orthosteric drugs. We believe this greater selectivity and modulatory control at extrasynaptic GABA_A receptors may allow us to develop CNS drugs that offer significant therapeutic and safety advantages over orthosteric drugs.

Allosteric modulation of extrasynaptic GABA_A receptors to treat SE

Our initial focus is on the development of positive allosteric modulators of both synaptic and extrasynaptic sites of the GABA_A receptor. BDZs are allosteric modulators that primarily act at a particular receptor, the synaptic GABA_A receptor, with little or no activity at extrasynaptic GABA_A receptors. BDZs have many positive drug-like attributes, including safety in overdose, reproducible dosing and predictable actions in humans. However, BDZs are inherently limited due to abuse potential, sedation, memory and performance impairment, and development of tolerance. We believe that our approach to GABA_A receptor allosteric modulation has the potential to be superior to BDZs because our products target both synaptic and extrasynaptic receptors. Therefore we believe we can enhance the potential utility of modulating the GABA_A receptor for new indications, and effectively avoid some of the limitations of BDZs.

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SE patients are often considered to be resistant to the action, or pharmacology, of drugs that only target the synaptic GABA_A receptors, such as BDZs, the first-line therapy for SE. Positively modulating, or up-regulating, GABA_A receptors results in a beneficial effect in some patients with seizures. However, in persistent SE, synaptic GABA_A receptors are down-regulated, or diminished in their activity.

Our initial product candidates are focused on allosteric modulation of both the synaptic and extrasynaptic GABA_A receptors unlike BDZs that primarily interact only with synaptic GABA_A receptors. The extrasynaptic GABA_A receptor is structurally distinct, possesses unique pharmacology and is located in a different place than the synaptic GABA_A receptor. In addition, the extrasynaptic GABA_A receptor remains intact during prolonged periods of seizure with no down-regulation. Since our selective allosteric GABA_A receptor modulators target both the extrasynaptic and synaptic GABA_A receptors, we believe they can treat seizures that are otherwise BDZ-resistant.

Published non-clinical testing utilizing well-validated animal models of SE and sophisticated instruments for identifying the expression of both synaptic and extrasynaptic GABA_A receptors on the surface of neurons support this hypothesis. These studies, performed in rats, demonstrate the reduced number and activity of synaptic GABA_A receptors during SE, in contrast to the preserved number and activity of extrasynaptic GABA_A receptors under the same conditions. These studies were done by measuring the amount of GABA_A synaptic and GABA_A extrasynaptic receptors that are present on the surface of the neurons. The analysis of protein present for each of the respective receptors in animals in the SE-state, versus normal animals, shows the difference in GABA_A receptor expression.

We believe animal models of seizure also portray the advantages of our allosteric approach over therapy with BDZs. The figure below shows the results of a rodent study where the subject animals were placed into an SE-like condition of prolonged seizure resulting in continuous spontaneous seizures. SAGE-547 was then administered to certain animals while the others received a BDZ. The results demonstrate that BDZs are unable to adequately control the seizure condition that we believe is due to down-regulation of synaptic GABA_A receptors. In contrast, SAGE-547, working at both synaptic and extrasynaptic GABA_A receptors, appears to have treated the seizures in these animals and resolved their SE.

Allosteric modulation of NMDA receptors to address other CNS conditions

Orthosteric drug candidate approaches to modulating the NMDA receptor have also been fraught with difficulties. NMDA receptor antagonists have been explored for treating Alzheimer's disease and neuropathic pain and for inducing anesthesia. Drugs that antagonize NMDA receptors are limited by

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adverse effects, such as neurotoxicity, deteriorating mental status and psychotomimetic, or the onset of psychotic symptoms following the administration of the drug. NMDA receptor agonists have been tested in schizophrenia and many believe may have a role in enhancing cognition and mood. However, their ability to be used at effective doses in humans is limited by non-clinical findings indicating these agents may induce cell death through excess excitation of nerve cells.

We are evaluating in non-clinical testing a number of positive and negative allosteric modulators of the NMDA receptor that we believe can overcome the difficulties associated with orthosteric approaches. Like our GABA_A allosteric modulators, our NMDA receptor allosteric modulators work at sites located in the synaptic and extrasynaptic spaces of the neuron and enhance, or modulate, the activity of the native molecule without directly activating the NMDA receptor. Initial non-clinical testing of our NMDA receptor allosteric modulators has indicated we can avoid the excitotoxicity and psychotomimesis seen with directly activating, orthosteric compounds. This in turn may allow us to discover and develop, alone or with partners, compounds to treat conditions such as depression, Alzheimer's disease, attention deficit hyperactivity disorder and schizophrenia, as well as certain aspects of Huntington's disease, and neuropathic pain. In addition, we are evaluating development of these molecules for rare and genetically defined populations where modulation of NMDA may have therapeutic benefit.

Our proprietary chemistry platform

Our proprietary chemistry platform is centered on novel chemical scaffolds of endogenous or chemically modified synthetic neuroactive steroid compounds that are allosteric modulators of GABA_A or NMDA receptors. We have leveraged this platform to assemble a chemistry portfolio of greater than 1,700 compounds. We believe our proprietary chemistry platform allows us to:

optimize the properties of neuroactive steroid compounds to develop proprietary, new chemical entities, with the potential to be used as oral, IV or intramuscular therapies;

control important properties such as half-life, brain penetration and the types of receptors our drugs act upon, thereby modulating either inhibition or excitation either acutely or chronically; and

create drugs that exert control over the intensity of receptor activation or deactivation, with the potential to hit targets in the brain with more precision, increased safety and tolerability and fewer off-target side effects than current CNS therapies.

Our Product Pipeline

We are focusing our efforts on developing product candidates that are derived from active endogenous steroids with properties that selectively target synaptic and extrasynaptic GABA_A or NMDA receptors. We believe that our proprietary approach to drug discovery and development enables us to create both IV and orally bioavailable selective allosteric modulators that can be applied to multiple CNS target indications. The lead product candidates, SAGE-547, SAGE-689 and SAGE-217, all allosteric modulators of the GABA_A receptor, are being developed as treatments for SRSE, RSE and orphan epilepsies, respectively. We intend to develop and commercialize these initial product candidates on our own, if approved. We are also developing additional potential product candidates from both our GABA_A and NMDA receptor programs that will serve mood and cognitive disorders, such as Alzheimer's Disease and depression, which we may choose to selectively partner in select geographies or commercial settings.

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Status Epilepticus (SE) Development Program

Status epilepticus (SE)

Seizures are brief episodes of abnormal excessive or synchronous neuronal activity in the brain. The outward effect can vary from rapid uncoordinated movement of the trunk and extremities, known as tonic-clonic seizure, to a brief loss of awareness, known as an absence seizure. An electroencephalogram, or EEG, is a measurement of electrical activity within neurons of the brain. Each line of an EEG represents a different region of the brain and becomes aberrant during a state of seizure. In cases of recurring or frequent seizures, or with persistent and long seizures, uncontrolled neurotransmission results in remodeling or changes to brain synaptic function. These physiological and anatomical changes to the brain include changes to the receptor systems of neurons, and shape of the neuron thereby impacting its ability to function, resulting in disorganization of brain proteins and potentially neuronal death.

SE is a life-threatening seizure condition in which the brain is in a state of persistent seizure and there is uncontrolled neuronal excitation. The Neurocritical Care Society defines SE as one continuous unremitting seizure lasting longer than five minutes, or recurrent seizures without regaining consciousness between seizures for greater than five minutes. Common causes of SE in adults include preexisting epilepsy, cerebrovascular disease, metabolic and electrolyte disturbances, encephalopathies, head trauma and drug or substance intoxication. SE is more common in children, often as a result of high fever during the first year of life. It is the most common neurologic emergency in pediatric practice.

SE is associated with substantial morbidity and mortality. We estimate that in the United States each year, there are up to 150,000 cases of SE, of which 30,000 SE patients die. We estimate that 35,000 patients with SE in the United States are hospitalized in the ICU each year. This results in an overall inpatient cost of \$3.8 billion to \$7.0 billion per year in the United States.

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SE treatment paradigm

The numbers in the chart above represent the estimated number of U.S. patients affected by SE at various stages each year.

SE is a medical emergency and is treated with aggressive pharmacological approaches. When a patient first presents with SE, medical personnel, typically emergency medical technicians at the scene of the seizure or during emergency transport, will treat the patient with IV BDZs such as diazepam, lorazepam or midazolam. Approximately 65% of patients treated with IV BDZs will respond to such treatment, and the seizure will be abated. If the patient does not respond, he or she will be brought to an emergency room where anti-seizure drugs such as phenytoin or valproic acid, will be administered.

If a patient's SE continues after administration of BDZs and anti-seizure drugs, the patient is diagnosed as having RSE, which must be treated quickly to cease the seizure activity, maintain the patient's airway and prevent brain damage. RSE patients are immediately admitted to the ICU and placed in a medically induced coma to stop all seizure-related activity. Currently, there are no therapies that have been specifically approved for RSE. The primary drugs used to induce coma are continuously infused IV agents such as propofol, midazolam or pentobarbital.

The RSE patient is continually monitored through EEG to ensure burst suppression is achieved. Burst suppression is a pattern of brain waves seen on an EEG characterized by periods of activity alternating with periods of little or no activity in the brain, informing medical personnel of the effectiveness of the medically induced coma. The goal of burst suppression is to allow the brain and corresponding neuronal tissue to restore function and reset to normal pre-seizure levels. After a short period of burst suppression, typically 24 hours, physicians attempt to wean the patient from the medically induced coma to evaluate EEG activity to assess if the neuronal activity has returned to normal levels. If unsuccessful, the patient is placed back into the medically induced coma in order to protect underlying neurological activity and brain function. At this point, patients are considered to be in a state of SRSE.

Currently, there are no therapies that have been specifically approved for SRSE. Treatment approaches consist of diagnosis and treatment of underlying medical conditions, adding and changing anti-seizure drugs, and repeated attempts at weaning in conjunction with IV anesthetic agents. The majority of SRSE patients either die or have significant comorbidities, such as decreased blood pressure and cardiorespiratory collapse.

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Our SE product candidates

SAGE-547

SAGE-547, a proprietary formulation of allopregnanolone, is a known metabolite of progesterone formed in the CNS in humans through the actions of two enzymes. SAGE-547 is being developed as an IV adjunctive therapy in conjunction with underlying anesthesia as a treatment for SRSE. SAGE-547 is currently entering Phase 3 clinical development for the treatment of SRSE. Over the course of 2014, the FDA, granted us orphan drug designation and Fast Track designation for our IND application for SAGE-547 as a treatment for SRSE. On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE and we expect to initiate the Phase 3 trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547.

We believe SAGE-547 has an optimal profile for the treatment of SRSE. SAGE-547 has a wide therapeutic window that allows for allosteric modulation of the GABA_A receptor both synaptically and extrasynaptically without inducing deep anesthesia. The pharmacological properties of SAGE-547, including a short half-life of one hour, allows for continuous IV administration. The ability to titrate SAGE-547 creates the opportunity to tailor therapy to a specific SRSE patient's needs as well as to efficiently administer and withdraw the compound.

Non-clinical results

We believe the clinical development program for SAGE-547 is supported by significant non-clinical data and strong scientific rationale. There are numerous reports that demonstrate the

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non-clinical efficacy of allopregnanolone as well as multiple studies that we have conducted showing the in vitro and in vivo pharmacologic efficacy of SAGE-547 in seizure models, thus providing a strong non-clinical rationale for SAGE-547 in certain forms of seizure, such as SRSE.

A comprehensive toxicology dose escalating study exploring the effects of SAGE-547 in multiple species (rat and dog) is currently underway. We submitted the first study to the FDA in the second quarter of 2014, and the next study was completed in the first quarter of 2015. We must complete these required studies in order to commence a pivotal clinical trial for SAGE-547. Remaining studies must be completed prior to NDA submission.

Emergency-use experience with SAGE-547

We have compiled evidence of activity with SAGE-547 in emergency-use settings that support the safety and potency of SAGE-547 for the treatment of SRSE. As of January 9, 2015, a total of ten patients, six males and four females with a mean age of 17, had been treated with SAGE-547 by independent centers under emergency-use INDs. Of note, each individual case of SRSE arose from a variety of underlying etiologies, the patients were of varying ages, and all patients had been placed in a long-duration medically induced coma. In each case SAGE-547 was administered with a target steady-state exposure similar to that used in our ongoing Phase 1/2 clinical trial. Seven of the nine patients that were evaluable and treated with SAGE-547 achieved resolution of SRSE either during the course of or soon after SAGE-547 treatment for an overall response rate of 78%.

* *Estimated*

We can provide no assurance that the positive results observed to date in emergency-use cases are attributable to SAGE-547; as such cases were not carried out in the controlled environment of a clinical trial. Further, we can provide no assurance that the administration of SAGE-547 to other patients in our clinical trials or otherwise will have positive results.

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Phase 1/2 Clinical Trial

In October 2013, we filed an IND for SAGE-547 for the treatment of SRSE with the FDA, and we received notification allowing us to proceed with our Phase 1/2 clinical trial of SAGE-547 in November 2013. We commenced our Phase 1/2 clinical trial to study safety, tolerability and efficacy of SAGE-547 in adult patients with SRSE in January 2014. This clinical trial is an open-label trial in at least ten patients diagnosed with SRSE. As of February 28, 2015, there were 17 active trial sites in the United States. An SE patient who has failed therapy with first- and second-line agents and has failed IV general anesthesia, or GA, administered over 24 hours is eligible to be included in this trial. Patients will be excluded from participation in this trial if their SE is due to anoxic brain injury or they have end-organ damage of any major organ, such as the brain, the liver or the heart, which would make recovery from either SE or the underlying condition highly unlikely.

The figure below demonstrates the design of the screening and treatment periods of this Phase 1/2 clinical trial. Following the treatment period, there is an acute two-day follow-up period and an extended three-week follow-up period.

The primary endpoints of this trial are to evaluate the safety and tolerability of SAGE-547 in SRSE patients. Safety and tolerability will be assessed by monitoring adverse events, EEG, physical examinations, neurological examinations, vital signs, clinical laboratory measures, electrocardiograms and concomitant medication usage. The secondary endpoint of this trial is to assess the efficacy of SAGE-547 on SRSE, assessed by the need to place the patient back into a medically induced coma for seizure control during administration of SAGE-547, as well as the duration of the observed response. In order to allow full assessment of pharmacologic activity, this trial employs broad inclusion criteria, primarily excluding patients only if there is major damage to the brain, such as anoxic injury, devastating stroke or the presence of a large lesion. Other secondary objectives used to measure efficacy include scores on global and specific scales relating to cognition, agitation and depth of coma and survival.

On January 9, 2015, we reported results from our Phase 1/2 clinical trial. Consistent with topline data announced in November 2014, the primary endpoint, safety and tolerability, was achieved in all patients. Of the 20 patients enrolled in the ongoing Phase 1/2 clinical trial, 17 patients were evaluable for efficacy. 71% of evaluable patients met the key efficacy endpoint of being successfully weaned off their anesthetic agents while SAGE-547 was being administered. In addition, 71% of evaluable patients were successfully weaned off SAGE-547 without recurrence of SRSE. As a group, patients who responded to SAGE-547 generally demonstrated rapid improvement over the first five days following treatment. Patients who responded also continued to improve over the 30-day follow-up period as assessed by several measures. SAGE-547 was generally well-tolerated and no drug-related serious adverse events, as determined by the Safety Review Committee, were reported in treated patients. The mean exposure level of SAGE-547 was approximately 200nM.

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At baseline, all patients were measured by the Clinical Global Impression of Severity, or CGI-S scale, which tracks patient progress and treatment response over time, as well as the Glasgow Coma Scale, which helps gauge the severity of an acute brain injury. At baseline, 19 patients were classified as most extremely ill as measured by CGI-S and the remaining patient was described as severely ill. By day 30, the group of patients who responded to SAGE-547 (12/17) had improved to mildly ill, which represents a three-step improvement in the severity of their illness. In contrast, the group of patients who did not respond to SAGE-547 (5/17) did not improve beyond severely ill throughout the study period. As measured by the Glasgow Coma Scale, the group of patients who responded to SAGE-547 showed rapid improvement in the first five days following treatment and continued improvement throughout the complete study period.

The underlying etiology was explored in the 20 patients enrolled. SRSE was attributed to brain hemorrhage in four patients, infections in four patients, worsening of seizures in two patients, primary or metastatic brain tumors in two patients and to unknown causes in three patients. The other five cases of SRSE were caused by each of the following in one case: stroke, sickle cell anemia, Lupus, posterior reversible encephalopathy syndrome and toxic ingestion.

Independent of treatment response, five patient deaths occurred within the trial period, all driven by underlying conditions. Although 13 patients (65%) reported serious adverse events, none were considered drug-related. Mean exposure levels of SAGE-547 were approximately 200 nM.

The results obtained from the patients in this trial may not be representative of results obtained from future patients treated with SAGE-547 in this clinical trial. For a further description of this risk, see Risk Factors Risks Related to Product Development, Regulatory Approval and Commercialization Positive results from early non-clinical studies and clinical trials of our product candidates are not necessarily predictive of the results of later non-clinical studies and clinical trials of our product candidates. If we cannot replicate the positive results from our earlier non-clinical studies and clinical trials of our product candidates in our later non-clinical studies and clinical trials, we may be unable to successfully develop, obtain regulatory approval for and commercialize our product candidates.

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In October 2014, the FDA approved a protocol amendment for our Phase 1/2 clinical trial that enables us to treat pediatric patients as young as two years old, to increase the dose of SAGE-547 being administered to patients and to increase the duration of treatment up to two weeks. We are continuing to enroll patients as an expansion cohort in this trial and we anticipate reporting final clinical data from this Phase 1/2 clinical trial of SAGE-547 at the Antiepileptic Drug and Device Trials XIII Conference, which is taking place May 13-15, 2015.

In addition, we continue to use SAGE-547 to explore additional potential uses of GABA_A receptor modulators in clinical trials for additional indications. In October 2014, we began patient enrollment in an exploratory Phase 2a clinical trial of SAGE-547 in patients with essential tremor, a debilitating neurological disorder that causes involuntary, rhythmic shaking with no known cause. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and activity of SAGE-547 in patients with essential tremor.

In January 2015, we also initiated a Phase 2a clinical trial of SAGE-547 in women with severe postpartum depression, or PPD, a distinct and readily identified form of major depressive disorder estimated to affect up to 20% of women following childbirth. This trial is designed to evaluate the safety, tolerability, pharmacokinetics and efficacy of SAGE-547 for the treatment of severe PPD. We plan to use the data from these exploratory trials to help guide the design of a second-generation GABA_A receptor modulator for the chronic treatment of these diseases.

Phase 3 Clinical Program Design

On April 2, 2015, we announced that at a recent End-of-Phase 2 meeting with the FDA, general agreement was reached on the design and key elements for our planned Phase 3 clinical program for SAGE-547 for the treatment of SRSE. Subject to submission and review by the FDA of a final protocol for the planned Phase 3 clinical trial and updated chemistry, manufacturing and controls information, we expect to initiate this trial in mid-2015. If successful, we believe the results from this Phase 3 clinical trial, together with other clinical data obtained from the SAGE-547 development program, could form the basis of an NDA submission for SAGE-547.

The Phase 3 clinical trial is planned as a randomized, double-blind, placebo-controlled trial designed to assess the efficacy and safety of SAGE-547 in approximately 126 evaluable patients with SRSE, aged two years or older, at up to 150 trial sites in the United States and Europe. It is expected that patients will be excluded from the trial for various reasons, including, pregnancy, instances where there is no reasonable expectation of recovery or life-expectancy, presence of certain underlying conditions, previous exposure to SAGE-547 or to an investigational medicine or device within 30 days of planned enrollment in the trial.

Patients will be randomized 1:1 to receive either SAGE-547 or placebo in addition to standard of care third-line anti-seizure agents for a total of six days, and will be followed after treatment for a period of 21 days. Based on data from the ongoing Phase 1/2 clinical trial, the trial is designed to provide 90% statistical power. The planned primary endpoint of this trial is expected to be successful resolution of SE after weaning the patient off all third-line anti-seizure agents, and SAGE-547 or placebo, without resumption of SE within 24 hours after completion of blinded SAGE-547 or placebo administration. Patients who fail to respond to initial blinded treatment with SAGE-547 or placebo may be eligible to be treated with an open-label, higher dose regimen of SAGE-547. Secondary endpoints are expected to explore the rate of recovery, regaining of consciousness, mental status and functional outcome of patients. It is expected that safety will be assessed through a review of adverse events and medications, laboratory testing, vital signs, ECG parameters and mortality.

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In conjunction with our planned Phase 3 clinical trial, we also plan to initiate an open-label, expanded access protocol designed to offer SAGE-547 to patients affected with SRSE with limited treatment options. The goal of this expanded access protocol is to allow emergency treatment of patients who are not admitted to trial sites and cannot be transferred to trial sites for medical or other reasons. The expanded access protocol will be identical to the protocol for the planned Phase 3 clinical trial for drug administration procedures, including dosing scheme and potential to retreat if patients do not respond to the initial six day course treatment. An important difference, however, is that all patients will be administered SAGE-547, there is no placebo group in this trial. While we will be collecting efficacy data from the expanded access trial, the data will be used primarily to support the safety database for our planned NDA submission. Patients treated in this protocol will be followed for two months following completion of treatment.

SAGE-689

SAGE-689 is being developed as an adjunctive IV therapy for the treatment of SE patients whose seizures have not resolved after treatment with BDZs in a non-hospital setting. Patients with SE at this stage are transported by ambulance to the hospital and frequently receive treatment in the emergency room with anti-seizure drugs. If their seizure does not resolve rapidly the patient must be transferred to the ICU and immediately placed into a medically induced coma to minimize the risk of brain damage. SAGE-689 is currently in IND-enabling toxicology and safety pharmacology testing.

We are developing SAGE-689 so that it will have what we believe is the optimal profile as a second-line therapy for the treatment of SE prior to a patient being placed into a medically induced coma. These characteristics include a wide therapeutic window to allow for modulation of the GABA_A receptor without inducing deep anesthesia, and a short half-life to permit rapid onset and loss of activity. The latter property will facilitate rapid discharge or transfer to the ICU without residual drug on board. SAGE-689 is being formulated for IV or intramuscular administration to optimize these characteristics in a clinical setting.

We plan on filing an IND for SAGE-689 by late 2015 and to begin a Phase 1 clinical trial thereafter. Our Phase 1 clinical development program for SAGE-689 will be designed to rapidly assess relevant product characteristics for this compound, such as quality of sedation, impact on EEG in normal patients and possibly in patients with epilepsy, pharmacokinetics and general safety. Our Phase 1 clinical trial will also inform us of SAGE-689's ability to induce EEG-confirmed burst suppression.

Depending upon the results of our Phase 1 clinical trial, we anticipate that subsequent development of SAGE-689 will involve studies of its utility as an adjunctive therapy, comparing it in combination with best practice, versus practice alone.

Non-clinical results

The non-clinical evaluation for SAGE-689 encompasses standard toxicology and pharmacology. In addition, our focus has been on understanding the potential for SAGE-689 to be deployed as a short half-life agent in the treatment of SE in an emergency situation. Thus we have looked at non-clinical models that will assess its sedative profile, its cardiovascular safety, its effect on EEG and its ability to induce burst suppression as a proxy for anti-seizure activity.

The results from non-clinical animal models may not be replicated in clinical trials. Many drug candidates, including many targeting CNS disorders, with promising non-clinical profiles have failed to demonstrate similar safety, non-toxicity and efficacy in humans. To the extent possible, animal models

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attempt to replicate both the phenotype of the human disease and its underlying causality to assess the putative efficacy of a drug candidate. However, in many cases, the cause of the human disease is not fully elucidated, thereby decreasing the likelihood that the animal model will accurately predict the efficacy of a drug candidate in humans. The non-clinical results reported for SAGE-689 below should be read with these limitations in mind.

Efficacy

SAGE-689, a selective positive allosteric modulator at GABA_A receptors, possesses anticonvulsant, anxiolytic and sedative properties in animal models. This activity provides a non-clinical rationale for potential efficacy of SAGE-689 in patients with various forms of seizure. In an SE model in rodents, a single IV bolus dose of SAGE-689 (5 mg/kg and 15 mg/kg) given up to 60 minutes following induction of SE produces complete cessation of seizure activity. In the same SE model, BDZs do not show effectiveness in seizure cessation. Furthermore, SAGE-689 effectively halts re-occurrence of seizure activity for up to three hours after treatment with the compound as measured by EEG. SAGE-689 produces dose-related protection from seizure activity in rodent model of SE.

With respect to sedation, SAGE-689 is an effective, fast acting and rapidly reversible sedative/hypnotic agent when given acutely in both rats and dogs. Single IV bolus injections of SAGE-689 produce a spectrum of sedative effects, from light sedation at low doses to general anesthesia at higher doses. In emergency room settings, propofol is often used as a sedative for emergency situations and for seizure control. We thus compared SAGE-689's safety and activity to propofol in several non-clinical animal experiments, and in general, SAGE-689 showed a more manageable profile for achieving varying levels of sedation. In particular, progressively deeper levels of sedation with SAGE-689 were achieved with more control and broader plasma exposures than with propofol, indicating that SAGE-689 in humans will allow, we believe, a more optimal level of controlled sedation than with propofol. Recovery from sedation after withdrawal of SAGE-689 was rapid, occurring within 15 minutes after cessation of a one hour continuous IV infusion in the rat. These recovery times are comparable with those observed after a one hour continuous infusion of propofol in animal studies.

Pharmacokinetics

SAGE-689 was found to have high systemic clearance and short half-life in both rodents and dogs. CNS penetration was observed in rats following both IV bolus and IV infusion doses, and brain to plasma ratios exceed one, showing ease of transport into the brain, which is necessary for efficacy in humans.

Non-clinical safety

Cardiovascular safety is an important attribute of any agent that is administered via IV infusion, whether used for sedation or other purposes. We compared SAGE-689 to propofol in these non-clinical studies. Safety studies were conducted in telemetered male beagle dogs administered SAGE-689. SAGE-689 exhibited less severe cardiovascular and respiratory effects than propofol across a wide range of exposures, with moderate sedation across a wide range of exposures. In addition, no apnea, or absence of breathing, was observed after IV administration of SAGE-689 in this study, and although a detailed analysis was not performed, there were no obvious, prevalent or reproducible abnormalities associated with the electrocardiogram during the administration of SAGE-689. Thus non-clinical data suggest an acceptable cardiovascular safety profile for SAGE-689 that we believe in humans has the potential to have a better risk-benefit profile than propofol.

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SAGE-217

SAGE-217 is being developed as an oral monotherapy for orphan epilepsies, such as Dravet syndrome and Rett syndrome. The chemical characteristics of SAGE-217 potentially allow formulation as both an IV and oral medication. SAGE is first intending to develop this compound as an oral agent where we believe an expedited pathway is possible. SAGE-217 is currently in IND-enabling toxicology and safety pharmacology testing.

We are developing SAGE-217 so that it will have what we believe is an optimal profile as a monotherapy for epilepsy in many forms. SAGE-217 is expected to have the ability to induce deep anesthesia and produce EEG-confirmed burst suppression. The long half-life of SAGE-217 will allow potential once-daily dosing, and will also allow it to auto-taper on cessation as well as avoiding rapid fluctuations of blood levels when administered. The long half-life, ability to induce deep and prolonged anesthesia, oral availability, and potency at extrasynaptic GABA_A receptors will distinguish it from other product candidates and other currently available therapies.

It is these target characteristics, we believe, that may make SAGE-217 useful as a treatment for orphan epilepsies, such as Dravet syndrome and Rett syndrome. In addition, potential anti-seizure activity may make SAGE-217 useful in general epilepsy and in forms of SE as an oral maintenance therapy after SE, RSE, or SRSE resolution is achieved, in order to prevent seizure recurrence.

We plan to file an IND for SAGE-217 by late 2015. In the Phase 1 development of SAGE-217, we intend to assess sedative qualities, safety profile, cardiovascular safety and impact on EEG.

Non-clinical results

The results from non-clinical animal models may not be replicated in clinical trials. Many drug candidates, including many targeting CNS disorders, with promising non-clinical profiles have failed to demonstrate similar safety, non-toxicity and efficacy in humans. To the extent possible, animal models attempt to replicate both the phenotype of the human disease and its underlying causality to assess the putative efficacy of a drug candidate. However, in many cases, the cause of the human disease is not fully elucidated, thereby decreasing the likelihood that an animal model will accurately predict the efficacy of a drug candidate in humans. The non-clinical results reported for SAGE-217 below should be read with these limitations in mind.

Efficacy

Similar to SAGE-689 and SAGE-547, SAGE-217 is pharmacologically active in various models of seizure. Efficacy studies in multiple non-clinical seizure models have shown good anti-seizure activity in pharmacoresistant models of SE. Through GABA_A receptor modulation, SAGE-217 possesses potent anticonvulsant, anxiolytic and sedative activity when administered in vivo.

In an SE model in rodents, SAGE-217 produces complete cessation of seizures at 3 mg/kg and 5 mg/kg when dosed via IV infusion. Furthermore SAGE-217 effectively halts any seizure recurrence up to three hours after the treatment when measured by EEG. Additional studies in seizure models are ongoing to fully understand the utility of SAGE-217 for other seizure indications, however, we believe the ability to prevent seizure recurrence in these models may be an attribute unique to this molecule.

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Pharmacokinetics

SAGE-217 was found to have low systemic clearance and long half-life in both rodents and dogs. CNS penetration was observed in rats following both oral and IV doses, and brain to plasma ratios exceed one, showing ease of transport into the brain, which is necessary for efficacy in humans. The pharmacokinetic profile of SAGE-217 suggests the compound will be amenable to once-a-day dosing in humans as an oral formulation, and will require only a very low infusion dose requirement when dosed via IV infusion.

Non-clinical safety

Safety studies on SAGE-217 are planned. Pharmacology studies completed suggest the molecule is well tolerated at efficacious doses demonstrating activity when administered either via IV infusion or orally.

Further Exploitation of GABA_A and NMDA Receptors

We are exploring additional potential products in a variety of CNS disorders based on modulation of both the GABA_A and NMDA receptors. In addition to our products focused on SE, other GABA_A mediated CNS disorders upon which we believe our approach can have a material impact include Rett syndrome, Dravet syndrome, fragile X syndrome, anxiety, depression, sleep disorders, mania, tremor, tinnitus and post-traumatic stress disorder.

NMDA receptors also serve a critical role in CNS related activities; however current attempts at exploiting these receptors have been suboptimal due to limited efficacy and adverse events. We have produced a large pool of highly selective product candidates which are allosteric modulators of the NMDA receptor that we believe can be used for the treatment of cognitive dysfunction in diseases such as depression, Alzheimer's disease, attention deficit hyperactivity disorder and schizophrenia, as well as certain aspects of Huntington's disease, and neuropathic pain.

Our initial focus will remain on those indications where we can independently develop and commercialize our products, if approved. However, our broad potential pipeline lessens our reliance on the success of any one program. We believe our ability to design and develop novel molecules with distinct profiles and receptor subtype selectivity will provide us with an opportunity to create value by either in-house development or by partnering these assets with third parties who possess the development and commercialization capabilities to pursue these programs.

Manufacturing and Supply

We do not own nor operate, and currently have no plans to establish, any manufacturing facilities. We currently resource all of our non-clinical and clinical compound supply through third-party contract manufacturing organizations, or CMOs.

We currently have sufficient SAGE-547 drug product on hand for our Phase 1/2 clinical trial in SRSE, Phase 2a clinical trial in essential tremor and Phase 2a clinical trial in severe PPD and ongoing non-clinical studies. Additionally, we have SAGE-547 drug substance on hand to support our planned Phase 3 clinical trial. We are working with our CMOs to modify the manufacturing process for SAGE-547 drug product to (i) increase the maximum shelf-life of this product candidate from one to two years to up to three years and (ii) eliminate the need for cold storage. We currently have sufficient SAGE-689 drug substance on hand for our ongoing non-clinical studies and have a scheduled manufacturing date for current good manufacturing practice, or cGMP, batches to support Phase 1 clinical trials. We currently have sufficient SAGE-217 drug substance on hand for our ongoing non-clinical trials and have begun manufacture of cGMP batches to support Phase 1 clinical trials.

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We have established relationships with several key CMOs to enable both the non-clinical and clinical supply chains for SAGE-547, SAGE-689 and SAGE-217 active pharmaceutical ingredient, or API, as well as drug product under cGMP protocols. Key intermediates to support the large-scale production of these candidates are performed by other CMOs on a purchase order basis. We do not currently have arrangements in place for redundant supply of bulk drug substance. It is our intent to identify and qualify additional manufacturers to provide API and fill-and-finish services prior to submission of a new drug application to the FDA for all product candidates.

SAGE-547, SAGE-689 and SAGE-217 are low molecular weight compounds isolated as stable crystalline solids. We believe the syntheses of SAGE-547, SAGE-689 and SAGE-217 are reliable and reproducible from readily available starting materials, and the synthetic routes are amenable to large-scale and do not require unusual equipment or handling in the manufacturing process. The enantiomeric purity of SAGE-547, SAGE-689 and SAGE-217 is very high (>99.9%) as a result of the chirality being derived from the backbone of the natural product steroid-based raw materials. We expect to continue to identify and develop drug candidates that are amenable to cost-effective production at contract manufacturing facilities.

Research and Development

See Management's Discussion and Analysis of Financial Condition and Results of Operations for more information regarding our research and development expenses in the years ended December 31, 2014, 2013, and 2012.

Sales and Marketing

Given our stage of development, we have not yet established a commercial organization or distribution capabilities, nor have we entered into any partnership or co-promotion arrangements with an established pharmaceutical company. We are concentrating our internal efforts on CNS disorders where we believe we can efficiently commercialize our product candidates on our own. For example, in the United States we believe that SE patients are easily identifiable in tertiary care centers where there are ICUs and staff trained in treating RSE or SRSE patients. In a recent secondary analysis of Premier Network Hospitals' 2012 billing data, we found approximately 70% of SRSE discharges, as determined by business rules we and our consultants established, occurred in 925 US hospitals of 300 beds or more. In addition, 56% of the SRSE discharges had a length of stay of 10 days or more thereby allowing hospitals to bill from approximately \$170,000 to \$550,000 in charges depending on the patient's length of stay and diagnosis-related group (DRG). As a result, we believe we can successfully launch and commercialize our initial product candidates on our own, using a small and highly specialized sales force similar to those of other rare-disease companies. We also think there may be potential pharmacoeconomic argument should hospital length of stays become reduced due to earlier resolution of SRSE although SAGE-547 will have no effect on the underlying etiology.

To develop the appropriate commercial infrastructure to launch our product candidates, we may establish alliances with one or more pharmaceutical company collaborators, depending on, among other things, the applicable indications, the related development costs and our available resources. We plan to selectively partner assets geared toward treating CNS disorders that impact large patient populations, such as depression and cognition where there are no reliable and predictive animal models to guide drug development. In order to effectively and efficiently develop product candidates for these larger markets or more difficult indications, we intend to partner at an appropriate stage with companies who have clinical expertise and pre-existing commercial infrastructure in these areas.

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Licenses

We have entered into several license agreements to support our various programs.

Washington University

In November 2013, we entered into a license agreement with Washington University, or WU. Under this agreement, and subject to certain rights of the U.S. government and rights retained by WU, WU granted to us an exclusive, worldwide license under certain patent rights to make, have made, sell and offer for sale, use and import products covered by certain of its patent rights. WU's rights in a patent application disclosing and claiming SAGE-689 is included in this license agreement. Under this agreement, WU also granted us non-exclusive license under certain technical information and tangible research information to use such technical information and/or tangible research information to make, have made, sell, offer for sale, use and import products that embody or were made using a method or process covered in the technical information and/or tangible research information. The WU license also grants us a right to sublicense our licensed rights to third parties, provided each sublicensee enters into a written agreement with us with terms consistent with our agreement with WU. We must pay to WU a percentage of the revenue we receive from sublicensing our rights under this agreement, initially in the mid-teens and decreasing to the mid-single digits over time.

Pursuant to the WU license, we are required to use commercially reasonable efforts to continue active, diligent development of licensed products and to use commercially reasonable efforts to manufacture, promote and sell licensed products throughout the territory and in the field during the term of the agreement. We must deliver written reports to WU describing our progress no later than January 31 and July 31 of the first two calendar years of the agreement, and no later than January 31 of each calendar year thereafter.

We must pay to WU an annual maintenance fee until and including the year in which our first Phase 2 clinical trial is initiated, and we must make up to \$0.7 million and \$0.5 million in clinical development and regulatory milestones, respectively, to WU, for each licensed product, upon reaching certain milestones relating to the clinical development of our product candidates. The license agreement also requires us to make low single-digit royalty payments to WU in connection with the sales of licensed products.

The WU agreement will expire on a licensed product-by-licensed product basis upon the later of (i) the last day that at least one valid patent claim covering the licensed product exists, or (ii) the tenth anniversary of the day of the first commercial sale of the licensed product. We may terminate the WU Agreement early for convenience upon providing WU with 90 days' written notice. WU may terminate this agreement early in the event of our failure to cure a material breach within the applicable cure period or our bankruptcy. In the event of early termination of this agreement before the expiration of the last to expire of the patent rights, we must immediately discontinue manufacture, sale and distribution of any licensed products.

CyDex Pharmaceuticals

In August 2013, we entered into a license agreement with CyDex Pharmaceuticals, Inc., or CyDex, a wholly owned subsidiary of Ligand Pharmaceuticals, Inc., which was amended in April 2014. Pursuant to the CyDex agreement, CyDex granted us an exclusive, non-transferable license under certain patent rights to research, develop, make, have made, import, use, offer for sale, and sell licensed products, which will consist of our compound, a certain neuroactive steroid known as allopregnanolone, formulated in CyDex's proprietary Captisol technology, in the licensed field. Captisol is an excipient that allows us to dissolve allopregnanolone, which has limited solubility in water, in an

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aqueous solution. Our field of use includes all fields for the treatment, prevention or diagnosis of any disease or symptom in humans or animals. We have also been granted a non-exclusive license to all toxicology/safety and other relevant scientific data owned, licensed or developed by CyDex and relating to Captisol for use in connection with licensed product in the licensed field. We have the right to grant sublicenses outright to third parties under the agreement, provided each sublicensee enters into a written agreement with us, and each sublicensee must abide by the restrictions of the CyDex license. Pursuant to the agreement, we granted CyDex a nonexclusive, royalty-free license to any Captisol improvements developed by us.

Pursuant to the CyDex license, we are required during the term of the agreement to use commercially reasonable efforts to continue active, diligent development of the licensed product, to seek regulatory approval of the licensed product and to commercialize the licensed product following regulatory approval. We must deliver periodic progress reports to CyDex.

We are obligated to make milestone payments of \$0.8 million and \$3.8 million, respectively, based on the achievement of clinical development and regulatory milestones for the development of SAGE-547, with the payments to be made once per field in the fields of SE and traumatic brain injury. For the development in two additional fields, we are obligated to make milestone payments, once per field, for the first two additional fields, on the achievement of clinical development and regulatory milestones of \$1.3 million and \$8.5 million, respectively. We must also pay low single-digit royalties to CyDex in connection with the sale of such licensed products. CyDex controls prosecution and enforcement of the licensed patent rights.

The CyDex license is perpetual until terminated. We may terminate the CyDex agreement for convenience upon providing 180 days prior written notice to CyDex. Either party has the right to terminate the agreement for failure to cure a material breach in the applicable cure period.

We have also entered into an amended supply agreement with CyDex pursuant to which we are required to purchase all of our supply of Captisol from CyDex and CyDex is required to supply us with Captisol, subject to certain limitations. Under this agreement, if we do not place an order for at least the quantity of Captisol we forecasted in the first quarter of any year, we will be required to pay 60% of the purchase price for the forecasted material not purchased to CyDex. CyDex has the right to raise prices for Captisol once per year based on an industry index maintained by the Bureau of Labor Statistics. The supply agreement also contains customary provisions regarding confidentiality, indemnification and non-solicitation, as well as customary representations and warranties. The supply agreement will terminate upon the termination of our license agreement with CyDex or upon breach by either party without cure after notice.

University of California

In October 2013, we entered into a license agreement with The Regents of the University of California, or the Regents, which was amended in May 2014. Pursuant to this agreement, and subject to certain rights of the U.S. government and rights retained by the Regents, the Regents granted us a non-exclusive, non-transferable license under all personal property rights of the Regents covering the tangible personal property in an IND application package owned by the Regents, or the Data, and a specified quantity of cGMP grade allopregnanolone, or the Material, to (i) use the Data for reference or incorporation in an IND for the use of the Material as a treatment of SE, essential tremor and/or severe PPD and (ii) use the Material or modifications of the Material to develop a pharmaceutical formulation for clinical trials for SE, essential tremor and/or postpartum depression. The rights licensed to us are not sublicenseable.

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Pursuant to this agreement, we are required to use commercially reasonable efforts to proceed with the development, manufacture and sale of one or more products containing allopregnanolone, a derived product under the agreement, for the treatment of SE, essential tremor and/or severe PPD depression. As of January 1, 2014, we must deliver written reports to the Regents describing our progress no later than 60 days subsequent to June 30 and December 31 of each fiscal year.

This agreement requires us to make up to \$0.1 million in milestone payments in connection with the first derived product that meets the relevant milestones and we must also pay royalties of less than 1% to the Regents for each derived product for a period of 15 years following the first commercial sale of such derived product. This agreement will terminate on the earlier to occur of (i) 27 years after the effective date or (ii) 15 years after the last-derived product is first commercially sold. We may terminate this agreement early for convenience upon providing 60 days prior written notice to the Regents. The Regents may terminate this agreement early in the event of material default, including failure to provide timely progress reports, after the applicable cure period, or in the event of our bankruptcy. In the event of early termination of this agreement, we have the right to sell any partially made derived products for a period of 120 days from the date of termination, but may not otherwise make, have made, use, sell, have sold, offer for sale or import products containing allopregnanolone.

Intellectual Property

We strive to protect the proprietary technology that we believe is important to our business, including seeking and maintaining patents intended to cover our product candidates and compositions, their methods of use and processes for their manufacture, and any other aspects of inventions that are commercially important to the development of our business. We also rely on trade secrets to protect aspects of our business that are not amenable to, or that we do not consider appropriate for, patent protection.

We plan to continue to expand our intellectual property estate by filing patent applications directed to compositions, methods of treatment and patient selection created or identified from our ongoing development of our product candidates. Our success will depend on our ability to obtain and maintain patent and other proprietary protection for commercially important technology, inventions and know-how related to our business, defend and enforce our patents, preserve the confidentiality of our trade secrets and operate without infringing the valid and enforceable patents and proprietary rights of third parties. We also rely on know-how, continuing technological innovation and in-licensing opportunities to develop and maintain our proprietary position. We seek to obtain domestic and international patent protection, and endeavor to promptly file patent applications for new commercially valuable inventions.

The patent positions of biopharmaceutical companies like us are generally uncertain and involve complex legal, scientific and factual questions. In addition, the coverage claimed in a patent application can be significantly reduced before the patent is issued, and patent scope can be reinterpreted by the courts after issuance. Moreover, many jurisdictions permit third parties to challenge issued patents in administrative proceedings, which may result in further narrowing or even cancellation of patent claims. We cannot predict whether the patent applications we are currently pursuing will issue as patents in any particular jurisdiction or whether the claims of any issued patents will provide sufficient protection from competitors.

Because patent applications in the United States and certain other jurisdictions are maintained in secrecy for 18 months or potentially even longer, and since publication of discoveries in the scientific or patent literature often lags behind actual discoveries, we cannot be certain of the priority of inventions covered by pending patent applications. Moreover, we may have to participate in interference proceedings or derivation proceedings declared by the United States Patent and Trademark Office, or U.S. PTO, to determine priority of invention.

Table of Contents**Patents**

Our patent portfolio includes patent applications in the early stages of prosecution and no patents have, as of yet, issued from our patent application estate. These patent applications fall into three categories: (1) SAGE-547; (2) GABA_A receptor modulators; including genus and species claims to SAGE-689; and (3) NMDA receptor modulators.

- (1) We own three patent families generally related to SAGE-547. One of these patent families includes a patent application having claims to compositions containing allopregnanolone and a cyclodextrin. The compositions can be used for the treatment of CNS disorders such as traumatic brain injury and SE. The second patent family includes patent applications having claims directed to methods of treating seizure disorders, such as SE, by administering allopregnanolone using particular dosing regimens or multiple dosage phases. The third patent family includes methods of treating essential tremor and depression such as severe PPD. Any U.S. patents that may issue from these families of patent applications would have a statutory expiration date in January and August of 2033, and September 2035 respectively. The time period for electing to pursue foreign patent protection for the inventions disclosed in the third patent family by filing national stage patent applications in individual jurisdictions has not yet expired, and we will need to decide whether and where to pursue ex-U.S. protection before expiration of the applicable deadlines.
- (2) We have exclusively licensed a portfolio of patent applications owned by WU, which are directed to certain GABA receptor modulating compounds and methods of using these compounds, for example in anesthesia or treatment of GABA-related disorders. This portfolio of patent applications includes seven families of patent applications. One of these seven families of patent applications is co-owned by us, and this co-owned family includes a pending U.S. patent application and a pending Patent Cooperation Treaty patent application. This co-owned application discloses and claims SAGE-689 and its use in anesthesia or treatment of GABA-related disorders. Any U.S. patents that may issue from the SAGE-689 patent family would have a statutory expiration date of December 2033. The time period for electing to pursue foreign patent protection for SAGE-689 by filing national stage patent applications in individual jurisdictions has not yet expired, and we will need to decide whether and where to pursue ex-U.S. protection before expiration of the applicable deadlines. In addition, U.S. 7,781,421, solely owned by WU, expires in September 2027. Any patents that may issue, if any, from the remaining five families solely owned by WU would have statutory expiration dates that range from 2032 to 2034.
- (3) In addition to the patent applications licensed from WU, we own thirteen patent families, resulting from work done exclusively by us and our contract research organizations, directed to additional GABA receptor modulating compounds and methods of using these compounds, for example in anesthesia or treatment of GABA-related disorders. Any U.S. patents that may issue from these patent families would have a statutory expiration ranging from October 2032 to January 2036. Other than SAGE-547 and SAGE-689, we have pending within these patent families genus and species claims to the majority of the compounds in our GABA_A receptor modulating compound collection, including SAGE-217. These patent families are in the early stages of patent prosecution and include families for which only provisional applications have been filed. The time period for electing to pursue foreign patent protection by filing national stage patent applications in individual jurisdictions has not yet expired for some of these patent families, and we will need to decide whether and where to pursue ex-U.S. protection before expiration of the applicable deadlines.
- (4) We also own six families of applications directed to modulators of NMDA receptors. Four of these patent families are directed to compounds that modulate NMDA receptors, which can be used to treat NMDA receptor-related disorders such as CNS related conditions. One of

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these patent families is directed to using a naturally occurring compound as a biomarker for a subject who would benefit from treatment with a modulator of NMDA receptors. One of these patent families is directed to using a modulator of NMDA receptors to treat a rare NMDA loss of function disorder. Any patents that may issue, if any, from these families of applications directed to modulators of NMDA receptors would have statutory expiration dates in September 2032 and October 2035.

Patent term

The base term of a U.S. patent is 20 years from the filing date of the earliest-filed non-provisional patent application from which the patent claims priority. The term of a U.S. patent can be lengthened by patent term adjustment, which compensates the owner of the patent for administrative delays at the U.S. PTO. In some cases, the term of a U.S. patent is shortened by terminal disclaimer that reduces its term to that of an earlier-expiring patent.

The term of a U.S. patent may be eligible for patent term extension under the Drug Price Competition and Patent Term Restoration Act of 1984, referred to as the Hatch-Waxman Act, to account for at least some of the time the drug is under development and regulatory review after the patent is granted. With regard to a drug for which FDA approval is the first permitted marketing of the active ingredient, the Hatch-Waxman Act allows for extension of the term of one U.S. patent that includes at least one claim covering the composition of matter of an FDA-approved drug, an FDA-approved method of treatment using the drug, and/or a method of manufacturing the FDA-approved drug. The extended patent term cannot exceed the shorter of five years beyond the non-extended expiration of the patent or 14 years from the date of the FDA approval of the drug. Some foreign jurisdictions, including Europe and Japan, have analogous patent term extension provisions, which allow for extension of the term of a patent that covers a drug approved by the applicable foreign regulatory agency. In the future, if and when our pharmaceutical products receive FDA approval, we expect to apply for patent term extension on patents covering those products, their methods of use, and/or methods of manufacture.

Trade secrets

In addition to patents, we rely on trade secrets and know-how to develop and maintain our competitive position. We typically rely on trade secrets to protect aspects of our business that are not amenable to, or that we do not consider appropriate for, patent protection. We protect trade secrets and know-how by establishing confidentiality agreements and invention assignment agreements with our employees, consultants, scientific advisors, contractors and partners. These agreements provide that all confidential information developed or made known during the course of an individual or entity's relationship with us must be kept confidential during and after the relationship. These agreements also provide that all inventions resulting from work performed for us or relating to our business and conceived or completed during the period of employment or assignment, as applicable, shall be our exclusive property. In addition, we take other appropriate precautions, such as physical and technological security measures, to guard against misappropriation of our proprietary information by third parties.

Competition

The biopharmaceuticals industry is highly competitive. There are many public and private biopharmaceutical companies, universities, governmental agencies and other research organizations actively engaged in the research and development of products that may be similar to our product candidates or address similar markets. It is probable that the number of companies seeking to develop products and therapies similar to our products will increase.

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Currently, there are no therapies that have been specifically approved for treatment of RSE or SRSE. However, many products approved for other indications, for example, general anesthetics and anti-seizure drugs, are used off-label for various stages of SE therapy. Additionally, though not indicated, acupuncture, hypothermia, and electroconvulsive therapy are sometimes used prior to withdrawal of care for patients with SRSE.

In the field of neuroactive steroids focused on modulation of GABA_A or NMDA receptors, our principal competitor is Marinus Pharmaceuticals, Inc., which we believe is developing a reformulated form of ganaxolone, a known GABA_A positive allosteric modulator neuroactive steroid, for the potential treatment of drug-resistant partial complex seizures and fragile X syndrome. With the development of SAGE's GABA platform, we see this as a prime opportunity to progress the second generation of neuroactive steroids which are pharmacologically active with fewer and less severe off-target effects than first generation compounds. Many of our potential competitors, alone or with their strategic partners, have substantially greater financial, technical and human resources than we do and significantly greater experience in the discovery and development of product candidates, obtaining FDA and other regulatory approvals of treatments and the commercialization of those treatments. Mergers and acquisitions in the biotechnology and pharmaceutical industries may result in even more resources being concentrated among a smaller number of our competitors. Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize products that are safer, more effective, have fewer or less severe side effects, are more convenient or are less expensive than any products that we may develop. Our competitors also may obtain FDA or other regulatory approval for their products more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market.

Government Regulation

Government authorities in the United States at the federal, state and local level and in other countries extensively regulate, among other things, the research, development, testing, manufacture, quality control, approval, labeling, packaging, storage, record-keeping, promotion, advertising, distribution, post-approval monitoring and reporting, marketing and export and import of drug products. Generally, before a new drug can be marketed, considerable data demonstrating its quality, safety and efficacy must be obtained, organized into a format specific to each regulatory authority, submitted for review and approved by the regulatory authority.

U.S. drug development

In the United States, the FDA regulates drugs under the Federal Food, Drug, and Cosmetic Act, or FDCA, and its implementing regulations. Drugs are also subject to other federal, state and local statutes and regulations. The process of obtaining regulatory approvals and the subsequent compliance with appropriate federal, state, local and foreign statutes and regulations require the expenditure of substantial time and financial resources. Failure to comply with the applicable U.S. requirements at any time during the product development process, approval process or after approval, may subject an applicant to administrative or judicial sanctions. These sanctions could include, among other actions, the FDA's refusal to approve pending applications, withdrawal of an approval, a clinical hold, warning letters, product recalls or withdrawals from the market, product seizures, total or partial suspension of production or distribution injunctions, fines, refusals of government contracts, restitution, disgorgement, or civil or criminal penalties. Any agency or judicial enforcement action could have a material adverse effect on us.

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Our product candidates must be approved by the FDA through the NDA process before they may be legally marketed in the United States. The process required by the FDA before a drug may be marketed in the United States generally involves the following:

Completion of extensive non-clinical, sometimes referred to as non-clinical laboratory tests, non-clinical animal studies and formulation studies in accordance with applicable regulations, including the FDA's current Good Laboratory Practice, or GLP, regulations;

Submission to the FDA of an IND application, which must become effective before human clinical trials may begin;

Approval by an independent institutional review board, or IRB, or ethics committee at each clinical trial site before each trial may be initiated;

Performance of adequate and well-controlled human clinical trials in accordance with applicable IND and other clinical trial-related regulations, sometimes referred to as good clinical practices, or GCPs, to establish the safety and efficacy of the proposed drug for each proposed indication;

Submission to the FDA of an NDA, for a new drug;

A determination by the FDA within 60 days of its receipt of an NDA to file the NDA for review;

Satisfactory completion of an FDA pre-approval inspection of the manufacturing facility or facilities where the drug is produced to assess compliance with cGMP requirements to assure that the facilities, methods and controls are adequate to preserve the drug's identity, strength, quality and purity;

Potential FDA audit of the non-clinical and/or clinical trial sites that generated the data in support of the NDA; and

FDA review and approval of the NDA, including consideration of the views of any FDA advisory committee, prior to any commercial marketing or sale of the drug in the United States.

The non-clinical and clinical testing and approval process requires substantial time, effort and financial resources, and we cannot be certain that any approvals for our product candidates will be granted on a timely basis, if at all. Non-clinical tests include laboratory evaluation of product chemistry, formulation, stability and toxicity, as well as animal studies to assess the characteristics and potential safety and efficacy of the product.

The data required to support an NDA is generated in two distinct development stages: non-clinical and clinical. For new chemical entities, the non-clinical development stage generally involves synthesizing the active component, developing the formulation and determining the manufacturing process, as well as carrying out non-human toxicology, pharmacology and drug metabolism studies in the laboratory, which support subsequent clinical testing. The conduct of the non-clinical tests must comply with federal regulations, including GLPs. The sponsor must submit the results of the non-clinical tests, together with manufacturing information, analytical data, any available clinical data or literature and a proposed clinical protocol, to the FDA as part of the IND. An IND is a request for authorization from the FDA to administer an investigational drug product to humans. Some non-clinical testing may continue even after the IND is submitted, but an IND must become effective before human clinical trials may begin. The central focus of an IND submission is on the general investigational plan and the protocol(s) for human trials. The IND automatically becomes effective 30 days after receipt by the FDA, unless the FDA raises concerns or questions regarding the proposed clinical trials, including subjects will be exposed to unreasonable health risks, and places the IND on clinical hold within that 30-day time period. In such a case, the IND sponsor and the FDA must resolve any outstanding concerns before the clinical trial

can begin. The FDA may also impose clinical holds on a drug

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candidate at any time before or during clinical trials due to safety concerns or non-compliance. Accordingly, we cannot be sure that submission of an IND will result in the FDA allowing clinical trials to begin, or that, once begun, issues will not arise that could cause the trial to be suspended or terminated.

The clinical stage of development involves the administration of the drug candidate to healthy volunteers or patients under the supervision of qualified investigators, generally physicians not employed by or under the trial sponsor's control, in accordance with GCPs, which include the requirement that all research subjects provide their informed consent for their participation in any clinical trial. Clinical trials are conducted under protocols detailing, among other things, the objectives of the clinical trial, dosing procedures, subject selection and exclusion criteria, and the parameters to be used to monitor subject safety and assess efficacy. Each protocol, and any subsequent amendments to the protocol, must be submitted to the FDA as part of the IND. Further, each clinical trial must be reviewed and approved by an independent institutional review board, or IRB, at or servicing each institution at which the clinical trial will be conducted. An IRB is charged with protecting the welfare and rights of trial participants and considers such items as whether the risks to individuals participating in the clinical trials are minimized and are reasonable in relation to anticipated benefits. The IRB also approves the informed consent form that must be provided to each clinical trial subject or his or her legal representative and must monitor the clinical trial until completed. There are also requirements governing the reporting of ongoing clinical trials and completed clinical trial results to public registries.

A sponsor who wishes to conduct a clinical trial outside the United States may, but need not, obtain FDA authorization to conduct the clinical trial under an IND. If a foreign clinical trial is not conducted under an IND, the sponsor may submit data from the clinical trial to the FDA in support of an NDA so long as the clinical trial is conducted in compliance with an international guideline for the ethical conduct of clinical research known as the Declaration of Helsinki and/or the laws and regulations of the country or countries in which the clinical trial is performed, whichever provides the greater protection to the participants in the clinical trial.

Clinical trials

Clinical trials are generally conducted in three sequential phases that may overlap, known as Phase 1, Phase 2 and Phase 3 clinical trials.

Phase 1 clinical trials generally involve a small number of healthy volunteers who are initially exposed to a single dose and then multiple doses of the product candidate. The primary purpose of these clinical trials is to assess the metabolism, pharmacologic action, side effect tolerability and safety of the drug.

Phase 2 clinical trials typically involve studies in disease-affected patients to determine the dose required to produce the desired benefits. At the same time, safety and further pharmacokinetic and pharmacodynamic information is collected, as well as identification of possible adverse effects and safety risks and preliminary evaluation of efficacy.

Phase 3 clinical trials generally involve large numbers of patients at multiple sites (from several hundred to several thousand subjects) and are designed to provide the data necessary to demonstrate the effectiveness of the product for its intended use, its safety in use, and to establish the overall benefit/risk relationship of the product and provide an adequate basis for product approval. Phase 3 clinical trials may include comparisons with placebo and/or other comparator treatments. The duration of treatment is often extended to mimic the actual use of a product during marketing.

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Post-approval trials, sometimes referred to as Phase 4 clinical trials, may be conducted after initial marketing approval. These trials are used to gain additional experience from the treatment of patients in the intended therapeutic indication. In certain instances, FDA may mandate the performance of Phase 4 clinical trials as a condition of approval of an NDA.

Progress reports detailing the results of the clinical trials must be submitted at least annually to the FDA and written IND safety reports must be submitted to the FDA and the investigators for serious and unexpected suspected adverse events, finding from other studies, or any finding from animal or in vitro testing that suggests a significant risk for human subjects. Phase 1, Phase 2 and Phase 3 clinical trials may not be completed successfully within any specified period, if at all. The FDA, the IRB, or the sponsor may suspend or terminate a clinical trial at any time on various grounds, including a finding that the research subjects or patients are being exposed to an unacceptable health risk. Similarly, an IRB can suspend or terminate approval of a clinical trial at its institution if the clinical trial is not being conducted in accordance with the IRB's requirements or if the drug has been associated with unexpected serious harm to patients. Additionally, some clinical trials are overseen by an independent group of qualified experts organized by the clinical trial sponsor, known as a data safety monitoring board or committee. This group provides authorization for whether or not a trial may move forward at designated check points based on access to certain data from the trial. Concurrent with clinical trials, companies usually complete additional animal studies and must also develop additional information about the chemistry and physical characteristics of the drug as well as finalize a process for manufacturing the product in commercial quantities in accordance with cGMP requirements. The manufacturing process must be capable of consistently producing quality batches of the drug candidate and, among other things; we must develop methods for testing the identity, strength, quality and purity of the final drug product. Additionally, appropriate packaging must be selected and tested and stability studies must be conducted to demonstrate that the drug candidate does not undergo unacceptable deterioration over its shelf life.

NDA and FDA review process

The results of non-clinical studies and of the clinical trials, together with other detailed information, including extensive manufacturing information and information on the composition of the drug and proposed labeling, are submitted to the FDA in the form of an NDA requesting approval to market the drug for one or more specified indications. The FDA reviews an NDA to determine, among other things, whether a drug is safe and effective for its intended use and whether the product is being manufactured in accordance with cGMP to assure and preserve the product's identity, strength, quality and purity. FDA approval of an NDA must be obtained before a drug may be offered for sale in the United States.

In addition, under the Pediatric Research Equity Act, or PREA, an NDA or supplement to an NDA must contain data to assess the safety and efficacy of the drug for the claimed indications in all relevant pediatric subpopulations and to support dosing and administration for each pediatric subpopulation for which the product is safe and effective. The FDA may grant deferrals for submission of pediatric data or full or partial waivers.

Under the Prescription Drug User Fee Act, or PDUFA, as amended, each NDA must be accompanied by a user fee. The FDA adjusts the PDUFA user fees on an annual basis. According to the FDA's fee schedule, effective through December 31, 2015, the user fee for an application requiring clinical data, such as an NDA, is \$2.3 million. PDUFA also imposes an annual product fee for human drugs of \$0.1 million and an annual establishment fee of \$0.6 million on facilities used to manufacture prescription drugs. Fee waivers or reductions are available in certain circumstances, including a waiver of the application fee for the first application filed by a small business. Additionally, no user fees are assessed on NDAs for products designated as orphan drugs, unless the product also includes a non-orphan indication.

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The FDA reviews all NDAs submitted before it accepts them for filing and may request additional information rather than accepting an NDA for filing. The FDA must make a decision on accepting an NDA for filing within 60 days of receipt. Once the submission is accepted for filing, the FDA begins an in-depth review of the NDA. Under the goals and policies agreed to by the FDA under PDUFA, the FDA has 10 months from the filing date in which to complete its initial review of a standard NDA and respond to the applicant, and six months from the filing date for a priority NDA. The FDA does not always meet its PDUFA goal dates for standard and priority NDAs, and the review process is often significantly extended by FDA requests for additional information or clarification.

After the NDA submission is accepted for filing, the FDA reviews the NDA to determine, among other things, whether the proposed product is safe and effective for its intended use, and whether the product is being manufactured in accordance with cGMP to assure and preserve the product's identity, strength, quality and purity. Before approving an NDA, the FDA will conduct a pre-approval inspection of the manufacturing facilities for the new product to determine whether they comply with cGMPs. The FDA will not approve the product unless it determines that the manufacturing processes and facilities are in compliance with cGMP requirements and adequate to assure consistent production of the product within required specifications. In addition, before approving an NDA, the FDA may also audit data from clinical trials to ensure compliance with GCP requirements. Additionally, the FDA may refer applications for novel drug products or drug products which present difficult questions of safety or efficacy to an advisory committee, typically a panel that includes clinicians and other experts, for review, evaluation and a recommendation as to whether the application should be approved and under what conditions. The FDA is not bound by the recommendations of an advisory committee, but it considers such recommendations carefully when making decisions. The FDA will likely re-analyze the clinical trial data, which could result in extensive discussions between the FDA and the applicant during the review process. The review and evaluation of an NDA by the FDA is extensive and time consuming and may take longer than originally planned to complete, and we may not receive a timely approval, if at all.

After the FDA evaluates an NDA, it may issue an approval letter or a Complete Response Letter. An approval letter authorizes commercial marketing of the drug with specific prescribing information for specific indications. A Complete Response Letter indicates that the review cycle of the application is complete and the application is not ready for approval. A Complete Response Letter usually describes all of the specific deficiencies in the NDA identified by the FDA. The Complete Response Letter may require additional clinical data and/or an additional pivotal Phase 3 clinical trial(s), and/or other significant and time-consuming requirements related to clinical trials, non-clinical studies or manufacturing. If a Complete Response Letter is issued, the applicant may either resubmit the NDA, addressing all of the deficiencies identified in the letter, or withdraw the application. Even if such data and information is submitted, the FDA may ultimately decide that the NDA does not satisfy the criteria for approval. Data obtained from clinical trials are not always conclusive and the FDA may interpret data differently than we interpret the same data.

There is no assurance that the FDA will ultimately approve a drug product for marketing in the United States and we may encounter significant difficulties or costs during the review process. If a product receives marketing approval, the approval may be significantly limited to specific diseases and dosages or the indications for use may otherwise be limited, which could restrict the commercial value of the product. Further, the FDA may require that certain contraindications, warnings or precautions be included in the product labeling or may condition the approval of the NDA on other changes to the proposed labeling, development of adequate controls and specifications, or a commitment to conduct post-marketing testing or clinical trials and surveillance to monitor the effects of approved products. For example, the FDA may require Phase 4 testing which involves clinical trials designed to further assess a drug's safety and efficacy and may require testing and surveillance programs to monitor the safety of approved products that have been commercialized. The FDA may also place other conditions on

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approvals including the requirement for a risk evaluation and mitigation strategy, or REMS, to assure the safe use of the drug. If the FDA concludes a REMS is needed, the sponsor of the NDA must submit a proposed REMS. The FDA will not approve the NDA without an approved REMS, if required. A REMS could include medication guides, physician communication plans, or elements to assure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. Any of these limitations on approval or marketing could restrict the commercial promotion, distribution, prescription or dispensing of products. Product approvals may be withdrawn for non-compliance with regulatory requirements or if problems occur following initial marketing.

Orphan drug designation

Under the Orphan Drug Act, the FDA may grant orphan designation to a drug product intended to treat a rare disease or condition, which is generally a disease or condition that affects fewer than 200,000 individuals in the United States, or more than 200,000 individuals in the United States and for which there is no reasonable expectation that the cost of developing and making a drug product available in the United States for this type of disease or condition will be recovered from sales of the product. Orphan product designation must be requested before submitting an NDA. After the FDA grants orphan product designation, the identity of the therapeutic agent and its potential orphan use are disclosed publicly by the FDA. Orphan product designation does not convey any advantage in or shorten the duration of the regulatory review and approval process.

If a product that has orphan designation subsequently receives the first FDA approval for the disease or condition for which it has such designation, the product is entitled to orphan product exclusivity, which means that the FDA may not approve any other applications to market the same drug for the same indication for seven years, except in limited circumstances, such as a showing of clinical superiority to the product with orphan exclusivity. Competitors, however, may receive approval of different products for the indication for which the orphan product has exclusivity or obtain approval for the same product but for a different indication than that for which the orphan product has exclusivity. Orphan product exclusivity also could block the approval of one of our products for seven years if a competitor obtains approval of the same product as defined by the FDA or if our product candidate is determined to be contained within the competitor's product for the same indication or disease. If a drug designated as an orphan product receives marketing approval for an indication broader than what is designated, it may not be entitled to orphan product exclusivity. Orphan drug status in the European Union has similar, but not identical, benefits.

Expedited development and review programs

The FDA has a Fast Track program that is intended to expedite or facilitate the process for reviewing new drugs that meet certain criteria. Specifically, new drugs are eligible for Fast Track designation if they are intended to treat a serious or life-threatening condition and demonstrate the potential to address unmet medical needs for the condition. Fast Track designation applies to the combination of the product and the specific indication for which it is being studied. The sponsor of a new drug or biologic may request the FDA to designate the drug as a Fast Track product at any time during the clinical development of the product. Unique to a Fast Track product, the FDA may review sections of the marketing application on a rolling basis before the complete NDA is submitted, if the sponsor provides a schedule for the submission of the sections of the application, the FDA agrees to accept sections of the application and determines that the schedule is acceptable, and the sponsor pays any required user fees upon submission of the first section of the application.

Any product submitted to the FDA for marketing, including under a Fast Track program, may be eligible for other types of FDA programs intended to expedite development and review, such as priority review and accelerated approval. Any product is eligible for priority review if it has the potential to

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provide safe and effective therapy where no satisfactory alternative therapy exists or offers a significant improvement in the treatment, diagnosis or prevention of a disease compared to marketed products. The FDA will attempt to direct additional resources to the evaluation of an application for a new drug designated for priority review in an effort to facilitate the review. A product may also be eligible for accelerated approval. Drugs studied for their safety and efficacy in treating serious or life-threatening illnesses and that provide meaningful therapeutic benefit over existing treatments may receive accelerated approval, which means that they may be approved on the basis of adequate and well-controlled clinical trials establishing that the product has an effect on a surrogate endpoint that is reasonably likely to predict a clinical benefit, or on the basis of an effect on a clinical endpoint other than survival or irreversible morbidity. As a condition of approval, the FDA may require that a sponsor of a drug receiving accelerated approval perform adequate and well-controlled post-marketing clinical trials. If the FDA concludes that a drug shown to be effective can be safely used only if distribution or use is restricted, it will require such post-marketing restrictions, as it deems necessary to assure safe use of the drug, such as:

distribution restricted to certain facilities or physicians with special training or experience; or

distribution conditioned on the performance of specified medical procedures.

The limitations imposed would be commensurate with the specific safety concerns presented by the drug. In addition, the FDA currently requires as a condition for accelerated approval pre-approval of promotional materials, which could adversely impact the timing of the commercial launch of the product. Additionally, a drug may be eligible for designation as a breakthrough therapy if the drug is intended, alone or in combination with one or more other drugs, to treat a serious or life-threatening disease or condition and preliminary clinical evidence indicates that the drug may demonstrate substantial improvement over existing therapies on one or more indications. The benefits of breakthrough therapy designation includes the same benefits as fast track designation, plus intensive guidance from FDA to ensure an efficient drug development program. Fast Track designation, priority review, accelerated approval and breakthrough designation do not change the standards for approval but may expedite the development or approval process.

Pediatric trials

The Food and Drug Administration Safety and Innovation Act, or FDASIA, which was signed into law on July 9, 2012, amended the FDCA to require that a sponsor who is planning to submit a marketing application for a drug that includes a new active ingredient, new indication, new dosage form, new dosing regimen or new route of administration submit an initial Pediatric Study Plan, or PSP, within sixty days of an end-of-Phase 2 meeting or as may be agreed between the sponsor and FDA. The initial PSP must include an outline of the pediatric study or studies that the sponsor plans to conduct, including study objectives and design, age groups, relevant endpoints and statistical approach, or a justification for not including such detailed information, and any request for a deferral of pediatric assessments or a full or partial waiver of the requirement to provide data from pediatric studies along with supporting information. FDA and the sponsor must reach agreement on the PSP. A sponsor can submit amendments to an agreed-upon initial PSP at any time if changes to the pediatric plan need to be considered based on data collected from non-clinical studies, early phase clinical trials, and/or other clinical development programs.

Post-marketing requirements

Following approval of a new product, a pharmaceutical company and the approved product are subject to continuing regulation by the FDA, including, among other things, monitoring and recordkeeping activities, reporting to the applicable regulatory authorities of adverse experiences with the product, providing the regulatory authorities with updated safety and efficacy information, product

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sampling and distribution requirements, and complying with promotion and advertising requirements, which include, among others, standards for direct-to-consumer advertising, restrictions on promoting drugs for uses or in patient populations that are not described in the drug's approved labeling (known as off-label use), limitations on industry-sponsored scientific and educational activities, and requirements for promotional activities involving the Internet. Although physicians may prescribe legally available drugs for off-label uses, manufacturers may not market or promote such off-label uses. Prescription drug promotional materials must be submitted to the FDA in conjunction with their first use. Further, if there are any modifications to the drug, including changes in indications, labeling, or manufacturing processes or facilities, the applicant may be required to submit and obtain FDA approval of a new NDA or NDA supplement, which may require the applicant to develop additional data or conduct additional non-clinical studies and clinical trials. As with new NDAs, the review process is often significantly extended by FDA requests for additional information or clarification. Any distribution of prescription drug products and pharmaceutical samples must comply with the U.S. Prescription Drug Marketing Act, or the PDMA, a part of the FDCA.

In the United States, once a product is approved, its manufacture is subject to comprehensive and continuing regulation by the FDA. The FDA regulations require that products be manufactured in specific approved facilities and in accordance with cGMP. We rely, and expect to continue to rely, on third parties for the production of clinical and commercial quantities of our products in accordance with cGMP regulations. NDA holders using contract manufacturers, laboratories or packagers are responsible for the selection and monitoring of qualified firms, and, in certain circumstances, qualified suppliers to these firms. These manufacturers must comply with cGMP regulations that require among other things, quality control and quality assurance as well as the corresponding maintenance of records and documentation and the obligation to investigate and correct any deviations from cGMP. Drug manufacturers and other entities involved in the manufacture and distribution of approved drugs are required to register their establishments with the FDA and certain state agencies, and are subject to periodic unannounced inspections by the FDA and certain state agencies for compliance with cGMP and other laws. Accordingly, manufacturers must continue to expend time, money, and effort in the area of production and quality control to maintain cGMP compliance. The discovery of violative conditions, including failure to conform to cGMP, could result in enforcement actions that interrupt the operation of any such facilities or the ability to distribute products manufactured, processed or tested by them. Discovery of problems with a product after approval may result in restrictions on a product, manufacturer, or holder of an approved NDA, including, among other things, recall or withdrawal of the product from the market.

Discovery of previously unknown problems with a product or the failure to comply with applicable FDA requirements can have negative consequences, including adverse publicity, judicial or administrative enforcement, warning letters from the FDA, mandated corrective advertising or communications with doctors, and civil or criminal penalties, among others. Newly discovered or developed safety or effectiveness data may require changes to a product's approved labeling, including the addition of new warnings and contraindications, and also may require the implementation of other risk management measures. Also, new government requirements, including those resulting from new legislation, may be established, or the FDA's policies may change, which could delay or prevent regulatory approval of our products under development.

Other regulatory matters

Manufacturing, sales, promotion and other activities following product approval are also subject to regulation by numerous regulatory authorities in addition to the FDA, including, in the United States, the Centers for Medicare & Medicaid Services, other divisions of the Department of Health and Human Services, the United States Department of Justice, the Drug Enforcement Administration, the Consumer Product Safety Commission, the Federal Trade Commission, the Occupational Safety &

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Health Administration, the Environmental Protection Agency and state and local governments. In the United States, sales, marketing and scientific/educational programs must also comply with state and federal fraud and abuse laws. These laws include the federal Anti-Kickback Statute, which makes it illegal for any person, including a prescription drug manufacturer (or a party acting on its behalf) to knowingly and willfully solicit, receive, offer, or pay any remuneration that is intended to induce the referral of business, including the purchase, order, or prescription of a particular drug, for which payment may be made under a federal healthcare program, such as Medicare or Medicaid. Violations of this law are punishable by up to five years in prison, criminal fines, administrative civil money penalties, and exclusion from participation in federal healthcare programs. In addition, the Patient Protection and Affordable Health Care Act, as amended by the Health Care and Education Reconciliation Act of 2010, or collectively the ACA, among other things, amends the intent requirement of the federal Anti-Kickback Statute and criminal healthcare fraud statutes created by the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA. A person or entity no longer needs to have actual knowledge of the statute or specific intent to violate it. Moreover, the ACA provides that the government may assert that a claim including items or services resulting from a violation of the federal anti-kickback statute constitutes a false or fraudulent claim for purposes of the False Claims Act.

Although we would not submit claims directly to payors, drug manufacturers can be held liable under the federal False Claims Act, which prohibits anyone from knowingly presenting, or causing to be presented, for payment to federal programs (including Medicare and Medicaid) claims for items or services, including drugs, that are false or fraudulent, claims for items or services not provided as claimed, or claims for medically unnecessary items or services. The government may deem manufacturers to have caused the submission of false or fraudulent claims by, for example, providing inaccurate billing or coding information to customers or promoting a product off-label. In addition, our future activities relating to the reporting of wholesaler or estimated retail prices for our products, the reporting of prices used to calculate Medicaid rebate information and other information affecting federal, state, and third-party reimbursement for our products, and the sale and marketing of our products, are subject to scrutiny under this law. Penalties for a False Claims Act violation include three times the actual damages sustained by the government, plus mandatory civil penalties of between \$5,500 and \$11,000 for each separate false claim, the potential for exclusion from participation in federal healthcare programs, and, although the federal False Claims Act is a civil statute, conduct that results in a False Claims Act violation may also implicate various federal criminal statutes. If the government were to allege that we were, or convict us of, violating these false claims laws, we could be subject to a substantial fine and may suffer a decline in our stock price. In addition, private individuals have the ability to bring actions under the federal False Claims Act and certain states have enacted laws modeled after the federal False Claims Act.

Pricing and rebate programs must comply with the Medicaid rebate requirements of the U.S. Omnibus Budget Reconciliation Act of 1990 and more recent requirements in ACA. If products are made available to authorized users of the Federal Supply Schedule of the General Services Administration, additional laws and requirements apply. The handling of any controlled substances must comply with the U.S. Controlled Substances Act and Controlled Substances Import and Export Act. Products must meet applicable child-resistant packaging requirements under the U.S. Poison Prevention Packaging Act. Manufacturing, sales, promotion and other activities are also potentially subject to federal and state consumer protection and unfair competition laws.

The distribution of pharmaceutical products is subject to additional requirements and regulations, including extensive record-keeping, licensing, storage and security requirements intended to prevent the unauthorized sale of pharmaceutical products.

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The failure to comply with any of these laws or regulatory requirements subjects firms to possible legal or regulatory action. Depending on the circumstances, failure to meet applicable regulatory requirements can result in criminal prosecution, fines or other penalties, injunctions, recall or seizure of products, total or partial suspension of production, denial or withdrawal of product approvals, or refusal to allow a firm to enter into supply contracts, including government contracts. Any action against us for violation of these laws, even if we successfully defend against it, could cause us to incur significant legal expenses and divert our management's attention from the operation of our business. Prohibitions or restrictions on sales or withdrawal of future products marketed by us could materially affect our business in an adverse way.

Changes in regulations, statutes or the interpretation of existing regulations could impact our business in the future by requiring, for example: (i) changes to our manufacturing arrangements; (ii) additions or modifications to product labeling; (iii) the recall or discontinuation of our products; or (iv) additional record-keeping requirements. If any such changes were to be imposed, they could adversely affect the operation of our business.

DEA Regulation

SAGE-547 will, and our other product candidates may, if approved, be regulated as controlled substances as defined in the Controlled Substances Act of 1970, or CSA, and the U.S. Drug Enforcement Agency, or DEA, implementing regulations, which establish registration, security, recordkeeping, reporting, storage, distribution, importation, exportation, inventory, quota and other requirements administered by the DEA. These requirements are directly applicable to us and also applicable to our CMOs and to distributors, prescribers and dispensers of our product candidates. The DEA regulates the handling of controlled substances through a closed chain of distribution. This control extends to the equipment and raw materials used in their manufacture and packaging in order to prevent loss and diversion into illicit channels of commerce.

The DEA regulates controlled substances as Schedule I, II, III, IV or V substances. Schedule I substances by definition have no established medicinal use, and may not be marketed or sold in the United States. A pharmaceutical product may be listed as Schedule II, III, IV or V.

We expect that SAGE-547 will be, and our other product candidates may be, listed by the DEA as Schedule IV controlled substances under the CSA. Consequently, the importation of APIs for our product candidates, as well as the manufacture, shipping, storage, sales and use of the products, will be subject to a high degree of regulation. Also, distribution and dispensing of these drugs are highly regulated.

Annual registration is required for any facility that manufactures, distributes, dispenses, imports or exports any controlled substance. The registration is specific to the particular location, activity and controlled substance schedule. For example, separate registrations are needed for import and manufacturing, and each registration will specify which schedules of controlled substances are authorized. Similarly, separate registrations are also required for separate facilities.

The DEA typically inspects a facility to review its security measures prior to issuing a registration and on a periodic basis. Reports must also be made for thefts or losses of any controlled substance, and to obtain authorization to destroy any controlled substance. In addition, special permits and notification requirements apply to imports and exports of narcotic drugs.

The DEA establishes annually an aggregate quota for how much of a controlled substance may be produced in total in the United States based on the DEA's estimate of the quantity needed to meet legitimate scientific and medicinal needs. The DEA may adjust aggregate production quotas and

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individual production and procurement quotas from time to time during the year, although the DEA has substantial discretion in whether or not to make such adjustments. Our, or our CMOs, quota of an active ingredient may not be sufficient to meet commercial demand or complete clinical trials. Any delay, limitation or refusal by the DEA in establishing our, or our contract manufacturers', quota for controlled substances could delay or stop our clinical trials or product launches, which could have a material adverse effect on our business, financial position and results of operations.

To enforce these requirements, the DEA conducts periodic inspections of registered establishments that handle controlled substances. Failure to maintain compliance with applicable requirements, particularly as manifested in loss or diversion, can result in administrative, civil or criminal enforcement action that could have a material adverse effect on our business, results of operations and financial condition. The DEA may seek civil penalties, refuse to renew necessary registrations, or initiate administrative proceedings to revoke those registrations. In some circumstances, violations could result in criminal proceedings.

Individual states also independently regulate controlled substances. We and our CMOs will be subject to state regulation on distribution of these products, including, for example, state requirements for licensures or registration.

U.S. patent term restoration and marketing exclusivity

Depending upon the timing, duration and specifics of the FDA approval of our drug candidates, some of our U.S. patents may be eligible for limited patent term extension under the Drug Price Competition and Patent Term Restoration Act of 1984, commonly referred to as the Hatch-Waxman Amendments. The Hatch-Waxman Amendments permit a patent restoration term of up to five years as compensation for patent term lost during product development and the FDA regulatory review process. However, patent term restoration cannot extend the remaining term of a patent beyond a total of 14 years from the product's approval date. The patent term restoration period is generally one-half the time between the effective date of an IND and the submission date of an NDA plus the time between the submission date of an NDA and the approval of that application. Only one patent applicable to an approved drug is eligible for the extension and the application for the extension must be submitted prior to the expiration of the patent. The U.S. PTO, in consultation with the FDA, reviews and approves the application for any patent term extension or restoration. In the future, we intend to apply for restoration of patent term for one of our currently owned or licensed patents to add patent life beyond its current expiration date, depending on the expected length of the clinical trials and other factors involved in the filing of the relevant NDA.

Marketing exclusivity provisions under the FDCA can also delay the submission or the approval of certain marketing applications. The FDCA provides a five-year period of non-patent marketing exclusivity within the United States to the first applicant to obtain approval of an NDA for a new chemical entity. A drug is a new chemical entity if the FDA has not previously approved any other new drug containing the same active moiety, which is the molecule or ion responsible for the action of the drug substance. During the exclusivity period, the FDA may not accept for review an abbreviated new drug application, or ANDA, or a 505(b)(2) NDA submitted by another company for another drug based on the same active moiety, regardless of whether the drug is intended for the same indication as the original innovator drug or for another indication, where the applicant does not own or have a legal right of reference to all the data required for approval. However, an application may be submitted after four years if it contains a certification of patent invalidity or non-infringement to one of the patents listed with the FDA by the innovator NDA holder. The FDCA also provides three years of marketing exclusivity for an NDA, or supplement to an existing NDA if new clinical investigations, other than bioavailability studies, that were conducted or sponsored by the applicant are deemed by the FDA to be essential to the approval of the application, for example new indications, dosages or strengths of an existing drug.

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This three-year exclusivity covers only the modification for which the drug received approval on the basis of the new clinical investigations and does not prohibit the FDA from approving ANDAs for drugs containing the active agent for the original indication or condition of use. Five-year and three-year exclusivity will not delay the submission or approval of a full NDA. However, an applicant submitting a full NDA would be required to conduct or obtain a right of reference to all of the non-clinical studies and adequate and well-controlled clinical trials necessary to demonstrate safety and efficacy. Orphan drug exclusivity, as described above, may offer a seven-year period of marketing exclusivity, except in certain circumstances. Pediatric exclusivity is another type of regulatory market exclusivity in the United States. Pediatric exclusivity, if granted, adds six months to existing exclusivity periods and patent terms. This six-month exclusivity, which runs from the end of other exclusivity protection or patent term, may be granted based on the voluntary completion of a pediatric trial in accordance with an FDA-issued Written Request for such a trial.

European Union drug development

In the European Union, our future products may also be subject to extensive regulatory requirements. As in the United States, medicinal products can only be marketed if a marketing authorization from the competent regulatory agencies has been obtained.

Similar to the United States, the various phases of non-clinical and clinical research in the European Union are subject to significant regulatory controls. Although the EU Clinical Trials Directive 2001/20/EC has sought to harmonize the EU clinical trials regulatory framework, setting out common rules for the control and authorization of clinical trials in the EU, the EU Member States have transposed and applied the provisions of the Directive differently. This has led to significant variations in the member state regimes. Under the current regime, before a clinical trial can be initiated it must be approved in each of the EU countries where the trial is to be conducted by two distinct bodies: the National Competent Authority, or NCA, and one or more Ethics Committees, or ECs. Under the current regime all suspected unexpected serious adverse reactions to the investigated drug that occur during the clinical trial have to be reported to the NCA and ECs of the Member State where they occurred.

The EU clinical trials legislation is currently undergoing a revision process mainly aimed at harmonizing and streamlining the clinical trials authorization process, simplifying adverse event reporting procedures, improving the supervision of clinical trials, and increasing their transparency.

European Union drug review and approval

In the European Economic Area, or EEA, (which is comprised of the 27 Member States of the European Union (excluding Croatia) plus Norway, Iceland and Liechtenstein), medicinal products can only be commercialized after obtaining a Marketing Authorization, or MA. There are two types of marketing authorizations:

The Community MA is issued by the European Commission through the Centralized Procedure, based on the opinion of the Committee for Medicinal Products for Human Use, or CHMP, of the European Medicines Agency, or EMA, and is valid throughout the entire territory of the EEA. The Centralized Procedure is mandatory for certain types of products, such as biotechnology medicinal products, orphan medicinal products, and medicinal products containing a new active substance indicated for the treatment of AIDS, cancer, neurodegenerative disorders, diabetes, auto-immune and viral diseases. The Centralized Procedure is optional for products containing a new active substance not yet authorized in the EEA, or for products that constitute a significant therapeutic, scientific or technical innovation or which are in the interest of public health in the EU.

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National MAs, which are issued by the competent authorities of the Member States of the EEA and only cover their respective territory, are available for products not falling within the mandatory scope of the Centralized Procedure. Where a product has already been authorized for marketing in a Member State of the EEA, this National MA can be recognized in another Member States through the Mutual Recognition Procedure. If the product has not received a National MA in any Member State at the time of application, it can be approved simultaneously in various Member States through the Decentralized Procedure. Under the Decentralized Procedure an identical dossier is submitted to the competent authorities of each of the Member States in which the MA is sought, one of which is selected by the applicant as the Reference Member State, or RMS. The competent authority of the RMS prepares a draft assessment report, a draft summary of the product characteristics, or SPC, and a draft of the labeling and package leaflet, which are sent to the other Member States (referred to as the Member States Concerned) for their approval. If the Member States Concerned raise no objections, based on a potential serious risk to public health, to the assessment, SPC, labeling, or packaging proposed by the RMS, the product is subsequently granted a national MA in all the Member States (i.e., in the RMS and the Member States Concerned).

Under the above described procedures, before granting the MA, the EMA or the competent authorities of the Member States of the EEA make an assessment of the risk-benefit balance of the product on the basis of scientific criteria concerning its quality, safety and efficacy.

European Union new chemical entity exclusivity

In the European Union, new chemical entities, sometimes referred to as new active substances, qualify for eight years of data exclusivity upon marketing authorization and an additional two years of market exclusivity. This data exclusivity, if granted, prevents regulatory authorities in the European Union from referencing the innovator's data to assess a generic application for eight years, after which generic marketing authorization can be submitted, and the innovator's data may be referenced, but not approved for two years. The overall ten-year period will be extended to a maximum of 11 years if, during the first eight years of those ten years, the marketing authorization holder obtains an authorization for one or more new therapeutic indications which, during the scientific evaluation prior to their authorization, are held to bring a significant clinical benefit in comparison with existing therapies.

European Union orphan designation and exclusivity

In the European Union, the EMA's Committee for Orphan Medicinal Products grants orphan drug designation to promote the development of products that are intended for the diagnosis, prevention or treatment of life-threatening or chronically debilitating conditions affecting not more than 5 in 10,000 persons in the European Union Community and for which no satisfactory method of diagnosis, prevention, or treatment has been authorized (or the product would be a significant benefit to those affected). Additionally, designation is granted for products intended for the diagnosis, prevention, or treatment of a life-threatening, seriously debilitating or serious and chronic condition and when, without incentives, it is unlikely that sales of the drug in the European Union would be sufficient to justify the necessary investment in developing the medicinal product.

In the European Union, orphan drug designation entitles a party to financial incentives such as reduction of fees or fee waivers and ten years of market exclusivity is granted following medicinal product approval. This period may be reduced to six years if the orphan drug designation criteria are no longer met, including where it is shown that the product is sufficiently profitable not to justify maintenance of market exclusivity. Orphan drug designation must be requested before submitting an application for marketing approval. Orphan drug designation does not convey any advantage in, or shorten the duration of, the regulatory review and approval process.

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Rest of the world regulation

For other countries outside of the European Union and the United States, such as countries in Eastern Europe, Latin America or Asia, the requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement vary from country to country. In all cases the clinical trials must be conducted in accordance with cGCP requirements and the applicable regulatory requirements and the ethical principles that have their origin in the Declaration of Helsinki.

If we fail to comply with applicable foreign regulatory requirements, we may be subject to, among other things, fines, suspension or withdrawal of regulatory approvals, product recalls, seizure of products, operating restrictions and criminal prosecution.

Reimbursement

Sales of our products will depend, in part, on the extent to which our products will be covered by third-party payors, such as government health programs, commercial insurance and managed healthcare organizations. In the United States no uniform policy of coverage and reimbursement for drug products exists. Accordingly, decisions regarding the extent of coverage and amount of reimbursement to be provided for any of our products will be made on a payor by payor basis. As a result, the coverage determination process is often a time-consuming and costly process that will require us to provide scientific and clinical support for the use of our product candidates to each payor separately, with no assurance that coverage and adequate reimbursement will be obtained.

Third-party payors are increasingly reducing reimbursements for medical products and services. Additionally, the containment of healthcare costs has become a priority of federal and state governments, and the prices of drugs have been a focus in this effort. The U.S. government, state legislatures and foreign governments have shown significant interest in implementing cost-containment programs, including price controls, restrictions on reimbursement and requirements for substitution of generic products. Adoption of price controls and cost-containment measures, and adoption of more restrictive policies in jurisdictions with existing controls and measures, could further limit our net revenue and results. Decreases in third-party reimbursement for our product candidate or a decision by a third-party payor to not cover our product candidate could reduce physician usage of the product candidate and have a material adverse effect on our sales, results of operations and financial condition.

The Medicare Prescription Drug, Improvement, and Modernization Act of 2003, or the MMA, established the Medicare Part D program to provide a voluntary prescription drug benefit to Medicare beneficiaries. Under Part D, Medicare beneficiaries may enroll in prescription drug plans offered by private entities that provide coverage of outpatient prescription drugs. Unlike Medicare Part A and B, Part D coverage is not standardized. Part D prescription drug plan sponsors are not required to pay for all covered Part D drugs, and each drug plan can develop its own drug formulary that identifies which drugs it will cover and at what tier or level. However, Part D prescription drug formularies must include drugs within each therapeutic category and class of covered Part D drugs, though not necessarily all the drugs in each category or class. Any formulary used by a Part D prescription drug plan must be developed and reviewed by a pharmacy and therapeutic committee. Government payment for some of the costs of prescription drugs may increase demand for products for which we receive marketing approval. However, any negotiated prices for our products covered by a Part D prescription drug plan will likely be lower than the prices we might otherwise obtain. Moreover, while the MMA applies only to drug benefits for Medicare beneficiaries, private payors often follow Medicare coverage policy and payment limitations in setting their own payment rates. Any reduction in payment that results from the MMA may result in a similar reduction in payments from non-governmental payors.

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The American Recovery and Reinvestment Act of 2009 provides funding for the federal government to compare the effectiveness of different treatments for the same illness. The plan for the research was published in 2012 by the Department of Health and Human Services, the Agency for Healthcare Research and Quality and the National Institutes for Health, and periodic reports on the status of the research and related expenditures will be made to Congress. Although the results of the comparative effectiveness studies are not intended to mandate coverage policies for public or private payors, it is not clear what effect, if any, the research will have on the sales of our product candidate, if any such product or the condition that it is intended to treat is the subject of a trial. It is also possible that comparative effectiveness research demonstrating benefits in a competitor's product could adversely affect the sales of our product candidate. If third-party payors do not consider our products to be cost-effective compared to other available therapies, they may not cover our products after approval as a benefit under their plans or, if they do, the level of payment may not be sufficient to allow us to sell our products on a profitable basis.

The ACA is expected to have a significant impact on the health care industry. The ACA is expected to expand coverage for the uninsured while at the same time containing overall healthcare costs. With regard to pharmaceutical products, among other things, the ACA is expected to expand and increase industry rebates for drugs covered under Medicaid programs and make changes to the coverage requirements under the Medicare Part D program. We cannot predict the full impact of the ACA on our business as many of the ACA reforms require the promulgation of detailed regulations implementing the statutory provisions that has not yet occurred. For example, the ACA imposed new reporting requirements on drug manufacturers for payments made to physicians and teaching hospitals, as well as ownership and investment interests held by physicians and their immediate family members. Failure to submit required information may result in civil monetary penalties of up to an aggregate of \$150,000 per year (or up to an aggregate of \$1 million per year for knowing failures), for all payments, transfers of value or ownership or investment interests that are not timely, accurately and completely reported in an annual submission. Drug manufacturers were required to begin collecting data on August 1, 2013 and were required to submit reports to CMS by March 31, 2014 (and by the 90th day of each subsequent calendar year). In addition, many states have adopted laws similar to the federal laws discussed above. Some of these state prohibitions apply to the referral of patients for healthcare services reimbursed by any insurer, not just federal healthcare programs such as Medicare and Medicaid. There has also been a recent trend of increased federal and state regulation of payments made to physicians. Certain states mandate implementation of compliance programs, impose restrictions on drug manufacturers' marketing practices and/or require the tracking and reporting of gifts, compensation and other remuneration to physicians. In addition, other legislative changes have been proposed and adopted in the United States since the ACA was enacted. On August 2, 2011, the Budget Control Act of 2011 among other things, created measures for spending reductions by Congress. A Joint Select Committee on Deficit Reduction, tasked with recommending a targeted deficit reduction of at least \$1.2 trillion for the years 2013 through 2021, was unable to reach required goals, thereby triggering the legislation's automatic reduction to several government programs. This includes aggregate reductions to Medicare payments to providers of up to 2% per fiscal year, started in April 2013. On January 2, 2013, President Obama signed into law the American Taxpayer Relief Act of 2012, or the ATRA, which delayed for another two months the budget cuts mandated by these sequestration provisions of the Budget Control Act of 2011. The ATRA, among other things, also reduced Medicare payments to several providers, including hospitals, imaging centers and cancer treatment centers, and increased the statute of limitations period for the government to recover overpayments to providers from three to five years. We expect that additional federal healthcare reform measures will be adopted in the future, any of which could limit the amounts that federal and state governments will pay for healthcare products and services, and in turn could significantly reduce the projected value of certain development projects and reduce our profitability.

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In addition, in some foreign countries, the proposed pricing for a drug must be approved before it may be lawfully marketed. The requirements governing drug pricing vary widely from country to country. For example, the European Union provides options for its member states to restrict the range of medicinal products for which their national health insurance systems provide reimbursement and to control the prices of medicinal products for human use. A member state may approve a specific price for the medicinal product or it may instead adopt a system of direct or indirect controls on the profitability of the company placing the medicinal product on the market. There can be no assurance that any country that has price controls or reimbursement limitations for pharmaceutical products will allow favorable reimbursement and pricing arrangements for any of our products. Historically, products launched in the European Union do not follow price structures of the United States and generally prices tend to be significantly lower.

Employees

As of February 28, 2015, we employed 30 full-time employees, including 19 in research and development and 11 in general and administrative, and no part-time employees. 13 of our employees hold M.D. or Ph.D. degrees. We have never had a work stoppage, and none of our employees is represented by a labor organization or under any collective-bargaining arrangements. We consider our employee relations to be good.

Facilities

We lease our office space, which consists of 13,300 square feet located in Cambridge, Massachusetts. Our lease expires on February 28, 2017. We expect to lease additional space prior to the expiration of our lease to meet the needs of the business.

Legal Proceedings

As of the date of this prospectus, we were not party to any legal matters or claims. In the future, we may become party to legal matters and claims arising in the ordinary course of business, the resolution of which we do not anticipate would have a material adverse impact on our financial position, results of operations or cash flows.

JOBS Act

On April 5, 2012, the Jumpstart Our Business Startups Act, or the JOBS Act, was signed into law. The JOBS Act contains provisions that, among other things, reduce certain reporting requirements for an emerging growth company. As an emerging growth company, we are electing not to take advantage of the extended transition period afforded by the JOBS Act for the implementation of new or revised accounting standards and, as a result, we will comply with new or revised accounting standards on the relevant dates on which adoption of such standards is required for non-emerging growth companies. Section 107 of the JOBS Act provides that our decision not to take advantage of the extended transition period is irrevocable.

In addition, we intend to rely on the other exemptions and reduced reporting requirements provided by the JOBS Act. Subject to certain conditions set forth in the JOBS Act, if as an emerging growth company we intend to rely on such exemptions, we are not required to, among other things, (i) provide an auditor's attestation report on our system of internal controls over financial reporting pursuant to Section 404, (ii) provide all of the compensation disclosure that may be required of non-emerging growth public companies under the Dodd-Frank Wall Street Reform and Consumer

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Protection Act, (iii) comply with any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation or a supplement to the auditor's report providing additional information about the audit and the financial statements (auditor discussion and analysis), and (iv) disclose certain executive compensation-related items such as the correlation between executive compensation and performance and comparisons of the Chief Executive Officer's compensation to median employee compensation. These exemptions will apply for a period of five years following the date of our IPO or until we no longer meet the requirements of being an emerging growth company, whichever is earlier.

Table of Contents**MANAGEMENT****Executive Officers and Directors**

The following table sets forth certain information about our executive officers and directors, including their ages as of February 28, 2015:

Name	Age	Position(s)
Executive Officers:		
Jeffrey M. Jonas, M.D.	62	President, Chief Executive Officer and Director
Stephen J. Kanes, M.D., Ph.D.	50	Chief Medical Officer
Albert J. Robichaud, Ph.D.	54	Chief Scientific Officer
Kimi Iguchi	52	Chief Financial Officer
Thomas D. Anderson	59	Chief Commercial Strategy Officer
Directors:		
Michael F. Cola(2)(3)(4)	55	Director
James Frates(1)(2)	47	Director
Robert T. Nelsen(3)	51	Director
Steven Paul, M.D.(1)(4)	64	Director
Kevin P. Starr(2)(3)	52	Director
Howard Pien(1)	57	Director

- (1) Member of the Compensation Committee.
- (2) Member of the Audit Committee.
- (3) Member of the Nominating and Corporate Governance Committee.
- (4) Member of the Science and Technology Committee.

The following paragraphs provide information as of the date of this prospectus about our executive officers, key employees and directors. The information presented includes information about each of our directors' specific experience, qualifications, attributes and skills that led our board of directors to the conclusion that he should serve as a director.

Jeffrey M. Jonas, M.D. Dr. Jonas has served as our Chief Executive Officer and President and a member of our board of directors since August 2013. From 2012 to 2013, Dr. Jonas served as the President of the Regenerative Medicine Division of Shire plc, or Shire, and from 2008 to 2012 as Senior Vice President of Research and Development, Pharmaceuticals at Shire. From 2007 to 2008, Dr. Jonas served as the Executive Vice President of ISIS Pharmaceuticals, Inc., as the Chief Medical Officer and from 2006 to 2007 Executive Vice President of Forest Laboratories, Inc. and from 1991 to 1996 in senior-level positions at Upjohn Laboratories. Dr. Jonas also founded AVAX Technologies, Inc. and SCEPTOR Industries, Inc., where he served as the Chief Executive Officer, President and a Director. Dr. Jonas has published more than 70 scientific papers and chapters, authored more than 100 books, scientific articles and abstracts, and has received numerous awards. Dr. Jonas received his B.A. from Amherst College and M.D. from Harvard Medical School. He completed a residency in psychiatry at Harvard Medical School, and he served as Chief Resident in psychopharmacology at McLean Hospital, Harvard Medical School. Dr. Jonas' qualifications to sit on our board includes more than 20 years of experience on both the scientific and business sides of the pharmaceutical and healthcare industries, particularly in the CNS field.

Stephen J. Kanes, M.D., Ph.D. Dr. Kanes has served as our Chief Medical Officer since July 2013. From 2012 to 2013, he served as the Chair of the neuroscience safety knowledge group at AstraZeneca plc, or AstraZeneca. From 2011 to 2013, Dr. Kanes served as the Executive Director Therapeutic Area Clinical Director for the inflammation, neuroscience and respiratory GMED Division of AstraZeneca. From 2008 to 2012, Dr. Kanes served as the Medical Science Senior Director for the

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neuroscience established brands and emerging anesthesia Group Product Team and in other positions of increasing responsibility in the Neuroscience Discovery Medicine, early and late development groups of AstraZeneca. From 1999 to 2006, Dr. Kanes, served as a practicing psychiatrist. Dr. Kanes was a faculty member in the Psychiatry Department at the University of Pennsylvania School of Medicine, where he continues to serve as an adjunct assistant professor of psychiatry. Dr. Kanes has authored or co-authored more than 30 peer-reviewed publications and has served as an ad hoc reviewer for the journals Neuropsychopharmacology, American Journal of Medical Genetics, and Biological Psychiatry. Dr. Kanes received his B.A. from the University of Pennsylvania and both his Ph.D. and M.D. from State University of New York Stony Brook. Dr. Kanes completed his psychiatry residency at Yale-New Haven Medical Center and postdoctoral fellowship at the University of Pennsylvania.

Albert J. Robichaud, Ph.D. Dr. Robichaud has served as our Chief Scientific Officer since November 2011. From 2010 to 2011, he was Vice President of Chemistry and Pharmacokinetic Sciences at Lundbeck, Inc., where he was responsible for the drug discovery, analytical, computational and pharmacokinetics departments focused on synaptic transmission and neuroinflammation. From 2002 to 2010, Dr. Robichaud was Senior Director and Head of the Neuroscience Discovery Chemistry department of Wyeth Research. During his tenure there, his group successfully delivered more than 15 drug candidates into clinical development in a broad range of neuroscience indications. Dr. Robichaud has co-authored more than 125 manuscripts and abstracts, and is a co-inventor on 45 patents and patent applications. Dr. Robichaud earned a B.S. in chemistry from Rensselaer Polytechnic Institute, a Ph.D. in organic chemistry from the University of California, Irvine and was an American Chemical Society postdoctoral fellow at Colorado State University.

Kimi Iguchi. Ms. Iguchi has served as our Chief Financial Officer since March 2013. From 2008 to 2011, Ms. Iguchi served as the Chief Operating Officer, North America for Santhera Pharmaceuticals Holding AG. From 2004 to 2007, Ms. Iguchi held the role of Vice President of Finance at Cyberkinetics Neurotechnology Systems, Inc. From 1998 to 2004, Ms. Iguchi was the Senior Director of Financial Reporting and Analysis at Millennium Pharmaceuticals, Inc., and from 1996 to 1998 the Senior Manager External Reporting at Biogen, Inc. From 1987 to 1995, Ms. Iguchi also worked as a business assurance manager at PricewaterhouseCoopers LLP. Ms. Iguchi received her B.A. in chemistry from Drew University and an M.B.A. from Northeastern University.

Thomas D. Anderson. Mr. Anderson has served as our Chief Commercial Strategy Officer since April 2014. From 2004 to 2014, Mr. Anderson served as Senior Vice President, Corporate Strategy and Commercial Assessment at Shire Pharmaceuticals Group where he held positions of increasing responsibility. Prior to that, he was Executive Director, Market Research, Business Information at Janssen Pharmaceuticals Inc., President and CEO, Anderson Corporation, President and CEO, Ranir-DCP Corporation, Executive Vice President/COO, Lander Company, Inc. and Product Director, Janssen Pharmaceuticals, Inc. Mr. Anderson received his M.B.A. in finance from Mendoza College of Business Administration at the University of Notre Dame and his B.S. in civil engineering from Lehigh University.

James Frates. Mr. Frates has served as a member of our board of directors since May 2014. He is the Senior Vice President and Chief Financial Officer of Alkermes plc, having held that position since September 2011. From 2007 to 2011, Mr. Frates served as Senior Vice President and Chief Financial Officer of Alkermes, Inc. From 1998 to 2007, Mr. Frates served as Vice President, Chief Financial Officer and Treasurer of Alkermes, Inc. From 1996 to 1998, he was employed at Robertson, Stephens & Company, most recently as a Vice President in Investment Banking. Prior to that time, he was employed at Morgan Stanley & Co. From 2004 to 2009, Mr. Frates served on the board of directors of GPC Biotech AG, a biotechnology company, and was a national director of the Association

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of Bioscience Financial Officers from 2004 to 2009. Mr. Frates is also a Trustee of St. Paul's School. We believe Mr. Frates' qualifications to sit on our board of directors include his leadership experience, financial expertise, business judgment and industry knowledge.

Robert T. Nelsen. Mr. Nelsen has served as a member of our board of directors since September 2013. Mr. Nelsen was a co-founder of ARCH Venture Partners, a venture capital firm, and has served in various capacities for ARCH and affiliated entities since 1986. He is currently a managing director of ARCH Venture Corporation. Mr. Nelsen has played a significant role in the early sourcing, financing and development of more than 30 companies. Mr. Nelsen is a director of Agios Pharmaceuticals, Inc., Kythera Biopharmaceuticals, Inc., Sapphire Energy, Inc., Fate Therapeutics, Inc., Ensemble Therapeutics Corporation, Syros Pharmaceuticals Inc., Bellerophon, LLC, Juno Therapeutics, Inc., and serves as chairman of the board of Hua Medicine. Mr. Nelsen also serves as a Trustee of the Fred Hutchinson Cancer Research Institute, the Institute for Systems Biology, and is a director of the National Venture Capital Association. Mr. Nelsen previously served on the boards of Illumina, Inc., Caliper Life Sciences, Inc., Adolor Corporation, Receptos, Inc., NeurogesX, Inc., Ikaria, Inc., and entities affiliated with deCode Genetics, Inc. among others. Mr. Nelsen received a B.S. with majors in biology and economics from the University of Puget Sound and an M.B.A. from the University of Chicago. Mr. Nelsen's qualifications to sit on our board include his extensive experience as an investor in, and director of, biopharmaceutical and life sciences companies.

Steven Paul, M.D. Dr. Paul has served as a member of our board of directors since September 2011. Dr. Paul is currently a professor of neuroscience, psychiatry and pharmacology at Weill Cornell Medical College. From 2003 to 2010, Dr. Paul, as the Executive Vice President of Eli Lilly and Company, or Eli Lilly, and President of Lilly Research Laboratories, was responsible for Eli Lilly's overall research and development efforts helping to expand Eli Lilly's R&D efforts in oncology and biotechnology resulting in a pipeline of approximately 70 new molecular entities. Dr. Paul spent 17 years at Eli Lilly, during which time he held several key leadership roles, including Vice President of Neuroscience (CNS) Research and Group Vice President of Discovery Research (all therapeutic areas) from 1993 to 2003. Prior to Eli Lilly, from 1988 to 1993 Dr. Paul served as Scientific Director of the National Institute of Mental Health (NIMH). Dr. Paul also served as Medical Director in the Commissioned Corps of the United States Public Health Service. Dr. Paul has been the recipient of many awards and honors and has served on numerous committees and advisory boards. Dr. Paul has also authored or co-authored over 500 papers and book chapters. Dr. Paul is an elected fellow of the American Association for the Advancement of Science and a member of the Institute of Medicine of the National Academy of Sciences. He is also currently on the board of directors or is a trustee of several organizations, including the Sigma-Aldrich Corporation, Alnylam Pharmaceuticals, Inc. and the Foundation for the NIH. Dr. Paul has also served as a member of the National Institute of General Medical Sciences (NIGMS) Advisory Council and was appointed by the Secretary of the Department of Health and Human Services (HHS) as a member of the advisory committee to the Director of the NIH from 2001-2006. Dr. Paul was also a member of the National Advisory Mental Health Council, NIMH, and is board certified by the American Board of Psychiatry and Neurology. Dr. Paul received his B.A. in Biology and Psychology from Tulane University, and his M.S. and M.D. degrees from the Tulane University School of Medicine. Dr. Paul's qualifications to sit on our board include his extensive career in neuroscience and his leadership and managerial experiences at various pharmaceutical and biotechnology companies and healthcare organizations.

Howard Pien. Mr. Pien has served as a member of our board of directors since March 2014. Mr. Pien was the Chairman of the Board and Chief Executive Officer of Medarex, Inc. from 2007 to its acquisition by Bristol-Myers Squibb Company in 2009. Prior to that, he was a private consultant from 2006 to 2007. Prior to 2006, he was President and Chief Executive Officer of Chiron Corporation from 2003 to its acquisition by Novartis in 2006, and before that Mr. Pien held positions of increasing responsibility at GlaxoSmithKline and its predecessor, SmithKline Beecham, at Abbott Laboratories

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and at Merck & Co. He is also a Director of ImmunoGen, Inc., ViroPharma Incorporated, Vanda Pharmaceuticals, Ikaria, Inc. and an Advisor for Warburg Pincus, a private equity firm. Mr. Pien received his MBA in finance from Carnegie Mellon University and his B.S. in civil engineering from Massachusetts Institute of Technology. We believe Mr. Pien's qualifications to sit on our board of directors include his extensive experience working for various pharmaceutical and biotechnology companies.

Michael F. Cola. Mr. Cola has served as a member of our board of directors since September 2014. He currently serves as President and Chief Executive Officer of Medgenics, Inc. Prior to joining Medgenics, from 2005 to 2012, he served as president of specialty pharmaceuticals at Shire plc, a global specialty pharmaceutical company. Previously from 2000 to 2005, Mr. Cola also served as a growth capital provider and president of the life sciences group for Safeguard Scientifics, Inc., where he served as Chairman and Chief Executive Officer of Clariant, Inc., and Chairman of Laureate Pharma, Inc. In addition, Mr. Cola has held senior positions in product development and commercialization at AstraMerck and AstraZeneca. Mr. Cola received a B.A. in biology and physics from Ursinus College and an M.S. in biomedical science from Drexel University. He also serves on the board of directors at Vanda Pharmaceuticals Inc. and Pennsylvania BIO, and serves as chairman for the board of governors of the Boys & Girls Clubs of Philadelphia. We believe Mr. Cola's qualifications to sit on our board of directors include his extensive experience working for various pharmaceutical and biotechnology companies.

Kevin P. Starr. Mr. Starr has served as a member of our board of directors since September 2011. In 2007, Mr. Starr co-founded Third Rock Ventures, a venture capital firm where he remains a partner. From 2003 to 2007, Mr. Starr undertook a number of entrepreneurial endeavors in the life science and entertainment industries. From 2001 to 2002, Mr. Starr served as chief operating officer of Millennium Pharmaceuticals, Inc. He also served as Millennium's chief financial officer from 1998 to 2002. Mr. Starr currently serves on the board of directors of Agios Pharmaceuticals, Inc., Alnylam Pharmaceuticals, Inc., PanOptica, Inc., MyoKardia, Inc., Global Blood Therapeutics, Inc., Afferent Pharmaceuticals, Inc., Ember Therapeutics, Inc. and Zafgen, Inc. Mr. Starr received an M.S. in corporate finance from Boston College and a B.S./B.A. in mathematics and business from Colby College. Mr. Starr's qualifications to serve on our board of directors include his executive management roles with responsibility over key financial and business planning functions and experience in the formation, development and business strategy of multiple start-up companies in the life sciences sector.

Composition of Our Board of Directors

As of February 28, 2015, our board of directors consisted of seven members. Our nominating and corporate governance committee and our board of directors may consider a broad range of factors relating to the qualifications and background of nominees, which may include diversity, which is not only limited to race, gender or national origin. We have no formal policy regarding board diversity. Our nominating and corporate governance committee's and our board of directors' priority in selecting board members is identification of persons who will further the interests of our stockholders through his or her established record of professional accomplishment, the ability to contribute positively to the collaborative culture among board members, knowledge of our business, understanding of the competitive landscape and professional and personal experiences and expertise relevant to our growth strategy. Our directors hold office until their successors have been elected and qualified or until the earlier of their resignation or removal. Our amended and restated certificate of incorporation and amended and restated bylaws also provide that our directors may be removed only for cause by the affirmative vote of the holders of at least 75% of the votes that all our stockholders would be entitled to cast in an annual election of directors, and that any vacancy on our board of directors, including a

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vacancy resulting from an enlargement of our board of directors, may be filled only by vote of a majority of our directors then in office. Notwithstanding the foregoing, Dr. Jonas will serve without further compensation as a member of the board of directors for as long as he serves as our chief executive officer.

Director Independence. Our board of directors has determined that all members of the board of directors, except Jeffrey M. Jonas, M.D. and Kevin P. Starr, are independent directors, including for purposes of the rules of The NASDAQ Stock Market and relevant federal securities laws and regulations. Jeffrey M. Jonas, M.D. is not an independent director under these rules because he is our President and Chief Executive Officer. Kevin P. Starr is not an independent director under these rules because he served as our Interim Chief Executive Officer from January 2011 until July 2013. There are no family relationships among any of our directors or executive officers.

Staggered Board. In accordance with the terms of our amended and restated certificate of incorporation and amended and restated bylaws, our board of directors is divided into three staggered classes of directors of the same or nearly the same number and each will be assigned to one of the three classes. At each annual meeting of the stockholders, a class of directors will be elected for a three-year term to succeed the directors of the same class whose terms are then expiring. The terms of the directors will expire upon the election and qualification of successor directors at the annual meeting of stockholders to be held during the years 2015 for Class I directors, 2016 for Class II directors and 2017 for Class III directors.

Our Class I directors are Steven Paul, M.D. and Robert T. Nelsen;

Our Class II directors are Kevin P. Starr and James Frates; and

Our Class III directors are Jeffrey M. Jonas, M.D., Howard Pien, and Michael F. Cola.

Our amended and restated certificate of incorporation and amended and restated by-laws provide that the number of our directors shall be fixed from time to time by a resolution of the majority of our board of directors. Any additional directorships resulting from an increase in the number of directors will be distributed among the three classes so that, as nearly as possible, each class shall consist of one third of the board of directors.

The division of our board of directors into three classes with staggered three-year terms may delay or prevent stockholder efforts to effect a change of our management or a change in control.

Committees of Our Board of Directors

Our board of directors has established an audit committee, a compensation committee, a nominating and corporate governance committee and a science and technology committee, each of which operate pursuant to a charter adopted by our board of directors. The composition and functioning of all of our committees comply with all applicable requirements of the Sarbanes-Oxley Act of 2002, The NASDAQ Stock Market and the SEC rules and regulations.

Audit committee. Michael F. Cola, James Frates and Kevin P. Starr serve on our audit committee, which is chaired by James Frates. Our board of directors has determined that Michael F. Cola and James Frates are independent for audit committee purposes as that term is defined in the rules of the SEC and the applicable NASDAQ Stock Market rules, and have sufficient knowledge in financial and auditing matters to serve on the audit committee. Our board of directors has designated James Frates as an audit committee financial expert, as defined under the applicable rules of the SEC. Our board has determined that Kevin P. Starr does not satisfy independence requirements under applicable NASDAQ Stock Market rules. The transition rules of the SEC provide that two members of

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the audit committee may be exempt from independence requirements for 90 days after the effectiveness of our registration statement for the IPO, and one member may be exempt for one year after the effectiveness of such registration statement. Our board of directors intends to cause our audit committee to comply with the transition rules within the applicable time periods. The audit committee's responsibilities include:

appointing, approving the compensation of, and assessing the independence of our independent registered public accounting firm;

pre-approving auditing and permissible non-audit services, and the terms of such services, to be provided by our independent registered public accounting firm;

reviewing the overall audit plan with our independent registered public accounting firm and members of management responsible for preparing our consolidated financial statements;

reviewing and discussing with management and our independent registered public accounting firm our annual and quarterly consolidated financial statements and related disclosures as well as critical accounting policies and practices used by us;

coordinating the oversight and reviewing the adequacy of our internal control over financial reporting;

establishing policies and procedures for the receipt and retention of accounting-related complaints and concerns;

recommending based upon the audit committee's review and discussions with management and our independent registered public accounting firm whether our audited consolidated financial statements shall be included in our Annual Report on Form 10-K;

monitoring the integrity of our consolidated financial statements and our compliance with legal and regulatory requirements as they relate to our consolidated financial statements and accounting matters;

preparing the audit committee report required by SEC rules to be included in our annual proxy statement;

reviewing all related person transactions for potential conflict of interest situations and approving all such transactions; and

reviewing quarterly earnings releases and scripts.

Compensation committee. Howard Pien, Steven Paul, M.D. and James Frates serve on our compensation committee, which is chaired by Howard Pien. Our board of directors has determined that each member of the compensation committee is independent as defined in the applicable NASDAQ Stock Market rules. The compensation committee's responsibilities include:

annually reviewing and approving corporate goals and objectives relevant to the compensation of our Chief Executive Officer;

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evaluating the performance of our Chief Executive Officer in light of such corporate goals and objectives and determining the compensation of our Chief Executive Officer;

reviewing and approving the compensation of our other executive officers;

reviewing and establishing our overall management compensation, philosophy and policy;

overseeing and administering our compensation and similar plans;

reviewing and approving our policies and procedures for the grant of equity-based awards;

reviewing and making recommendations to the board of directors with respect to director compensation;

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reviewing and discussing with management the compensation discussion and analysis to be included in our annual proxy statement or Annual Report on Form 10-K; and

reviewing and discussing with the board of directors the corporate succession plans for the Chief Executive Officer and other key officers.

Nominating and corporate governance committee. Michael F. Cola, Kevin P. Starr and Robert T. Nelsen serve on our nominating and corporate governance committee, which is chaired by Michael F. Cola. Our board of directors has determined that Robert T. Nelsen and Michael F. Cola are independent as defined in the applicable NASDAQ Stock Market rules. Our board has determined that Kevin P. Starr does not satisfy the independence requirements under applicable NASDAQ Stock Market rules. The transition rules of the SEC provide that two members of the nominating and corporate governance committee may be exempt from independence requirements for 90 days after the effectiveness of our registration statement for our IPO, and one member may be exempt for one year after the effectiveness of such registration statement. Our board of directors intends to cause our nominating and corporate governance committee to comply with the transition rules within the applicable time periods. The nominating and corporate governance committee's responsibilities include:

developing and recommending to the board of directors criteria for board and committee membership;

establishing procedures for identifying and evaluating board of director candidates, including nominees recommended by stockholders;

reviewing the size and composition of the board of directors to ensure that it is composed of members containing the appropriate skills and expertise to advise us;

identifying individuals qualified to become members of the board of directors;

recommending to the board of directors the persons to be nominated for election as directors and to each of the board's committees;

developing and recommending to the board of directors a code of business conduct and ethics and a set of corporate governance guidelines;

developing a mechanism by which violations of the code of business conduct and ethics can be reported in a confidential manner; and

overseeing the evaluation of the board of directors and management.

Science and Technology Committee. Jeffrey M. Jonas, M.D., Michael F. Cola and Steven Paul, M.D. serve on our science and technology committee. Michael F. Cola and Steven Paul, M.D. serve as co-chairs on our science and technology committee. Our board of directors has determined that Michael F. Cola and Steven Paul, M.D. are independent as defined in the applicable NASDAQ Stock Market rules. The science and technology committee's responsibilities include:

reviewing, evaluating, and advising the board of directors and management regarding the long-term strategic goals and objectives and the quality and direction of our research and development programs;

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monitoring and evaluating trends in research and development, and recommend to our board of directors and management emerging technologies for building the company's technological strength;

recommending approaches to acquiring and maintaining technology positions (including but not limited to contracts, grants, collaborative efforts, alliances, and capital); advising the board of directors and management on the scientific aspects of business development transactions;

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regularly reviewing our research and development pipeline;

assisting the board of directors with its oversight responsibility for enterprise risk management in areas affecting the Company's research and development; and

review such other topics as delegated to the Committee from time to time by the board of directors. Our board of directors may from time to time establish other committees.

Compensation Committee Interlocks and Insider Participation

None of the members of our compensation committee has at any time during the prior three years been one of our officers or employees. None of our executive officers currently serve, or in the past fiscal year has served, as a member of the board of directors or compensation committee of any entity that has one or more executive officers serving on our board of directors or compensation committee.

Corporate Governance

We adopted a code of business conduct and ethics that applies to all of our employees, officers and directors, including those officers responsible for financial reporting. Our code of business conduct and ethics is available on our website at www.sagerx.com. We intend to disclose any amendments to the code, or any waivers of its requirements, on our website or in a Current Report on Form 8-K.

Board Leadership Structure and Board's Role in Risk Oversight

Currently, the role of chairman of the board is separated from the role of Chief Executive Officer, and we plan to keep these roles separated following the completion of this offering. We believe that separating these positions allows our Chief Executive Officer to focus on our day-to-day business, while allowing the chairman of the board to lead the board of directors in its fundamental role of providing advice to and independent oversight of management. Our board of directors recognizes the time, effort and energy that the Chief Executive Officer is required to devote to his position in the current business environment, as well as the commitment required to serve as our chairman, particularly as the board of directors' oversight responsibilities continue to grow. While our amended and restated by-laws and corporate governance guidelines do not require that our chairman and Chief Executive Officer positions be separate, our board of directors believes that having separate positions is the appropriate leadership structure for us at this time and demonstrates our commitment to good corporate governance.

Risk is inherent with every business, and how well a business manages risk can ultimately determine its success. We face a number of risks, including risks relating to our operations, strategic direction and intellectual property as more fully discussed under "Risk Factors" in this prospectus. Management is responsible for the day-to-day management of risks we face, while our board of directors, as a whole and through its committees, has responsibility for the oversight of risk management. In its risk oversight role, our board of directors has the responsibility to satisfy itself that the risk management processes designed and implemented by management are adequate and functioning as designed.

The role of the board of directors in overseeing the management of our risks is conducted primarily through committees of the board of directors, as disclosed in the descriptions of each of the

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committees above and in the charters of each of the committees. The full board of directors (or the appropriate board committee in the case of risks that are under the purview of a particular committee) discusses with management our major risk exposures, their potential impact on us, and the steps we take to manage them. When a board committee is responsible for evaluating and overseeing the management of a particular risk or risks, the chairman of the relevant committee reports on the discussion to the full board of directors during the committee reports portion of the next board meeting. This enables the board of directors and its committees to coordinate the risk oversight role, particularly with respect to risk interrelationships.

Table of Contents**EXECUTIVE AND DIRECTOR COMPENSATION****Summary Compensation Table**

The following table sets forth the compensation paid or accrued during the fiscal year ended December 31, 2014 and 2013 to our Chief Executive Officer and our next two highest-paid executive officers as of December 31, 2014 and 2013. We refer to these officers as our named executive officers.

Name and Principal Position	Year	Salary (\$)	Bonus (\$)	Option Awards⁽¹⁾ (\$)	Non-equity Incentive Plan Compensation (\$)	All Other Compensation (\$)⁽³⁾	Total (\$)
Jeffrey M. Jonas, M.D. ⁽²⁾ <i>Chief Executive Officer</i>	2014	425,000	195,500			125,000	745,500
	2013	152,973		204,774		228,000	585,747
Stephen J. Kanés, M.D., Ph.D. <i>Chief Medical Officer</i>	2014	325,000	112,125			65,000	502,125
	2013	148,958		65,150		65,000	279,108
Albert J. Robichaud, Ph.D. <i>Chief Scientific Officer</i>	2014	300,000	99,000				399,000
	2013	300,000	2,500				302,500

- (1) Amounts represent the aggregate grant-date fair value of option awards granted to our named executive officers in 2014 and 2013 computed in accordance with FASB ASC Topic 718. The assumptions used in the valuation of these awards are consistent with the valuation methodologies specified in the notes to our consolidated financial statements and discussions in Management's Discussion and Analysis of Financial Condition and Results of Operations included elsewhere in this prospectus. The amounts above reflect our aggregate accounting expense for these awards and do not necessarily correspond to the actual value that will be recognized by the named executive officers.
- (2) Dr. Jonas became our Chief Executive Officer in August 2013, and therefore his salary for 2013 is prorated.
- (3) Amounts represent payments of sign-on bonuses.

Narrative to Summary Compensation Table***Employment arrangements with our named executive officers***

We have entered into an employment agreement or letter agreement with each of our named executive officers in connection with their employment with us. Except as noted below, these employment agreements and offer letters provide for at will employment.

We entered into a letter agreement with Dr. Jonas in July 2013 and he assumed the role of Chief Executive Officer in August 2013. The agreement entitles Dr. Jonas to an initial base salary of \$425,000 and eligibility in our bonus pool of up to 40% of his base salary, based upon achievements agreed to between Dr. Jonas and the Board. Dr. Jonas received a signing bonus, with \$225,000 paid out during his first month of employment and an additional \$125,000 paid on the one year anniversary of his employment. Dr. Jonas was also granted options to purchase 701,587 shares of our common stock with an exercise price equal to the fair market value of our common stock on the day of the option grant, with one quarter of such options vesting on the one year anniversary of Dr. Jonas' employment and the rest on a monthly basis for three years thereafter. Dr. Jonas received a bonus of \$195,500 for 2014. As of February 1, 2015, Dr. Jonas' base salary is \$450,000 and his bonus target is 50%.

We entered into a letter agreement with Dr. Kanés in May 2013 and he assumed the role of Chief Medical Officer in August 2013. The agreement entitles Dr. Kanés to an initial base salary of \$325,000.

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Dr. Kanes received a signing bonus, with \$65,000 paid out during his first month of employment and an additional \$65,000 paid on the one year anniversary of his employment. Dr. Kanes was also granted options to purchase 222,222 shares of our common stock with an exercise price equal to the fair market value of our common stock on the day of the option grant, with one quarter of such options vesting on the one year anniversary of Dr. Kanes' employment and the rest on a monthly basis for three years thereafter. Dr. Kanes received a bonus of \$112,125 for 2014. As of February 1, 2015, Dr. Kanes' base salary is \$357,500. As of the completion of our IPO on July 23, 2014, Dr. Kanes was eligible to participate in our bonus pool of up to 30% of his base salary, based upon attainment of corporate and individual goals, as agreed between Dr. Kanes and the Chief Executive Officer and as approved by the Compensation Committee.

We entered into a letter agreement with Dr. Robichaud in September 2011 and he assumed the role of Chief Scientific Officer in November 2011. The agreement entitles Dr. Robichaud to an initial base salary of \$300,000 and eligibility in our bonus pool of up to 30% of his base salary, based upon achievements agreed to between Dr. Robichaud and our Chief Executive Officer. Dr. Robichaud received a signing bonus, with \$65,000 paid out during his first month of employment and an additional \$50,000 paid on the one year anniversary of his employment. Dr. Robichaud was also granted options to purchase 222,222 shares of our common stock with an exercise price equal to the fair market value of our common stock on the day of the option grant, with one quarter of such options vesting on the one year anniversary of Dr. Robichaud's employment and the rest on a monthly basis for three years thereafter. As of the completion of our IPO on July 23, 2014, Dr. Robichaud was eligible to participate in our bonus pool of up to 30% of his base salary, based upon attainment of corporate and individual goals, as agreed between Dr. Robichaud and the Chief Executive Officer and as approved by the Compensation Committee. Dr. Robichaud received a bonus of \$99,000 for 2014. As of February 1, 2015, Dr. Robichaud's base salary is \$315,000.

Payments provided upon termination without cause

We have entered into severance agreements with each of Dr. Jonas, Dr. Kanes and Dr. Robichaud. Pursuant to their severance agreements, each of Dr. Jonas, Dr. Kanes and Dr. Robichaud is eligible to receive certain payments and benefits in the event that such officer's employment is terminated by us without cause (as defined in the severance agreements), or in the event that such officer terminates his employment with good reason (as defined in the severance agreements).

In the event that Dr. Jonas terminates his employment with good reason or is terminated without cause, he is eligible to receive 12 months of base salary continuation and 12 months of COBRA continuation medical benefits subsidized by us, provided that he executes and does not revoke a separation agreement and release of us and our affiliates. In the event that either Dr. Kanes or Dr. Robichaud terminates his employment with good reason or is terminated without cause, such officer is eligible to receive 9 months of base salary continuation and 9 months of COBRA continuation medical benefits subsidized by us, provided that he or she executes and does not revoke a separation agreement and release of us and our affiliates.

Payments provided upon change in control

Pursuant to their change of control agreements, in the event that any of Dr. Jonas, Dr. Kanes or Dr. Robichaud terminates his employment with good reason or is terminated without cause within 12 months of a change in control (as defined in each respective change in control agreement), such officer will be eligible to receive a pro rata portion of such individual's target bonus for that fiscal year and the pro rata portion of that individual's target bonus for that fiscal year based on the number of

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days worked in that fiscal year at the time of termination, and all options and other stock-based awards of such officer shall immediately accelerate and become fully exercisable or nonforfeitable as of the date of termination.

Definitions

For purposes of the severance agreement with each of Dr. Jonas, Dr. Kanes and Dr. Robichaud, **cause** means:

indictment for any felony, any crime involving the Company, or any crime involving fraud, moral turpitude or dishonesty;

any unauthorized use or disclosure of the Company's proprietary information;

any intentional misconduct or gross negligence on the officer's part which has a materially adverse effect on the Company's business or reputation; or

the officer's repeated and willful failure to perform the duties, functions and responsibilities of the officer's position after a written warning from the Company.

For purposes of the severance agreements with each of Dr. Jonas, Dr. Kanes and Dr. Robichaud, **good reason** means:

a material diminution in the officer's responsibilities, authority or duties;

a material diminution in the officer's base salary except for across-the-board salary reductions based on the Company's financial performance similarly affecting all or substantially all senior management employees of the Company;

a material change in the geographic location at which such officer is required to provide services to the Company, not including business travel and short-term assignments; or

a material breach of the severance agreement by the Company.

For purposes of the severance agreements with each of Dr. Jonas, Dr. Kanes and Dr. Robichaud, a **change in control** shall be deemed to have occurred upon the occurrence of any one of the following events:

the sale of all or substantially all of the assets of the Company on a consolidated basis to an unrelated person or entity;

a merger, reorganization or consolidation pursuant to which the holders of the Company's outstanding voting power and outstanding stock immediately prior to such transaction do not own a majority of the outstanding voting power and outstanding stock or other equity interests of the resulting or successor entity (or its ultimate parent, if applicable) immediately upon completion of such transaction;

the sale of all of the stock of the Company to an unrelated person, entity or group thereof acting in concert; or

any other transaction in which the owners of the Company's outstanding voting power immediately prior to such transaction do not own at least a majority of the outstanding voting power of the Company or any successor entity immediately upon completion of the transaction other than as a result of the acquisition of securities directly from the Company.

Employee confidentiality, non-competition, non-solicitation and assignment agreements

Each of our named executive officers has entered into a standard form agreement with respect to confidential information and assignment of inventions. Among other things, this agreement obligates

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each named executive officer to refrain from disclosing any of our proprietary information received during the course of employment and to assign to us any inventions conceived or developed during the course of employment. Such agreement also provides that during the period of the named executive officer's employment and for 12 months thereafter, the named executive officer will not compete with us and will not solicit our employees, consultants, customers or suppliers.

Outstanding Equity Awards at Fiscal Year End

The following table presents the outstanding equity awards held by each of our named executive officers as of December 31, 2014.

Name	Option Awards			Stock Awards		
	Number of Securities Underlying Unexercised Options (#) Exercisable	Number of Securities Underlying Unexercised Options (#) Unexercisable	Option Exercise Price (\$)	Option Expiration Date	Number of Shares or Units of Stock That Have Not Vested (#)	Market Value of Shares or Units of Stock That Have Not Vested (\$) ⁽⁴⁾
Jeffrey M. Jonas, M.D. ⁽¹⁾	233,865	467,722	\$ 0.45	8/12/2023		
Stephen J. Kanes, M.D., Ph.D. ⁽²⁾	78,706	83,330	\$ 0.45	7/23/2023		
Albert J. Robichaud, Ph.D. ⁽³⁾	5,097	17,125	\$ 8.01	1/22/2024	50,919	1,863,635

- (1) Dr. Jonas' options represent options to purchase shares of our common stock granted on August 12, 2013. The shares underlying these options vest as follows: 25% vest on August 12, 2014, with the remainder of the shares vesting in equal monthly installments over the following three years.
- (2) Dr. Kanes' options represent options to purchase shares of our common stock granted on July 23, 2013. The shares underlying these options vest as follows: 25% vest on July 18, 2014, with the remainder of the shares vesting in equal monthly installments over the following three years.
- (3) Dr. Robichaud was granted restricted stock on January 6, 2012. The shares underlying this grant vest as follows: 25% vest on November 7, 2012 with the remainder of the shares vesting in equal monthly installments over the following 36 months. Dr. Robichaud's options represent options to purchase shares of our common stock granted on January 22, 2014. The shares underlying these options vest as follows: 25% vest on January 22, 2015, with the remainder of the shares vesting in equal monthly installments over the following three years.
- (4) Amounts represent the market value using a \$36.60 fair market value of one share of common stock as of the close of business on the NASDAQ Stock Market.

Director Compensation

The following table sets forth a summary of the compensation we paid to our nonemployee directors during 2014. Other than as set forth in the table and described more fully below, we did not pay any compensation, reimburse any expense of, make any equity awards or non-equity awards to, or pay any other compensation to any of the other nonemployee members of our board of directors in 2014. We reimburse nonemployee directors for reasonable travel expenses. Dr. Jonas, our President and Chief Executive Officer, receives no compensation for his service as a director, and, consequently, is not included in this table. The compensation received by Dr. Jonas as an employee during 2014 is presented in the Summary Compensation Table above.

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Name	Fees Earned or Paid in Cash (\$)	Option Awards (\$) ⁽¹⁾	All Other Compensation (\$)	Total (\$)
Robert T. Nelsen	17,395 ⁽³⁾			17,395
Steven Paul, M.D.	18,966 ⁽⁴⁾		76,400 ⁽²⁾	95,366
Kevin P. Starr	37,542 ⁽⁵⁾			37,542
Howard Pien	31,394 ⁽⁶⁾	135,857		167,251
James Frates	29,711 ⁽⁷⁾	150,737		180,448
Michael F. Cola	17,273 ⁽⁸⁾	421,475		438,748

- (1) Amounts represent the aggregate grant-date fair value of option awards granted to our directors in 2014 computed in accordance with FASB ASC Topic 718. The assumptions used in the valuation of these awards are consistent with the valuation methodologies specified in the notes to our consolidated financial statements and discussions in Management's Discussion and Analysis of Financial Condition and Result of Operations, included elsewhere in this prospectus. The amounts above reflect our aggregate accounting expense for these awards and do not necessarily correspond to the actual value that will be recognized by the directors.
- (2) The amounts paid to Steven Paul, M.D. is made through our consulting arrangement with Third Rock Ventures LLC.
- (3) As of December 31, 2014, \$9,500 was not paid.
- (4) As of December 31, 2014, \$10,000 was not paid.
- (5) As of December 31, 2014, \$20,125 was not paid.
- (6) As of December 31, 2014, \$11,250 was not paid.
- (7) As of December 31, 2014, \$13,802 was not paid.
- (8) As of December 31, 2014, \$13,996 was not paid.

In March 2014, our board of directors adopted a nonemployee director compensation policy, as amended on December 12, 2014, that is designed to provide a total compensation package that enables us to attract and retain, on a long-term basis, high caliber nonemployee directors. Under the policy, all nonemployee directors will be paid cash compensation from and after the completion of this offering, as set forth below:

	Annual Retainer
Board of Directors:	
All nonemployee members	\$ 35,000
Additional retainer for Non-Executive Chairman of the Board	\$ 35,000
Audit Committee:	
Chairman	\$ 15,000
Non-Chairman members	\$ 7,500
Compensation Committee:	
Chairman	\$ 10,000
Non-Chairman members	\$ 5,000
Nominating and Corporate Governance Committee:	
Chairman	\$ 7,500
Non-Chairman members	\$ 3,000
Science and Technology Committee:	
Chairman	\$ 10,000
Non-Chairman members	\$ 5,000

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Under the nonemployee director compensation policy, each person who is initially appointed or elected to the board of directors will be eligible for an option grant to purchase up to 20,883 shares of our common stock under our stock option plan on the date he or she first becomes a nonemployee director, of which one third of such options will vest on the one year anniversary of the grant date and the rest on a monthly basis for two years thereafter. In addition, on the date of the annual meeting of stockholders, each continuing nonemployee director who has served on the board of directors for a minimum of six months will be eligible to receive an annual option grant to purchase up to 13,922 shares of our common stock, which will vest in full upon the earlier of the first anniversary of the date of grant or the date of the following annual meeting of stockholders. All of the foregoing options will be granted at fair market value on the date of grant.

Compensation Risk Assessment

We believe that although a portion of the compensation provided to our executive officers and other employees is performance-based, our executive compensation program does not encourage excessive or unnecessary risk-taking. This is primarily due to the fact that our compensation programs are designed to encourage our executive officers and other employees to remain focused on both short-term and long-term strategic goals, in particular in connection with our pay-for-performance compensation philosophy. As a result, we do not believe that our compensation programs are reasonably likely to have a material adverse effect on the company.

Stock Option Plans

2011 Stock Option Plan

The 2011 Stock Option and Grant Plan (the 2011 Stock Option Plan), was approved by our board of directors and our stockholders on September 30, 2011 and was most recently amended on February 12, 2014. Under the 2011 Stock Option Plan, 3,142,857 shares of common stock have been reserved for issuance in the form of incentive stock options, non-qualified stock options, restricted stock, unrestricted stock or any combination of the foregoing. The shares issuable pursuant to awards granted under the 2011 Stock Option Plan are authorized but unissued shares.

As of February 28, 2015, options to purchase 1,717,497 shares of common stock were outstanding under the 2011 Stock Option Plan. Our board determined not to make any further awards under the 2011 Stock Option Plan following the completion of our IPO.

2014 Stock Option Plan

On July 2, 2014, the Company's stockholders approved the 2014 Stock Option and Incentive Plan (the 2014 Stock Option Plan), which became effective upon completion of the IPO. The 2014 Stock Option Plan provides for the grant of restricted stock awards, incentive stock options and non-statutory stock options. The 2014 Stock Option Plan replaced the Company's 2011 Stock Option Plan. Any options or awards outstanding under the 2011 Stock Option Plan remained outstanding and effective.

We initially reserved 1,768,508 shares of common stock for the issuance of awards under the 2014 Stock Option Plan. The 2014 Stock Option Plan provides for an annual increase, to be added on the first day of each fiscal year, by up to 4% of the Company's issued and outstanding shares of common stock on the immediately preceding December 31.

Under the 2014 Stock Option Plan, stock options or stock appreciation rights with respect to no more than 1,768,508 shares may be granted to any one individual in any one calendar year and the

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maximum aggregate number of shares that may be issued in the form of incentive stock options shall not exceed the initial number of shares reserved and available for issuance under the 2014 Stock Option Plan.

The 2014 Stock Option Plan is administered by the compensation committee of the board of directors. The compensation committee has full power to select, from among the individuals eligible for awards, the individuals to whom awards will be granted, to make any combination of awards to participants, and to determine the specific terms and conditions of each award, subject to the provisions of the 2014 Stock Option Plan. The compensation committee has the right to grant the CEO the authority to make awards under specified parameters, including limits on the number of shares to be awarded and the time period over which they may be awarded, and has granted this authority to our CEO in the past. Employees, nonemployee directors and other key persons (including consultants) are eligible to receive awards under the 2014 Stock Option Plan.

The 2014 Stock Option Plan permits the granting of both options to purchase common stock intended to qualify as incentive stock options under Section 422 of the Code and options that do not so qualify. The exercise price of each stock option will be determined by our compensation committee but may not be less than 100% of the fair market value of our common stock on the date of grant or, in the case of an incentive stock option granted to a 10% owner, less than 110% of the fair market value of our common stock on the date of grant. The term of each stock option will be fixed by the compensation committee and may not exceed ten years from the date of grant (or five years in the case of an incentive stock option granted to a 10% owner). The compensation committee will also determine vesting schedule for granted stock options.

The compensation committee may award stock appreciation rights subject to such conditions and restrictions as it may determine. Stock appreciation rights entitle the recipient to shares of common stock, or cash, equal to the value of the appreciation in our stock price over the exercise price. The exercise price of each stock appreciation right may not be less than 100% of the fair market value of the common stock on the date of grant.

The compensation committee may award restricted stock or restricted stock units to participants subject to such conditions and restrictions as it may determine. These conditions and restrictions may include the achievement of certain performance goals and/or continued employment or service with us through a specified vesting period. The compensation committee may also grant cash-based awards to participants subject to such conditions and restrictions as it may determine. Our compensation committee may also grant shares of common stock that are free from any restrictions under the 2014 Stock Option Plan. Unrestricted stock may be granted to participants in recognition of past services or for other valid consideration and may be issued in lieu of cash compensation due to such participant.

The compensation committee may grant performance share awards to participants that entitle the recipient to receive share awards of common stock upon the achievement of certain performance goals and such other conditions as our compensation committee shall determine.

The compensation committee may grant cash bonuses under the 2014 Stock Option Plan to participants, subject to the achievement of certain performance goals.

The compensation committee may grant performance-based awards to participants in the form of restricted stock, restricted stock units, performance shares or cash-based awards upon the achievement of certain performance goals and such other conditions as the compensation committee shall determine. The compensation committee may grant such performance-based awards under the 2014 Stock Option Plan that are intended to qualify as performance-based compensation under Section 162(m) of the Code. Those awards would only vest or become payable upon the attainment of

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performance goals that are established by our compensation committee and related to one or more performance criteria. The performance criteria that could be used with respect to any such awards include: total shareholder return, earnings before interest, taxes, depreciation and amortization, net income (loss) (either before or after interest, taxes, depreciation and/or amortization), changes in the market price of our common stock, economic value-added, sales or revenue, development, clinical or regulatory milestones, acquisitions or strategic transactions, operating income (loss), cash flow (including, but not limited to, operating cash flow and free cash flow), return on capital, assets, equity, or investment, return on sales, gross or net profit levels, productivity, expense, margins, operating efficiency, customer satisfaction, working capital, earnings (loss) per share of stock, sales or market shares and number of customers, any of which may be measured either in absolute terms or as compared to any incremental increase or as compared to results of a peer group. From and after the time that we become subject to Section 162(m) of the Code, the maximum award that is intended to qualify as performance-based compensation under Section 162(m) of the Code that may be made to any one employee during any one calendar year period is 1,768,508 shares with respect to a stock-based award and \$2,000,000 with respect to a cash-based award.

The 2014 Stock Option Plan provides that upon the effectiveness of a sale event, as defined in the 2014 Stock Option Plan, all options and stock appreciation rights that are not exercisable immediately prior to the effective time of the sale event shall become fully exercisable as of the effective time of the sale event, all other awards with time-based vesting, conditions or restrictions, shall become fully vested and nonforfeitable as of the effective time of the sale event and all awards with conditions and restrictions relating to the attainment of performance goals may become vested and nonforfeitable in the discretion of the compensation committee and all awards granted under the 2014 Stock Option Plan shall terminate. In addition, in connection with the termination of the 2014 Stock Option Plan upon a sale event, we may make or provide for a cash payment to participants holding options and stock appreciation rights equal to the difference between the per share cash consideration payable to stockholders in the sale event and the exercise price of the options or stock appreciation rights.

Our board of directors may amend or discontinue the 2014 Stock Option Plan and our compensation committee may amend or cancel outstanding awards for purposes of satisfying changes in law or any other lawful purpose, including option repricing, but no such action may adversely affect rights under an award without the holder's consent. Certain amendments to the 2014 Stock Option Plan may require the approval of our stockholders.

No awards may be granted under the 2014 Stock Option Plan after the date that is ten years from the date of stockholder approval of the 2014 Stock Option Plan.

2014 Employee Stock Purchase Plan

Our 2014 Employee Stock Purchase Plan was adopted by our board of directors on July 2, 2014 and approved by our stockholders on July 2, 2014. Our 2014 Employee Stock Purchase Plan authorizes the initial issuance of up to a total of 282,000 shares of our common stock to participating employees.

All employees who have been employed by us or our designated subsidiaries for at least six months and whose customary employment is for more than 20 hours a week are eligible to participate in our 2014 Employee Stock Purchase Plan. Any employee who owns, or would own upon such purchase under our 2014 Employee Stock Purchase Plan, 5% or more of the voting power or value of our stock is not eligible to purchase shares under our 2014 Employee Stock Purchase Plan.

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We may make one or more offerings to our employees to purchase stock under our 2014 Employee Stock Purchase Plan. Unless otherwise determined by the administrator of our 2014 Employee Stock Purchase Plan, each offering will begin on the first business day occurring on or after each January 1st and July 1st and will end on the last business day occurring on or before the following June 30th and December 31st, respectively, each referred to as offering periods. The administrator may designate different offering periods in its discretion but no offering shall exceed 12 months in duration or overlap with another offering.

Each employee who is a participant in our 2014 Employee Stock Purchase Plan may purchase shares by authorizing payroll deductions of up to 10% of his or her eligible compensation during an offering period. Unless the participating employee has previously withdrawn from the offering, his or her accumulated payroll deductions will be used to purchase common stock on the last business day of the offering period at a price equal to 85% of the fair market value of the common stock on either the first or the last day of the offering period, whichever is lower, provided that no more than 2,500 shares of common stock or such other lesser maximum number established by the plan administrator may be purchased by any one employee during each offering period. Under applicable tax rules, an employee may purchase no more than \$25,000 worth of common stock, valued at the start of the purchase period, under our 2014 Employee Stock Purchase Plan in any calendar year.

The accumulated payroll deductions of any employee who is not a participant on the last day of an offering period will be refunded. An employee's rights under our 2014 Employee Stock Purchase Plan terminate upon voluntary withdrawal from the plan or when the employee ceases employment for any reason.

Our 2014 Employee Stock Purchase Plan may be terminated or amended by our board of directors at any time. Amendments that increase the number of shares of our common stock authorized under our 2014 Employee Stock Purchase Plan and certain other amendments require the approval of our stockholders.

Senior Executive Cash Incentive Bonus Plan

On April 30, 2014, our board of directors adopted the Senior Executive Cash Incentive Bonus Plan, or the Bonus Plan. The Bonus Plan provides for cash bonus payments based upon the attainment of performance targets established by our compensation committee. The payment targets will be related to corporate, financial and operational measures or objectives, or Corporate Performance Goals, as well as individual performance objectives.

Our compensation committee may select Corporate Performance Goals from among the following: cash flow (including, but not limited to, operating cash flow and free cash flow); sales or revenue; corporate revenue; earnings before interest, taxes, depreciation and amortization; net income (loss) (either before or after interest, taxes, depreciation and/or amortization); changes in the market price of our common stock; economic value-added; development, clinical or regulatory milestones; acquisitions or strategic transactions; operating income (loss); return on capital, assets, equity, or investment; stockholder returns; return on sales; gross or net profit levels; productivity; expense efficiency; margins; operating efficiency; customer satisfaction; working capital; earnings (loss) per share of our common stock; bookings, new bookings or renewals; sales or market shares; number of customers; number of new customers or customer references; operating income and/or net annual recurring revenue, any of which may be measured in absolute terms, as compared to any incremental increase, in terms of growth, as compared to results of a peer group, against the market as a whole, compared to applicable market indices and/or measured on a pre-tax or post-tax basis.

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Each employee who is selected to participate in the Bonus Plan will have a target bonus opportunity set for each performance period. The bonus formulas will be adopted in each performance period by the compensation committee and communicated to each employee. The Corporate Performance Goals will be measured at the end of each performance period after our financial reports have been published or such other appropriate time as the compensation committee determines. If the Corporate Performance Goals and individual performance objectives are met, payments will be made as soon as practicable following the end of each performance period. Subject to the rights contained in any agreement between the executive officer and the company, an executive officer must be employed by the company on the bonus payment date to be eligible to receive a bonus payment. The Bonus Plan also permits the compensation committee to approve additional bonuses to executive officers in its sole discretion.

Other Compensation

We currently maintain broad-based benefits that are provided to all employees, including health insurance, life and disability insurance and dental insurance.

Table of Contents**CERTAIN RELATIONSHIPS AND RELATED PARTY TRANSACTIONS**

Other than compensation arrangements, we describe below the transactions, and series of similar transactions, since January 1, 2012, to which we were a party or will be a party, in which:

the amounts involved exceeded or will exceed \$120,000; and

any of our directors, executive officers or holders of more than 5% of our capital stock, or any member of the immediate family of the foregoing persons, had or will have a direct or indirect material interest.

In connection with the completion of our IPO in July 2014, we adopted a related party policy that requires all future transactions between us and any director, executive officer, holder of 5% or more of any class of our capital stock or any member of the immediate family of, or entities affiliated with, any of them, or any other related persons (as defined in Item 404 of Regulation S-K) or their affiliates, in which the amount involved is equal to or greater than \$120,000, be approved in advance by our audit committee. Any request for such a transaction must first be presented to our audit committee for review, consideration and approval. In approving or rejecting any such proposal, our audit committee is to consider the relevant facts and circumstances available and deemed relevant to the audit committee, including, but not limited to, the extent of the related party's interest in the transaction, and whether the transaction is on terms no less favorable to us than terms we could have generally obtained from an unaffiliated third party under the same or similar circumstances.

All of the transactions described below were entered into prior to the adoption of this written policy but each was approved by our board of directors. Prior to our board of directors' consideration of a transaction with a related person, the material facts as to the related person's relationship or interest in the transaction were disclosed to our board of directors, and the transaction was not approved by our board of directors unless a majority of the directors approved the transaction. Our current policy with respect to approval of related person transactions is not set forth in writing.

Private Placements of Securities***Series A preferred stock financing***

On September 30, 2011, we entered into a securities purchase agreement with TRV pursuant to which we agreed to issue, in a series of closings, an aggregate of up to 35,000,000 shares of our Series A redeemable convertible preferred stock at a price of \$1.00 per share. On September 12, 2013, we entered into a joinder and amendment agreement with TRV and ARCH, pursuant to which we increased the number of shares of Series A redeemable convertible preferred stock to be issued under such agreement to 37,500,000.

The following table summarizes the participation in the Series A preferred stock financing by any of our directors, executive officers, holders of more than 5% of our voting securities, or any member of the immediate family of the foregoing persons.

Name	Shares of Series A Preferred	Aggregate Purchase Price Paid	Date Purchased
Third Rock Ventures II, L.P.	6,000,000	\$ 6,000,000	9/30/2011
Third Rock Ventures II, L.P.	4,000,000	\$ 4,000,000	4/9/2012
Third Rock Ventures II, L.P.	5,000,000	\$ 5,000,000	11/12/2012
Third Rock Ventures II, L.P.	5,000,000	\$ 5,000,000	3/18/2013
Third Rock Ventures II, L.P.	5,000,000	\$ 5,000,000	7/1/2013
Third Rock Ventures II, L.P.	7,500,000	\$ 7,500,000	9/12/2013
ARCH Venture Fund VII, L.P.	5,000,000	\$ 5,000,000	9/12/2013

Table of Contents**Series B preferred stock financing**

On October 15, 2013, we entered into a securities purchase agreement with TRV and ARCH pursuant to which we agreed to issue, in a series of closings, up to an aggregate of 13,333,332 shares of our Series B redeemable convertible preferred stock at a price of \$1.50 per share. On February 12, 2014, we entered into an amendment to the securities purchase agreement with TRV and ARCH pursuant to which we decreased the number of shares of Series B redeemable convertible preferred stock to be issued thereunder to 9,999,999.

The following table summarizes the participation in the Series B preferred stock financing by any of our directors, executive officers, holders of more than 5% of our voting securities, or any member of the immediate family of the foregoing persons.

Name	Shares of Series B Preferred	Aggregate Purchase Price Paid	Date Purchased
ARCH Venture Fund VII, L.P.	5,000,000	\$ 7,500,000	1/7/2014
ARCH Venture Fund VII, L.P.	2,500,000	\$ 3,750,000	2/12/2014
Third Rock Ventures II, L.P.	1,666,666	\$ 2,500,000	1/7/2014
Third Rock Ventures II, L.P.	833,333	\$ 1,250,000	2/12/2014

Series C preferred stock financing

On March 11, 2014, we entered into a securities purchase agreement with TRV, ARCH, and certain other investors pursuant to which we agreed to issue in one closing an aggregate of 8,973,905 shares of our Series C redeemable convertible preferred stock at a price of \$4.2345 per share.

The following table summarizes the participation in the Series C preferred stock financing by any of our directors, executive officers, holders of more than 5% of our voting securities, or any member of the immediate family of the foregoing persons.

Name	Shares of Series C Preferred	Aggregate Purchase Price Paid	Date Purchased
ARCH Venture Fund VII, L.P.	885,583	\$ 3,750,000	3/11/2014
Third Rock Ventures II, L.P.	295,194	\$ 1,250,000	3/11/2014
Fidelity Management Research Company & Affiliates	3,542,331	\$ 15,000,000	3/11/2014

Participation in IPO

Certain of our pre-IPO stockholders, officers and directors purchased an aggregate of approximately 40,000 shares of our common stock in our IPO at a price of \$18.00 per share, or approximately \$0.7 million in the aggregate.

Purchaser	Initial Public Offering Shares
Jeffrey M. Jonas, M.D	6,000
Stephen J. Kaness, M.D., Ph.D	3,000
Albert J. Robichaud, Ph.D	4,000
Kimi Iguchi	6,000
Thomas Anderson	6,000

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Purchaser	Initial Public Offering Shares
James Frates	5,000
Steven Paul, M.D.	10,000

Agreements with Stockholders

In connection with the Series C preferred stock financing, we entered into the Second Amended and Restated Investors Rights Agreement, or investor rights agreement, dated as of March 11, 2014, with certain of our stockholders, including our principal stockholders and their affiliates and the Second Amended and Restated Stockholders Agreement, or Stockholders Agreement, dated as of March 11, 2014, with certain of our stockholders, including our principal stockholders and their affiliates. All of the provisions of these agreements terminated immediately upon completion of our IPO, other than the provisions relating to registration rights, which will continue in effect following completion of this offering and entitle the holders of such rights to have us register their shares of our common stock for sale in the United States. See Description of Capital Stock Registration Rights.

During the fiscal years ended December 31, 2012, 2013 and 2014, we incurred consulting and management fees to Third Rock Ventures, LLC, or TRV, in the amount of \$908, \$598 and \$157, respectively. TRV is a management company that is party to a services agreement with Third Rock Ventures, L.P., the beneficial owner of more than 5% of our voting securities. Mr. Starr and Dr. Paul are members of our board of directors, and Mr. Starr is a managing member of TRV GP, LLC, which is the general partner of Third Rock Ventures GP, L.P., the general partner of Third Rock Ventures, L.P. and a managing member of TRV. These consulting fees were paid to TRV in amounts mutually agreed upon in advance by us and TRV in consideration of certain strategic and ordinary course business operations consulting services provided to us on an as-needed basis, from time to time and at our request, by individuals related to TRV, including Dr. Paul but not including Mr. Starr. Such fees were payable pursuant to invoices submitted to us by TRV from time to time. None of these consulting fees were paid directly or indirectly to Mr. Starr and Dr. Paul. The consulting fees paid to TRV did not exceed 5% of the consolidated gross revenue of TRV during any of these fiscal years.

Executive Officer and Director Compensation

See Executive and Director Compensation for information regarding compensation of directors and executive officers.

Employment Agreements

We have entered into offer letters with our executive officers. For more information regarding our agreements with our named executive officers for the fiscal year ended December 31, 2014, see Executive and Director Compensation Narrative to Summary Compensation Table Employment arrangements with our named executive officers.

Indemnification Agreements

We have entered into indemnification agreements with each of our directors and officers, the form of which is attached as an exhibit to the registration statement of which this prospectus is a part. The indemnification agreements and our amended and restated certificate of incorporation and amended and restated by-laws require us to indemnify our directors and officers to the fullest extent permitted by Delaware law.

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The following table presents information concerning the beneficial ownership of the shares of our common stock as of February 28, 2015, by:

each person we know to be the beneficial owner of 5% or more of our outstanding shares of our capital stock;

each of our directors;

each of our named executive officers; and

all of our executive officers and directors as a group.

We have determined beneficial ownership in accordance with SEC rules. The information does not necessarily indicate beneficial ownership for any other purpose. Under these rules, a person is deemed to be a beneficial owner of our common stock if that person has a right to acquire ownership within 60 days by the exercise of vested options or the conversion of our redeemable convertible preferred stock. A person is also deemed to be a beneficial holder of our common stock if that person has or shares voting power, which includes the power to vote or direct the voting of our common stock, or investment power, which includes the power to dispose of or to direct the disposition of such capital stock. Except in cases where community property laws apply or as indicated in the footnotes to this table, we believe that each stockholder identified in the table possesses sole voting and investment power over all shares of common stock shown as beneficially owned by the stockholder.

Percentage of beneficial ownership in the table below is based on 25,808,688 shares of common stock deemed to be outstanding as of February 28, 2015, and _____ shares of common stock outstanding after the completion of this offering. The table below assumes that the underwriters do not exercise their option to purchase additional shares. If the underwriters' option to purchase additional shares is exercised in full, we will sell an aggregate of _____ additional shares of common stock. Shares of common stock subject to options that are currently exercisable or exercisable within 60 days of February 28, 2015 are considered outstanding and beneficially owned by the person holding the options for the purpose of computing the percentage ownership of that person but are not treated as outstanding for the purpose of computing the percentage ownership of any other person. Unless indicated below, the address of each individual listed below is c/o Sage Therapeutics, Inc., 215 First Street Cambridge, MA 02142.

Name and Address of Beneficial Owner ⁽¹⁾	Shares Beneficially Owned Prior to Offering		Shares Beneficially Owned After the Offering	
	Number	Percent	Number	Percent
5% Stockholders				
Third Rock Ventures II, L.P. ⁽²⁾	9,773,073	37.9%		
Entities Affiliated with Fidelity Investment ⁽³⁾	3,868,893	15.0%		
ARCH Venture Fund VII, L.P. ⁽⁴⁾	3,187,044	12.4%		
Named Executive Officers and Directors				
Jeffrey M. Jonas, M.D. ⁽⁵⁾	298,333	1.1%		
Named Executive Officers				
Albert J. Robichaud, Ph.D. ⁽⁶⁾	233,172	*		
Kimi Iguchi ⁽⁷⁾	124,475	*		
Stephen J. Kaness, M.D., Ph.D. ⁽⁸⁾	90,716	*		

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Name and Address of Beneficial Owner ⁽¹⁾	Shares Beneficially Owned Prior to Offering		Shares Beneficially Owned After the Offering	
	Number	Percent	Number	Percent
Other Directors				
Robert T. Nelsen ⁽⁹⁾	3,187,044	12.4%		
Steven Paul, M.D. ⁽¹⁰⁾	803,650	3.1%		
Kevin P. Starr ⁽¹¹⁾				
Howard Pien ⁽¹²⁾	7,937	*		
James Frates ⁽¹³⁾	5,000	*		
Michael F. Cola ⁽¹⁴⁾				
All directors and executive officers as a group (10 persons)	4,750,327	18.4%		

* Indicates beneficial ownership of less than one percent.

- (1) Unless otherwise indicated, the address for each beneficial owner is c/o Sage Therapeutics, Inc., 215 First Street, Cambridge, Massachusetts 02142.
- (2) The address for Third Rock Ventures II, L.P. (TRV LP) is 29 Newbury Street, 3rd Floor, Boston, MA 02116. Consists of 9,773,073 shares of common stock. All shares are held directly by TRV LP. Each of Third Rock Ventures II GP, L.P. (TRV GP), the general partner of TRV LP, Third Rock Ventures II GP, LLC (TRV LLC), the general partner of TRV GP, and Mark Levin, Kevin Starr and Robert Tepper, the managers of TRV LLC, may be deemed to have voting and investment power over the shares held of record by TRV LP.
- (3) The address for Fidelity Management & Research Company, or Fidelity, is 82 Devonshire Street, Boston, Massachusetts 02109, a wholly owned subsidiary of FMR LLC and an investment adviser registered under Section 203 of the Investment Advisers Act of 1940, is the beneficial owner of shares of common stock as a result of acting as investment adviser to various investment companies, or Fidelity Funds, registered under Section 8 of the Investment Company Act of 1940. Based solely on a Schedule 13G filed by FMR LLC on February 13, 2015, consists of 3,868,893 shares of common stock held by entities affiliated with FMR LLC. Edward C. Johnson 3d and FMR LLC, through its control of Fidelity, and the funds each has sole power to dispose of the shares owned by the Funds. Members of the family of Edward C. Johnson 3d, Chairman of FMR LLC, are the predominant owners, directly or through trusts, of Series B voting common shares of FMR LLC, representing 49% of the voting power of FMR LLC. The Johnson family group and all other Series B shareholders have entered into a shareholders' voting agreement under which all Series B voting common shares will be voted in accordance with the majority vote of Series B voting common shares. Accordingly, through their ownership of voting common shares and the execution of the shareholders' voting agreement, members of the Johnson family may be deemed, under the Investment Company Act of 1940, to form a controlling group with respect to FMR LLC. Neither FMR LLC nor Edward C. Johnson 3d, Chairman of FMR LLC, has the sole power to vote or direct the voting of the shares owned directly by the Fidelity Funds, which power resides with the Fidelity Funds' Boards of Trustees. Fidelity carries out the voting of the shares under written guidelines established by the Fidelity Funds' Boards of Trustees.
- (4) The address for ARCH Venture Fund VII, L.P., or ARCH, is 8725 West Higgins Road, Suite 290, Chicago, IL 60631. Consists of 3,187,044 shares of common stock. ARCH Venture Partners VII, L.P. (the GPLP), as the sole general partner of ARCH, may be deemed to beneficially own certain of the shares held by ARCH. The GPLP disclaims beneficial ownership of all shares held by ARCH in which the GPLP does not have an actual pecuniary interest. ARCH Venture Partners VII, LLC (the GPLLC), as the sole general partner of the GPLP, may be deemed to beneficially own certain of the shares held by ARCH. The GPLLC disclaims beneficial ownership of all shares held by ARCH in which it does not have an actual pecuniary interest. The managing directors of the GPLLC, Robert T. Nelsen, Keith Crandell and Clinton Bybee (together, the Managing Directors), are deemed to have voting and dispositive power over the shares held by ARCH, and may be deemed to beneficially own certain of the shares held by ARCH. Mr. Nelsen, a member of

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- our board of directors is one of the Managing Directors. The Managing Directors disclaim beneficial ownership of all shares held by ARCH in which they do not have an actual pecuniary interest.
- (5) Consists of 1,025,587 options to purchase shares of our common stock, 292,333 of which will vest within 60 days of February 28, 2015. Owns 6,000 shares of our common stock.
 - (6) Consists of 222,222 shares of restricted stock and 73,222 options to purchase shares of our common stock. 6,950 of Dr. Robichaud's options will vest within 60 days of February 28, 2015. Owns 4,000 shares of our common stock.
 - (7) Consists of 92,063 shares of restricted stock and 128,315 options to purchase shares of our common stock. 26,412 of Ms. Iguchi's options will vest within 60 days of February 28, 2015. Owns 6,000 shares of our common stock.
 - (8) Consists of 236,810 options to purchase shares of our common stock, 38,368 of which will vest within 60 days of February 28, 2015. Owns 52,348 shares of our common stock.
 - (9) Consists of the shares described in note (3) above. Mr. Nelsen is a managing director of GPLLC, which is the sole general partner of GPLP, which is the sole general partner of ARCH, and as such may be deemed to beneficially own such shares. Mr. Nelsen disclaims beneficial ownership of such shares except to the extent of any pecuniary interest therein.
 - (10) Consists of 793,650 shares of our common stock purchased by Dr. Paul on September 1, 2011, pursuant to a restricted stock purchase agreement with us. Owns 10,000 shares of our common stock.
 - (11) Investment decisions with respect to the shares held by TRV LP are made by an investment committee at TRV GP comprised of Mark Levin, Kevin Starr, and Bob Tepper. No stockholder, director, officer, manager, member or employee of TRV GP or TRV LLC has beneficial ownership (within the meaning of Rule 13d-3 promulgated under the Exchange Act) of any shares held by TRV LP.
 - (12) Consists of 23,809 options to purchase shares of our common stock, 7,937 of which will vest within 60 days of February 28, 2015.
 - (13) Consists of 23,809 options to purchase shares of our common stock, none of which will vest within 60 days of February 28, 2015. Owns 5,000 shares of our common stock.
 - (14) Consists of 20,833 options to purchase shares of our common stock, none of which will vest within 60 days of February 28, 2015.

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DESCRIPTION OF CAPITAL STOCK

Upon the completion of this offering, our authorized capital stock will consist of 120,000,000 shares of common stock, par value \$0.0001 per share, and 5,000,000 shares of preferred stock, par value \$0.0001 per share, all of which will be undesignated, and there will be _____ shares of common stock outstanding and no shares of preferred stock outstanding. As of February 28, 2015, we had approximately 38 record holders of our capital stock. The actual number of stockholders is greater than this number of record holders and includes stockholders who are beneficial owners but whose shares are held in street name by brokers and other nominees. This number of holders of record also does not include stockholders whose shares may be held in trust by other entities.

In addition, upon completion of this offering, and assuming the same capitalization as of February 28, 2015, 2,841,775 options to purchase shares of our common stock will be outstanding and 1,703,807 shares of our common stock will be reserved for future grants under our equity incentive plans.

The following description of our capital stock and provisions of our amended and restated certificate of incorporation and amended and restated by-laws are summaries of material terms and provisions and are qualified by reference to our amended and restated certificate of incorporation and amended and restated by-laws, copies of which have been filed with the SEC as exhibits to the registration statement of which this prospectus is a part. The descriptions of our common stock and preferred stock reflect amendments to our amended and restated certificate of incorporation and amended and restated by-laws.

Common Stock

We are authorized to issue one class of common stock. Holders of our common stock are entitled to one vote for each share of common stock held of record for the election of directors and on all matters submitted to a vote of stockholders. Holders of our common stock are entitled to receive dividends ratably, if any, as may be declared by our board of directors out of legally available funds, subject to any preferential dividend rights of any preferred stock then outstanding. Upon our dissolution, liquidation or winding up, holders of our common stock are entitled to share ratably in our net assets legally available after the payment of all our debts and other liabilities, subject to the preferential rights of any preferred stock then outstanding. Holders of our common stock have no preemptive, subscription, redemption or conversion rights. The rights, preferences and privileges of holders of common stock are subject to, and may be adversely affected by, the rights of the holders of shares of any series of preferred stock that we may designate and issue in the future. Except as described under "Antitakeover Effects of Delaware Law and Provisions of our Amended and Restated Certificate of Incorporation and Amended and Restated By-laws" below, a majority vote of the holders of common stock is generally required to take action under our amended and restated certificate of incorporation and amended and restated by-laws.

Preferred Stock

Our board of directors is authorized, without action by the stockholders, to designate and issue up to an aggregate of 5,000,000 shares of preferred stock in one or more series. Our board of directors can designate the rights, preferences and privileges of the shares of each series and any of its qualifications, limitations or restrictions. Our board of directors may authorize the issuance of preferred stock with voting or conversion rights that could adversely affect the voting power or other rights of the holders of common stock. The issuance of preferred stock, while providing flexibility in connection with possible future financings and acquisitions and other corporate purposes could, under certain circumstances, have the effect of restricting dividends on our common stock, diluting the voting power

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of our common stock, impairing the liquidation rights of our common stock, or delaying, deferring or preventing a change in control of our company, which might harm the market price of our common stock. See also *Antitakeover Effects of Delaware Law and Provisions of our Amended and Restated Certificate of Incorporation and Amended and Restated By-laws* Provisions of our amended and restated certificate of incorporation and amended and restated by-laws *Undesignated preferred stock* below.

Our board of directors will make any determination to issue such shares based on its judgment as to our company's best interests and the best interests of our stockholders. We have no shares of preferred stock outstanding and we have no current plans to issue any shares of preferred stock following completion of this offering.

Registration Rights

The holders of 14,084,664 shares of our common stock or their permitted transferees are entitled to rights with respect to the registration of these securities under the Securities Act. These rights are provided under the terms of the investor rights agreement. The investor rights agreement includes demand registration rights, short-form registration rights and piggyback registration rights. All fees, costs and expenses of underwritten registrations under the investor rights agreement will be borne by us and all selling expenses, including underwriting discounts and selling commissions, will be borne by the holders of the shares being registered.

Demand registration rights

The holders of 14,084,664 shares of our common stock or their permitted transferees are entitled to demand registration rights. Under the terms of the investor rights agreement, we will be required, upon the written request of holders of 25% of these securities, to file a registration statement and use commercially reasonable efforts to effect the registration of all or a portion of these shares for public resale. We are required to effect only two registrations pursuant to this provision of the investor rights agreement. A demand for registration may not be made until 180 days after the completion of our IPO.

Short form registration rights

The holders of 14,084,664 shares of our common stock or their permitted transferees are also entitled to short form registration rights. Pursuant to the investor rights agreement, if we are eligible to file a registration statement on Form S-3, upon the request of 15% of these holders to sell registrable securities at an aggregate price of at least \$1.0 million, we will be required to use our commercially reasonable efforts to effect a registration of such shares. We are required to effect only two registrations in any twelve month period pursuant to this provision of the investor rights agreement.

Piggyback registration rights

The holders of 14,084,664 shares of our common stock or their permitted transferees are entitled to piggyback registration rights. If we register any of our securities either for our own account or for the account of other security holders, the holders of these shares are entitled to include their shares in the registration. Subject to certain exceptions contained in the investor rights agreement, we and the underwriters may limit the number of shares included in the underwritten offering if the underwriters determine in good faith that marketing factors require a limitation of the number of shares to be underwritten.

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Indemnification

Our investor rights agreement contains customary cross-indemnification provisions, under which we are obligated to indemnify holders of registrable securities in the event of material misstatements or omissions in the registration statement attributable to us, and they are obligated to indemnify us for material misstatements or omissions attributable to them.

Expiration of registration rights

The registration rights granted under the investor rights agreement will terminate upon the earlier of (i) a deemed liquidation event, as defined in our amended and restated certificate of incorporation, (ii) at such time when all registrable securities could be sold without restriction under Rule 144 of the Securities Act or (iii) the fifth anniversary of our initial public offering.

Antitakeover Effects of Delaware Law and Provisions of Our Amended and Restated Certificate of Incorporation and Amended and Restated By-Laws

Certain provisions of the Delaware General Corporation Law and of our amended and restated certificate of incorporation and amended and restated by-laws could have the effect of delaying, deferring or discouraging another party from acquiring control of us. These provisions, which are summarized below, are expected to discourage certain types of coercive takeover practices and inadequate takeover bids and, as a consequence, they might also inhibit temporary fluctuations in the market price of our common stock that often result from actual or rumored hostile takeover attempts. These provisions are also designed in part to encourage anyone seeking to acquire control of us to first negotiate with our board of directors. These provisions might also have the effect of preventing changes in our management. It is possible that these provisions could make it more difficult to accomplish transactions that stockholders might otherwise deem to be in their best interests. However, we believe that the advantages gained by protecting our ability to negotiate with any unsolicited and potentially unfriendly acquirer outweigh the disadvantages of discouraging such proposals, including those priced above the then-current market value of our common stock, because, among other reasons, the negotiation of such proposals could improve their terms.

Delaware takeover statute

Upon completion of this offering, we will be subject to the provisions of Section 203 of the Delaware General Corporation Law. In general, Section 203 prohibits a publicly held Delaware corporation from engaging in a business combination with an interested stockholder for a three-year period following the time that this stockholder becomes an interested stockholder, unless the business combination is approved in a prescribed manner. Under Section 203, a business combination between a corporation and an interested stockholder is prohibited unless it satisfies one of the following conditions:

before the stockholder became interested, our board of directors approved either the business combination or the transaction which resulted in the stockholder becoming an interested stockholder;

upon consummation of the transaction which resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction commenced, excluding for purposes of determining the voting stock outstanding, shares owned by persons who are directors and also officers, and employee stock plans, in some instances, but not the outstanding voting stock owned by the interested stockholder; or

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at or after the time the stockholder became interested, the business combination was approved by our board of directors and authorized at an annual or special meeting of the stockholders by the affirmative vote of at least two-thirds of the outstanding voting stock which is not owned by the interested stockholder.

Section 203 defines a business combination to include:

any merger or consolidation involving the corporation and the interested stockholder;

any sale, transfer, lease, pledge, exchange, mortgage or other disposition involving the interested stockholder of 10% or more of the assets of the corporation;

subject to exceptions, any transaction that results in the issuance or transfer by the corporation of any stock of the corporation to the interested stockholder;

subject to exceptions, any transaction involving the corporation that has the effect of increasing the proportionate share of the stock of any class or series of the corporation beneficially owned by the interested stockholder; or

the receipt by the interested stockholder of the benefit of any loans, advances, guarantees, pledges or other financial benefits provided by or through the corporation.

In general, Section 203 defines an interested stockholder as any entity or person beneficially owning 15% or more of the outstanding voting stock of the corporation and any entity or person affiliated with or controlling or controlled by the entity or person.

Provisions of our amended and restated certificate of incorporation and amended and restated by-laws

Our amended and restated certificate of incorporation and amended and restated by-laws include a number of provisions that may have the effect of delaying, deferring or discouraging another party from acquiring control of us and encouraging persons considering unsolicited tender offers or other unilateral takeover proposals to negotiate with our board of directors rather than pursue non-negotiated takeover attempts. These provisions include the items described below.

Board composition and filling vacancies. In accordance with our amended and restated certificate of incorporation, our board is divided into three classes serving staggered three-year terms, with one class being elected each year. Our amended and restated certificate of incorporation also provides that directors may be removed only for cause and then only by the affirmative vote of the holders of 75% or more of the shares then entitled to vote at an election of directors. Furthermore, any vacancy on our board of directors, however occurring, including a vacancy resulting from an increase in the size of our board, may only be filled by the affirmative vote of a majority of our directors then in office even if less than a quorum.

No written consent of stockholders. Our amended and restated certificate of incorporation provides that all stockholder actions are required to be taken by a vote of the stockholders at an annual or special meeting, and that stockholders may not take any action by written consent in lieu of a meeting. This limit may lengthen the amount of time required to take stockholder actions and would prevent the amendment of our by-laws or removal of directors by our stockholder without holding a meeting of stockholders.

Meetings of stockholders. Our amended and restated by-laws provide that only a majority of the members of our board of directors then in office may call special meetings of stockholders and only those matters set forth in the notice of the special meeting may be considered or acted upon at a

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special meeting of stockholders. Our amended and restated by-laws limit the business that may be conducted at an annual meeting of stockholders to those matters properly brought before the meeting.

Advance notice requirements. Our amended and restated by-laws establish advance notice procedures with regard to stockholder proposals relating to the nomination of candidates for election as directors or new business to be brought before meetings of our stockholders. These procedures provide that notice of stockholder proposals must be timely given in writing to our corporate secretary prior to the meeting at which the action is to be taken. Generally, to be timely, notice must be received at our principal executive offices not less than 90 days or more than 120 days prior to the first anniversary date of the annual meeting for the preceding year. The notice must contain certain information specified in our amended and restated by-laws.

Amendment to certificate of incorporation and by-laws. As required by the Delaware General Corporation Law, any amendment of our amended and restated certificate of incorporation must first be approved by a majority of our board of directors, and if required by law or our amended and restated certificate of incorporation, must thereafter be approved by a majority of the outstanding shares entitled to vote on the amendment, and a majority of the outstanding shares of each class entitled to vote thereon as a class, except that the amendment of the provisions relating to stockholder action, directors, limitation of liability and the amendment of our amended and restated certificate of incorporation must be approved by not less than 75% of the outstanding shares entitled to vote on the amendment, and not less than 75% of the outstanding shares of each class entitled to vote thereon as a class. Our amended and restated by-laws may be amended by the affirmative vote of a majority vote of the directors then in office, subject to any limitations set forth in the amended and restated by-laws; and may also be amended by the affirmative vote of at least 75% of the outstanding shares entitled to vote on the amendment, or, if the board of directors recommends that the stockholders approve the amendment, by the affirmative vote of the majority of the outstanding shares entitled to vote on the amendment, in each case voting together as a single class.

Undesignated preferred stock. Our amended and restated certificate of incorporation provides for authorized shares of preferred stock. The existence of authorized but unissued shares of preferred stock may enable our board of directors to render more difficult or to discourage an attempt to obtain control of us by means of a merger, tender offer, proxy contest or otherwise. For example, if in the due exercise of its fiduciary obligations, our board of directors were to determine that a takeover proposal is not in the best interests of us or our stockholders, our board of directors could cause shares of preferred stock to be issued without stockholder approval in one or more private offerings or other transactions that might dilute the voting or other rights of the proposed acquirer or insurgent stockholder or stockholder group. In this regard, our amended and restated certificate of incorporation grants our board of directors broad power to establish the rights and preferences of authorized and unissued shares of preferred stock. The issuance of shares of preferred stock could decrease the amount of earnings and assets available for distribution to holders of shares of common stock. The issuance may also adversely affect the rights and powers, including voting rights, of these holders and may have the effect of delaying, deterring or preventing a change in control of us.

Transfer Agent and Registrar

The transfer agent and registrar for our common stock is Computershare Trust Company, N.A. The transfer agent and registrar's address is 250 Royall Street, Canton, Massachusetts, 02021.

Listing

Our common stock is listed on The NASDAQ Global Market under the symbol SAGE .

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SHARES ELIGIBLE FOR FUTURE SALE

Future sales of our common stock in the public market, or the availability of such shares for sale in the public market, could adversely affect market prices prevailing from time to time. As described below, only a limited number of shares will be available for sale shortly after this offering due to contractual and legal restrictions on resale. Nevertheless, sales of our common stock in the public market after such restrictions lapse, or the perception that those sales may occur, could adversely affect the prevailing market price at such time and our ability to raise equity capital in the future.

Based on the number of shares of common stock outstanding as of February 28, 2015, upon completion of this offering, _____ shares of common stock will be outstanding, assuming no exercise of the underwriter's option to purchase additional shares and no exercise of options. All of the shares sold in this offering will be freely tradable. The remaining shares of common stock outstanding after this offering will be restricted as a result of securities laws or lock-up agreements as described below. Following the expiration of the lock-up period, all shares will be eligible for resale in compliance with Rule 144 or Rule 701 under the Securities Act. Restricted securities as defined under Rule 144 of the Securities Act were issued and sold by us in reliance on exemptions from the registration requirements of the Securities Act. These shares may be sold in the public market only if registered or qualified for an exemption from registration, such as under Rule 144 or Rule 701 under the Securities Act.

Rule 144

In general, under Rule 144 under the Securities Act, as in effect on the date of this prospectus, a person who is one of our affiliates and has beneficially owned shares of our common stock for at least six months would be entitled to sell within any three-month period a number of shares that does not exceed the greater of:

one percent of the number of shares of common stock then outstanding, which will equal approximately _____ shares immediately after the completion of this offering; or

the average weekly trading volume of our common stock on the NASDAQ Global Market during the four calendar weeks preceding the filing of a notice on Form 144 with respect to the sale.

Sales under Rule 144 by our affiliates or persons selling shares on behalf of our affiliates are also subject to manner of sale provisions and notice requirements and to the availability of current public information about us. For a person who has not been deemed to have been one of our affiliates at any time during the 90 days preceding a sale, sales of our securities held longer than six months, but less than one year, will be subject only to the current public information requirement.

In general, under Rule 144 under the Securities Act, as in effect on the date of this prospectus, a person who is not deemed to have been one of our affiliates at any time during the 90 days preceding a sale, and who has beneficially owned the shares proposed to be sold for at least six months, including the holding period of any prior owner other than an affiliate, is entitled to sell the shares without complying with the manner of sale, volume limitation or notice provisions of Rule 144, and will be subject only to the public information requirements of Rule 144. If such a person has beneficially owned the shares proposed to be sold for at least one year, including the holding period of any prior owner other than our affiliates, then such person is entitled to sell such shares without complying with any of the requirements of Rule 144.

Rule 701

Rule 701 under the Securities Act, or Rule 701, as in effect on the date of this prospectus, permits resales of shares in reliance upon Rule 144 but without compliance with certain restrictions of

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Rule 144, including the holding period requirement. Most of our employees, executive officers or directors who purchased shares under a written compensatory plan or contract may be entitled to rely on the resale provisions of Rule 701, but all holders of Rule 701 shares are required to wait until 90 days after the date of this prospectus before selling their shares. However, substantially all Rule 701 shares are subject to lock-up agreements as described below and under "Underwriting" included elsewhere in this prospectus and will become eligible for sale upon the expiration of the restrictions set forth in those agreements.

Lock-up Agreements

In connection with this offering, we, each of our directors and executive officers, and holders holding shares of our outstanding stock have agreed that, subject to limited exceptions, which include:

sales of securities acquired in open market transactions after the completion of this offering;

transfers of securities (i) as a bona fide gift or gifts or (ii) by will or intestacy to the legal representative, heir, beneficiary or a member of the immediate family of the undersigned in a transaction not involving a disposition for value;

if the holder is an individual, transfers of shares of our common stock or any security convertible into our common stock to any trust for the benefit of the holder or the immediate family of the undersigned, or limited partnerships the partners of which are the holder and/or the immediate family members of the holder, in each case for estate planning purposes;

if the holder is a trust, distributions of shares of our common stock or any security convertible into our common stock to its beneficiaries in a transaction not involving a disposition for value;

if the holder is a corporation, limited liability company, partnership or other entity, distribution of shares of our common stock or any security convertible into our common stock to members, stockholders, limited partners, subsidiaries or affiliates (as defined in Rule 405 promulgated under the Securities Act of 1933, as amended) of the holder or to any investment fund or other entity that controls or manages the holder in a transaction not involving a disposition for value;

transfers to us pursuant to agreements under which we have the option to repurchase such shares or securities upon termination of service of the holder;

the receipt by the holder from us of shares of our common stock upon the exercise of options;

the establishment of a trading plan that satisfies the requirements of Rule 10b5-1 under the Exchange Act for the transfer of shares of our common stock; and

sales or transfers of our common stock made pursuant to a trading plan that satisfies the requirements of Rule 10b5-1 under the Exchange Act that has been entered into prior to the date of the lock-up agreement;

without the prior written consent of J.P. Morgan Securities LLC and Goldman, Sachs & Co. on behalf of the underwriters, we and they will not, during the period ending 90 days after the date of this prospectus for us and 60 days after the date of this prospectus for our directors, executive officers and certain holders holding shares of our outstanding stock, (1) offer, pledge, sell, contract to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase, lend, or otherwise transfer or dispose of, directly or indirectly, any shares of our common stock or any securities convertible into, or exercisable or exchangeable for, our common stock (including, without limitation, common stock or such other securities which may be deemed to be beneficially owned by such directors,

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executive officers, managers and members in accordance with the rules and regulations of the SEC and securities which may be issued upon exercise of a stock option or warrant) or publicly disclose the intention to make any offer, sale, pledge or disposition,

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(2) enter into any swap or other arrangement that transfers, in whole or in part, any of the economic consequences of ownership of our common stock or such other securities, whether any such transaction is to be settled by delivery of our common stock or such other securities, in cash or otherwise or (3) make any demand for or exercise any right with respect to the registration of any shares of our common stock or any security convertible into or exercisable or exchangeable for our common stock.

Registration Rights

We are party to an investor rights agreement which provides that holders holding 14,084,664 of the shares of our common stock have the right to demand that we file a registration statement or request that their shares of our common stock be covered by a registration statement that we are otherwise filing. See [Description of Capital Stock Registration Rights](#) in this prospectus. Except for shares purchased by affiliates, registration of their shares under the Securities Act would result in these shares becoming freely tradable without restriction under the Securities Act immediately upon effectiveness of the registration, subject to the expiration of the lock-up period described above and under [Underwriting](#) in this prospectus, and to the extent such shares have been released from any repurchase option that we may hold.

Stock Option Plans

We have filed a Form S-8 registration statement under the Securities Act to register shares of our common stock subject to options outstanding or reserved for issuance under our stock plans. Shares covered by this registration statement will thereupon be eligible for sale in the public markets, subject to Rule 144 limitations applicable to affiliates and any lock-up agreements. For a more complete discussion of our stock plans, see [Executive and Director Compensation Stock Option Plans](#).

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MATERIAL U.S. FEDERAL INCOME AND ESTATE TAX CONSIDERATIONS TO NON-U.S. HOLDERS

The following is a general discussion of material U.S. federal income and estate tax considerations relating to ownership and disposition of our common stock by a non-U.S. holder. For purposes of this discussion, the term "non-U.S. holder" means a beneficial owner of our common stock that is not, for U.S. federal income tax purposes:

an individual who is a citizen or resident of the United States;

a corporation, or other entity treated as a corporation for U.S. federal income tax purposes, created or organized in or under the laws of the United States or of any political subdivision of the United States;

an estate the income of which is subject to U.S. federal income taxation regardless of its source; or

a trust, if a U.S. court is able to exercise primary supervision over the administration of the trust and one or more U.S. persons (as defined in the Code) have authority to control all substantial decisions of the trust or if the trust has a valid election in effect to be treated as a U.S. person under applicable U.S. Treasury Regulations.

A modified definition of "non-U.S. holder" applies for U.S. federal estate tax purposes (as discussed below).

This discussion is based on current provisions of the Code, existing and proposed U.S. Treasury Regulations promulgated thereunder, current administrative rulings and judicial decisions, all as in effect as of the date of this prospectus and all of which are subject to change or to differing interpretation, possibly with retroactive effect. Any change could alter the tax consequences to non-U.S. holders described in this prospectus. In addition, the Internal Revenue Service, or the IRS, could challenge one or more of the tax consequences described in this prospectus.

We assume in this discussion that each non-U.S. holder holds shares of our common stock as a capital asset (generally, property held for investment) within the meaning of Section 1221 of the Code. This discussion does not address all aspects of U.S. federal income and estate taxation that may be relevant to a particular non-U.S. holder in light of that non-U.S. holder's individual circumstances nor does it address any aspects of state, local or non-U.S. taxes, or U.S. federal taxes other than income and estate taxes. This discussion also does not consider any specific facts or circumstances that may apply to a non-U.S. holder and does not address the special tax rules applicable to particular non-U.S. holders, such as:

insurance companies;

tax-exempt organizations;

financial institutions;

brokers or dealers in securities;

regulated investment companies;

pension plans;

controlled foreign corporations;

passive foreign investment companies;

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owners that hold our common stock as part of a straddle, hedge, conversion transaction, synthetic security or other integrated investment;

certain U.S. expatriates;

persons who have elected to mark securities to market;

persons subject to the unearned income Medicare contribution tax;

persons subject to the alternative minimum tax; or

persons that acquire our common stock as compensation for services.

In addition, this discussion does not address the tax treatment of partnerships (including any entity or arrangement treated as a partnership for U.S. federal income tax purposes) or other entities that are transparent for U.S. federal income tax purposes or persons who hold their common stock through partnerships or other entities that are transparent for U.S. federal income tax purposes. In the case of a holder that is classified as a partnership for U.S. federal income tax purposes, the tax treatment of a person treated as a partner in such partnership for U.S. federal income tax purposes generally will depend on the status of the partner and the activities of the partner and the partnership. A person treated as a partner in a partnership or who holds their stock through another transparent entity should consult his, her or its own tax advisor regarding the tax consequences of the ownership and disposition of our common stock through a partnership or other transparent entity, as applicable.

Prospective investors should consult their own tax advisors regarding the U.S. federal, state, local and non-U.S. income and other tax considerations of acquiring, holding and disposing of our common stock.

Distributions on Our Common Stock

We do not currently expect to pay dividends. See **Dividend Policy** above in this prospectus. However, in the event that we do pay distributions of cash or property on our common stock (or in the case of certain redemptions that are treated as distributions with respect to our common stock), those distributions generally will constitute dividends for U.S. federal income tax purposes to the extent paid from our current or accumulated earnings and profits, as determined under U.S. federal income tax principles. If a distribution exceeds our current and accumulated earnings and profits, the excess will be treated as a tax-free return of the non-U.S. holder's investment, up to such holder's tax basis in the common stock. Any remaining excess will be treated as capital gain, subject to the tax treatment described below under the heading **Gain on Sale, Exchange or Other Taxable Disposition of Common Stock**.

Subject also to the discussions below under the headings **Information Reporting and Backup Withholding Tax** and **Foreign Account Tax Compliance Act**, dividends paid to a non-U.S. holder generally will be subject to withholding of U.S. federal income tax at a 30% rate or such lower rate as may be specified by an applicable income tax treaty between the United States and such holder's country of residence. If we determine, at a time reasonably close to the date of payment of a distribution on our common stock, that the distribution will not constitute a dividend because we do not anticipate having current or accumulated earnings and profits, we intend not to withhold any U.S. federal income tax on the distribution as permitted by U.S. Treasury Regulations. If we or another withholding agent apply over-withholding, a non-U.S. holder may be entitled to a refund or credit of any excess tax withheld by timely filing an appropriate claim with the IRS.

Dividends that are treated as effectively connected with a trade or business conducted by a non-U.S. holder within the United States, and, if an applicable income tax treaty so provides, that are

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attributable to a permanent establishment or a fixed base maintained by the non-U.S. holder within the United States, are generally exempt from the 30% withholding tax if the non-U.S. holder satisfies applicable certification and disclosure requirements. To obtain this exemption, a non-U.S. holder must generally provide us with a properly executed original and unexpired IRS Form W-8ECI properly certifying such exemption. However, such U.S. effectively connected income, net of specified deductions and credits, is taxed at the same graduated U.S. federal income tax rates applicable to U.S. persons (as defined in the Code). Any U.S. effectively connected income received by a non-U.S. holder that is a corporation may also, under certain circumstances, be subject to an additional branch profits tax at a 30% rate or such lower rate as may be specified by an applicable income tax treaty between the United States and such holder's country of residence.

A non-U.S. holder of our common stock who claims the benefit of an applicable income tax treaty between the United States and such holder's country of residence generally will be required to provide a properly executed IRS Form W-8BEN (or successor form) and satisfy applicable certification and other requirements. Non-U.S. holders are urged to consult their own tax advisors regarding their entitlement to benefits under a relevant income tax treaty.

A non-U.S. holder that is eligible for a reduced rate of U.S. withholding tax under an income tax treaty may obtain a refund or credit of any excess amounts withheld by timely filing an appropriate claim with the IRS.

Any documentation provided to an applicable withholding agent may need to be updated in certain circumstances. The certification requirements described above also may require a non-U.S. holder to provide its U.S. taxpayer identification number.

Gain on Sale, Exchange or Other Taxable Disposition of Common Stock

Subject to the discussions below under the headings Information Reporting and Backup Withholding Tax and Foreign Account Tax Compliance Act, a non-U.S. holder generally will not be subject to U.S. federal income tax or withholding tax on gain recognized on a sale, exchange or other taxable disposition of our common stock (other than a redemption that is treated as a distribution for U.S. federal income tax purposes and taxed as described above) unless:

the gain is effectively connected with the non-U.S. holder's conduct of a trade or business in the United States, and, if an applicable income tax treaty so provides, the gain is attributable to a permanent establishment or fixed base maintained by the non-U.S. holder in the United States; in these cases, the non-U.S. holder will be taxed on a net income basis at the regular graduated rates and in the manner applicable to U.S. persons, and, if the non-U.S. holder is a foreign corporation, an additional branch profits tax at a rate of 30%, or a lower rate as may be specified by an applicable income tax treaty, may also apply;

the non-U.S. holder is an individual present in the United States for 183 days or more in the taxable year of the disposition and certain other conditions are met, in which case the non-U.S. holder will be subject to a 30% tax (or such lower rate as may be specified by an applicable income tax treaty) on the amount by which the non-U.S. holder's capital gains allocable to U.S. sources exceed capital losses allocable to U.S. sources during the taxable year of the disposition; or

we are or were a U.S. real property holding corporation during a certain look-back period, unless our common stock is regularly traded on an established securities market and the non-U.S. holder held no more than five percent of our outstanding common stock, directly or indirectly, during the shorter of the five-year period ending on the date of the disposition or the period that the non-U.S. holder held our common stock. Generally, a corporation is a U.S. real

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property holding corporation if the fair market value of its U.S. real property interests equals or exceeds 50% of the sum of the fair market value of its worldwide real property interests plus its other assets used or held for use in a trade or business. Although there can be no assurance, we believe that we have not been and are not currently, and we do not anticipate becoming, a U.S. real property holding corporation for U.S. federal income tax purposes.

If these exceptions do not apply, gain on the disposition of shares of our common stock will generally be taxed in the same manner as gain that is effectively connected with the conduct of a U.S. trade or business, except that the branch profits tax will not apply.

Information Reporting and Backup Withholding Tax

We must report annually to the IRS and to each non-U.S. holder the gross amount of the distributions on our common stock paid to such holder and the tax withheld, if any, with respect to such distributions. Non-U.S. holders may have to comply with specific certification procedures to establish that the holder is not a U.S. person (as defined in the Code) in order to avoid backup withholding at the applicable rate with respect to dividends on our common stock. Generally, a holder will comply with such procedures if it provides a properly executed IRS Form W-8BEN or otherwise meets documentary evidence requirements for establishing that it is a non-U.S. holder, or otherwise establishes an exemption.

Information reporting and backup withholding generally will apply to the proceeds of a disposition of our common stock by a non-U.S. holder effected by or through the U.S. office of any broker, U.S. or foreign, unless the holder certifies its status as a non-U.S. holder and satisfies certain other requirements, or otherwise establishes an exemption. Generally, information reporting and backup withholding will not apply to a payment of disposition proceeds to a non-U.S. holder where the transaction is effected outside the United States through a non-U.S. office of a broker. However, for information reporting purposes, dispositions effected through a non-U.S. office of a broker with substantial U.S. ownership or operations generally will be treated in a manner similar to dispositions effected through a U.S. office of a broker. Non-U.S. holders should consult their own tax advisors regarding the application of the information reporting and backup withholding rules to them.

Copies of information returns may be made available to the tax authorities of the country in which the non-U.S. holder resides or is incorporated under the provisions of a specific treaty or agreement. Any documentation provided to an applicable withholding agent may need to be updated in certain circumstances.

Backup withholding is not an additional tax. Any amounts withheld under the backup withholding rules from a payment to a non-U.S. holder can be refunded or credited against the non-U.S. holder's U.S. federal income tax liability, if any, provided that an appropriate claim is timely filed with the IRS.

Foreign Account Tax Compliance Act

Legislation commonly referred to as the Foreign Account Tax Compliance Act and associated guidance, or collectively, FATCA, will generally impose a 30% withholding tax on any withholdable payment (as defined below) to a foreign financial institution, unless such institution enters into an agreement with the U.S. government to collect and provide to the U.S. tax authorities substantial information regarding U.S. account holders of such institution (which would include certain equity and debt holders of such institution, as well as certain account holders that are foreign entities with United States owners) or another applicable exception applies or such institution is compliant with applicable foreign law enacted in connection with an applicable intergovernmental agreement between the United

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States and a foreign jurisdiction. FATCA will also generally impose a 30% withholding tax on any withholdable payment (as defined below) to a foreign entity that is not a financial institution, unless such entity provides the withholding agent with a certification identifying the substantial U.S. owners of the entity (which generally includes any U.S. person who directly or indirectly owns more than 10% of the entity), if any, or another applicable exception applies or such entity is compliant with applicable foreign law enacted in connection with an applicable intergovernmental agreement between the United States and a foreign jurisdiction. Under certain circumstances, a non-U.S. holder might be eligible for refunds or credits of such taxes.

Under final regulations and other current guidance, withholdable payments will generally include dividends on our common stock paid on or after July 1, 2014 and the gross proceeds of a disposition of our common stock paid on or after January 1, 2017. The FATCA withholding tax will apply regardless of whether a payment would otherwise be exempt from or not subject to U.S. nonresident withholding tax (e.g., under the portfolio interest exemption or as capital gain). The IRS is authorized to provide, and has begun the process of providing, rules for coordinating the FATCA withholding regime with the existing nonresident withholding tax rules.

Federal Estate Tax

Common stock owned or treated as owned by an individual who is a non-U.S. holder (as specially defined for U.S. federal estate tax purposes) at the time of death will be included in the individual's gross estate for U.S. federal estate tax purposes and, therefore, may be subject to U.S. federal estate tax, unless an applicable estate tax or other treaty provides otherwise.

The preceding discussion of material U.S. federal tax considerations is for general information only. It is not tax advice. Prospective investors should consult their own tax advisors regarding the particular U.S. federal, state, local and non-U.S. tax consequences of purchasing, holding and disposing of our common stock, including the consequences of any proposed changes in applicable laws.

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UNDERWRITING

We are offering the shares of common stock described in this prospectus through a number of underwriters. J.P. Morgan Securities LLC and Goldman, Sachs & Co. are acting as joint book-running managers of the offering and as representatives of the underwriters. We have entered into an underwriting agreement with the underwriters. Subject to the terms and conditions of the underwriting agreement, we have agreed to sell to the underwriters, and each underwriter has severally agreed to purchase, at the public offering price less the underwriting discounts and commissions set forth on the cover page of this prospectus, the number of shares of common stock listed next to its name in the following table:

Name	Number of Shares
J.P. Morgan Securities LLC	
Goldman, Sachs & Co.	
Leerink Partners LLC	
Cowen & Company	
Total	

The underwriters are committed to purchase all the common shares offered by us if they purchase any shares. The underwriting agreement also provides that if an underwriter defaults, the purchase commitments of non-defaulting underwriters may also be increased or the offering may be terminated.

The underwriters propose to offer the common shares directly to the public at the public offering price set forth on the cover page of this prospectus and to certain dealers at that price less a concession not in excess of \$ _____ per share. Any such dealers may resell shares to certain other brokers or dealers at a discount of up to \$ _____ per share from the public offering price. After the public offering of the shares, the offering price and other selling terms may be changed by the underwriters. Sales of shares made outside of the United States may be made by affiliates of the underwriters.

The underwriters have an option to purchase up to _____ additional shares of common stock from us to cover sales of shares by the underwriters which exceed the number of shares specified in the table above. The underwriters have 30 days from the date of this prospectus to exercise this option. If any shares are purchased with this option, the underwriters will purchase shares in approximately the same proportion as shown in the table above. If any additional shares of common stock are purchased, the underwriters will offer the additional shares on the same terms as those on which the shares are being offered.

The underwriting fee is equal to the public offering price per share of common stock less the amount paid by the underwriters to us per share of common stock. The underwriting fee is \$ _____ per share. The following table shows the per share and total underwriting discounts and commissions to be paid to the underwriters assuming both no exercise and full exercise of the underwriters' option to purchase additional shares.

Per Share	No Exercise	Full Exercise
Total	\$ _____	\$ _____

We estimate that the total expenses of this offering, including registration, filing and listing fees, printing fees and legal and accounting expenses, but excluding the underwriting discounts and commissions, will be approximately \$ _____. We have agreed to reimburse the underwriters for expenses relating to clearance of this offering with the Financial Industry Regulatory Authority in an amount not to exceed \$ _____.

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A prospectus in electronic format may be made available on the web sites maintained by one or more underwriters, or selling group members, if any, participating in the offering. The underwriters may agree to allocate a number of shares to underwriters and selling group members for sale to their online brokerage account holders. Internet distributions will be allocated by the representatives to underwriters and selling group members that may make Internet distributions on the same basis as other allocations.

We have agreed that we will not (i) offer, pledge, announce the intention to sell, sell, contract to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase or otherwise dispose of, directly or indirectly, or file with the SEC a registration statement under the Securities Act relating to, any shares of our common stock or securities convertible into or exchangeable or exercisable for any shares of our common stock, or publicly disclose the intention to make any offer, sale, pledge, disposition or filing, or (ii) enter into any swap or other arrangement that transfers all or a portion of the economic consequences associated with ownership of any shares of common stock or any such other securities (regardless of whether any of these transactions are to be settled by the delivery of shares of common stock or such other securities, in cash or otherwise), in each case without the prior written consent of J.P. Morgan Securities LLC and Goldman, Sachs & Co. for a period of 90 days after the date of this prospectus, other than (a) the shares of our common stock to be sold hereunder, (b) any shares of our common stock issued upon the exercise of options granted under company stock plans, and (c) shares of our common stock or other securities issued in connection with a joint venture, marketing or distribution arrangement, collaboration agreement, intellectual property license agreements, or any acquisition of assets or not less than a majority or controlling portion of the equity of another entity; provided that the aggregate number of shares of our common stock issued pursuant to clause (c) shall not exceed 5.0% of the total number of outstanding shares of our common stock immediately following the issuance and sale of the underwritten shares pursuant to the underwriting agreement and provided, further, the recipient of any such shares of our common stock and securities issued pursuant to clause (c) during the 90-day restricted period described above shall enter into an agreement substantially in the form described thereby.

Our directors and executive officers, and certain of our significant shareholders have entered into lock-up agreements with the underwriters prior to the commencement of this offering pursuant to which each of these persons or entities, for a period of 60 days after the date of this prospectus, may not, without the prior written consent of J.P. Morgan Securities LLC and Goldman, Sachs & Co., (1) offer, pledge, announce the intention to sell, contract to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase, lend or otherwise transfer or dispose of, directly or indirectly, any shares of our common stock or any securities convertible into, or exercisable or exchangeable for, our common stock (including, without limitation, common stock or such other securities which may be deemed to be beneficially owned by such directors, executive officers, managers and members in accordance with the rules and regulations of the SEC and securities which may be issued upon exercise of a stock option or warrant) or (2) enter into any swap or other agreement that transfers, in whole or in part, any of the economic consequences of ownership of our common stock or such other securities, whether any such transaction described in clause (1) or (2) above is to be settled by delivery of our common stock or such other securities, in cash or otherwise, or (3) make any demand for or exercise any right with respect to the registration of any shares of our common stock or any security convertible into or exercisable or exchangeable for our common stock, in each case subject to certain exceptions, including (A) sales of securities acquired in open market transactions after the date of the this offering, (B) transfers of shares of our common stock or other securities as bona fide gifts or by will or intestacy to the legal representative, heir, beneficiary or a member of the immediate family of the person or entity in a transaction not involving a disposition for value, (C) in the case of lock-up agreements signed by directors and officers, transfers or dispositions of shares of our common stock or other securities to

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any trust for the direct or indirect benefit of the director or officer signing the lock-up agreement or the immediate family of such person, in each case for estate planning purposes; (D) in the case of a lock-up agreement signed by a trust, distributions of shares of our common stock or any security directly or indirectly convertible into our common stock to its beneficiaries in a transaction not involving a disposition for value, (E) in the case of lock-up agreements signed by a corporation, limited liability company, partnership or other entity, distribution of shares of our common stock or any security directly or indirectly convertible into shares of our common stock to members, stockholders, limited partners, subsidiaries or affiliates of the undersigned or to any investment fund or other entity that controls or manages the undersigned in a transaction not involving a disposition for value; (F) the receipt by the person or entity signing the lock-up of shares of our common stock in connection with the conversion of the outstanding preferred stock of the Company upon the consummation of this offering into shares of our common stock, provided that any such shares of common stock received upon such conversion shall be subject to the restrictions described herein, (G) transfers to the Company pursuant to agreements under which the Company has the option to repurchase shares or securities upon termination of service of the person or entity, provided that the repurchase price for any such shares or securities shall not exceed the original purchase price paid by the undersigned to the Company for such shares or securities, (H) the receipt by the person or entity from the Company of shares of our common stock upon the exercise of options, provided that any such shares of our common stock received upon such exercise shall be subject to the restrictions described herein, and (I) the establishment of a trading plan pursuant to Rule 10b5-1 under the Exchange Act for the transfer of common stock, provided that such plan does not provide for any transfers of common stock, and no filing with the SEC or other public announcement shall be required or voluntarily made by the director or officer or any other person in connection therewith, in each case during the 60-day restricted period or any extension thereof pursuant to the lock-up agreement.

In the case of any transfer, disposition or distribution pursuant to clause (B), (C), (D) or (E), each transferee, donee or distributee must execute and deliver to the Representatives a lock-up agreement. In addition, in the case of any transfer, disposition or distribution pursuant to clause (A), (B), (C), (D), (E), (F), (G), (H) or (I) no filing by any party under the Exchange Act, or other public announcement may be required or voluntarily made in connection with such transfer, disposition or distribution, other than a filing on a Form 5 made after the expiration of the 60-day restricted period referred to above. In addition, notwithstanding the foregoing restrictions, the director or officer may transfer shares of common stock pursuant to sales in the public market undertaken by such person under a trading plan pursuant to Rule 10b5-1 under the Exchange Act, provided that such trading plan shall have been in effect prior to the date of the lock-up agreement, provided that no amendments or other modifications are made to such plans.

We have agreed to indemnify the underwriters against certain liabilities, including liabilities under the Securities Act.

Our common stock is listed on The NASDAQ Global Market under the symbol SAGE .

In connection with this offering, the underwriters may engage in stabilizing transactions, which involves making bids for, purchasing and selling shares of common stock in the open market for the purpose of preventing or retarding a decline in the market price of the common stock while this offering is in progress. These stabilizing transactions may include making short sales of the common stock, which involves the sale by the underwriters of a greater number of shares of common stock than they are required to purchase in this offering, and purchasing shares of common stock on the open market to cover positions created by short sales. Short sales may be covered shorts, which are short positions in an amount not greater than the underwriters' option to purchase additional shares referred to above, or may be naked shorts, which are short positions in excess of that amount. The underwriters may close out any covered short position either by exercising their option to purchase

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additional shares, in whole or in part, or by purchasing shares in the open market. In making this determination, the underwriters will consider, among other things, the price of shares available for purchase in the open market compared to the price at which the underwriters may purchase shares through their option. A naked short position is more likely to be created if the underwriters are concerned that there may be downward pressure on the price of the common stock in the open market that could adversely affect investors who purchase in this offering. To the extent that the underwriters create a naked short position, they will purchase shares in the open market to cover the position.

The underwriters have advised us that, pursuant to Regulation M of the Securities Act, they may also engage in other activities that stabilize, maintain or otherwise affect the price of the common stock, including the imposition of penalty bids. This means that if the representatives of the underwriters purchase common stock in the open market in stabilizing transactions or to cover short sales, the representatives can require the underwriters that sold those shares as part of this offering to repay the underwriting discount received by them.

These activities may have the effect of raising or maintaining the market price of the common stock or preventing or retarding a decline in the market price of the common stock, and, as a result, the price of the common stock may be higher than the price that otherwise might exist in the open market. If the underwriters commence these activities, they may discontinue them at any time. The underwriters may carry out these transactions on The NASDAQ Stock Market, in the over-the-counter market or otherwise.

In addition, in connection with this offering certain of the underwriters (and selling group members) may engage in passive market making transactions in our common stock on The NASDAQ Stock Market prior to the pricing and completion of this offering. Passive market making consists of displaying bids on The NASDAQ Stock Market no higher than the bid prices of independent market makers and making purchases at prices no higher than these independent bids and effected in response to order flow. Net purchases by a passive market maker on each day are generally limited to a specified percentage of the passive market maker's average daily trading volume in the common stock during a specified period and must be discontinued when such limit is reached. Passive market making may cause the price of our common stock to be higher than the price that otherwise would exist in the open market in the absence of these transactions. If passive market making is commenced, it may be discontinued at any time.

Neither we nor the underwriters can assure investors that the shares will trade in the public market at or above the public offering price.

Other than in the United States, no action has been taken by us or the underwriters that would permit a public offering of the securities offered by this prospectus in any jurisdiction where action for that purpose is required. The securities offered by this prospectus may not be offered or sold, directly or indirectly, nor may this prospectus or any other offering material or advertisements in connection with the offer and sale of any such securities be distributed or published in any jurisdiction, except under circumstances that will result in compliance with the applicable rules and regulations of that jurisdiction. Persons into whose possession this prospectus comes are advised to inform themselves about and to observe any restrictions relating to the offering and the distribution of this prospectus. This prospectus does not constitute an offer to sell or a solicitation of an offer to buy any securities offered by this prospectus in any jurisdiction in which such an offer or a solicitation is unlawful.

This document is only being distributed to and is only directed at (i) persons who are outside the United Kingdom or (ii) investment professionals falling within Article 19(5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005 (the "Order") or (iii) high net worth entities, and other persons to whom it may lawfully be communicated, falling within Article 49(2)(a) to (d) of the Order

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(all such persons together being referred to as relevant persons). The securities are only available to, and any invitation, offer or agreement to subscribe, purchase or otherwise acquire such securities will be engaged in only with, relevant persons. Any person who is not a relevant person should not act or rely on this document or any of its contents.

In relation to each Member State of the European Economic Area (each, a Relevant Member State), no offer of shares may be made to the public in that Relevant Member State other than:

- A. to any legal entity which is a qualified investor as defined in the Prospectus Directive;
- B. to fewer than 100 or, if the Relevant Member State has implemented the relevant provision of the 2010 PD Amending Directive, 150, natural or legal persons (other than qualified investors as defined in the Prospectus Directive), as permitted under the Prospectus Directive, subject to obtaining the prior consent of the representatives; or
- C. in any other circumstances falling within Article 3(2) of the Prospectus Directive,

provided that no such offer of shares shall require the Company or the representatives to publish a prospectus pursuant to Article 3 of the Prospectus Directive or supplement a prospectus pursuant to Article 16 of the Prospectus Directive.

Each person in a Relevant Member State who initially acquires any shares or to whom any offer is made will be deemed to have represented, acknowledged and agreed that it is a qualified investor within the meaning of the law in that Relevant Member State implementing Article 2(1)(e) of the Prospectus Directive. In the case of any shares being offered to a financial intermediary as that term is used in Article 3(2) of the Prospectus Directive, each such financial intermediary will be deemed to have represented, acknowledged and agreed that the shares acquired by it in the offer have not been acquired on a non-discretionary basis on behalf of, nor have they been acquired with a view to their offer or resale to, persons in circumstances which may give rise to an offer of any shares to the public other than their offer or resale in a Relevant Member State to qualified investors as so defined or in circumstances in which the prior consent of the representatives has been obtained to each such proposed offer or resale.

We, the representatives and their affiliates will rely upon the truth and accuracy of the foregoing representations, acknowledgements and agreements.

This prospectus has been prepared on the basis that any offer of shares in any Relevant Member State will be made pursuant to an exemption under the Prospectus Directive from the requirement to publish a prospectus for offers of shares. Accordingly any person making or intending to make an offer in that Relevant Member State of shares which are the subject of the offering contemplated in this prospectus may only do so in circumstances in which no obligation arises for the Company or any of the underwriters to publish a prospectus pursuant to Article 3 of the Prospectus Directive in relation to such offer. Neither the Company nor the underwriters have authorized, nor do they authorize, the making of any offer of shares in circumstances in which an obligation arises for the Company or the underwriters to publish a prospectus for such offer.

For the purpose of the above provisions, the expression an offer to the public in relation to any shares in any Relevant Member State means the communication in any form and by any means of sufficient information on the terms of the offer and the shares to be offered so as to enable an investor to decide to purchase or subscribe the shares, as the same may be varied in the Relevant Member State by any measure implementing the Prospectus Directive in the Relevant Member State and the expression Prospectus Directive means Directive 2003/71/EC (including the 2010 PD Amending Directive, to the extent implemented in the Relevant Member States) and includes any relevant

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implementing measure in the Relevant Member State and the expression "2010 PD Amending Directive" means Directive 2010/73/EU.

In addition, in the United Kingdom, this document is being distributed only to, and is directed only at, and any offer subsequently made may only be directed at persons who are "qualified investors" (as defined in the Prospectus Directive) (i) who have professional experience in matters relating to investments falling within Article 19 (5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005, as amended (the "Order") and/or (ii) who are high net worth companies (or persons to whom it may otherwise be lawfully communicated) falling within Article 49(2)(a) to (d) of the Order (all such persons together being referred to as "relevant persons").

Any person in the United Kingdom that is not a relevant person should not act or rely on the information included in this document or use it as basis for taking any action. In the United Kingdom, any investment or investment activity that this document relates to may be made or taken exclusively by relevant persons. Any person in the United Kingdom that is not a relevant person should not act or rely on this document or any of its contents.

Certain of the underwriters and their affiliates have provided in the past to us and our affiliates and may provide from time to time in the future certain commercial banking, financial advisory, investment banking and other services for us and such affiliates in the ordinary course of their business, for which they have received and may continue to receive customary fees and commissions. In addition, from time to time, certain of the underwriters and their affiliates may effect transactions for their own account or the account of customers, and hold on behalf of themselves or their customers, long or short positions in our debt or equity securities or loans, and may do so in the future.

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LEGAL MATTERS

The validity of the common stock offered hereby will be passed upon for us by Goodwin Procter LLP, Boston, Massachusetts and for the underwriters by Ropes & Gray LLP, Boston, Massachusetts.

EXPERTS

The financial statements as of December 31, 2014 and 2013 and for each of the three years in the period ended December 31, 2014 included in this Prospectus have been so included in reliance on the report of PricewaterhouseCoopers LLP, an independent registered public accounting firm, given on the authority of said firm as experts in auditing and accounting.

WHERE YOU CAN FIND MORE INFORMATION

We have filed with the SEC a registration statement on Form S-1 under the Securities Act that registers the shares of our common stock to be sold in this offering. This prospectus does not contain all of the information set forth in the registration statement and the exhibits and schedules filed as part of the registration statement. For further information with respect to us and our common stock, we refer you to the registration statement and the exhibits and schedules filed as a part of the registration statement. Statements contained in this prospectus concerning the contents of any contract or any other document are not necessarily complete. If a contract or document has been filed as an exhibit to the registration statement, we refer you to the copy of the contract or document that has been filed. Each statement in this prospectus relating to a contract or document filed as an exhibit is qualified in all respects by the filed exhibit. The reports and other information we file with the SEC can be read and copied at the SEC's Public Reference Room at 100 F Street, NE, Washington D.C. 20549. Copies of these materials can be obtained at prescribed rates from the Public Reference Section of the SEC at the principal offices of the SEC, 100 F Street, NE, Washington D.C. 20549. You may obtain information regarding the operation of the public reference room by calling 1(800) SEC-0330. The SEC also maintains a web site (<http://www.sec.gov>) that contains reports, proxy and information statements and other information regarding issuers like us that file electronically with the SEC.

We are subject to the reporting and information requirements of the Exchange Act and, as a result, we file periodic reports, proxy statements and other information with the SEC. These periodic reports, proxy statements and other information will be available for inspection and copying at the SEC's public reference room and the web site of the SEC referred to above.

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Report of Independent Registered Public Accounting Firm

To the Board of Directors and Stockholders of

Sage Therapeutics, Inc.:

In our opinion, the accompanying consolidated balance sheets and the related consolidated statements of operations and comprehensive loss, of changes in redeemable convertible preferred stock and shareholders' equity (deficit) and of cash flows present fairly, in all material respects, the financial position of Sage Therapeutics, Inc. and its subsidiary at December 31, 2014 and December 31, 2013 and the results of their operations and their cash flows for each of the three years in the period ended December 31, 2014 in conformity with accounting principles generally accepted in the United States of America. These financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements based on our audits. We conducted our audits of these statements in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. An audit includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements, assessing the accounting principles used and significant estimates made by management, and evaluating the overall financial statement presentation. We believe that our audits provide a reasonable basis for our opinion.

/s/ PricewaterhouseCoopers LLP

Boston, Massachusetts

March 6, 2015

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Table of Contents**Sage Therapeutics, Inc. and Subsidiary****Consolidated Balance Sheets**

(in thousands, except share and per share data)

	December 31, 2014	December 31, 2013
Assets		
Current assets:		
Cash and cash equivalents	\$ 127,766	\$ 8,066
Prepaid expenses and other current assets	1,056	341
Total current assets	128,822	8,407
Property and equipment, net	163	86
Restricted cash	39	39
Deferred tax assets	641	
Total assets	\$ 129,665	\$ 8,532
Liabilities, Redeemable Convertible Preferred Stock and Stockholders Equity (Deficit)		
Current liabilities:		
Accounts payable	\$ 2,429	\$ 1,988
Accrued expenses	4,687	327
Deferred tax liabilities	641	
Total current liabilities	7,757	2,315
Other liabilities	23	44
Total liabilities	7,780	2,359
Commitments and contingencies (Note 4)		
Redeemable convertible preferred stock (Series A, B and C), \$0.0001 par value; no shares and 37,750,000 shares authorized at December 31, 2014 and 2013, respectively; no shares and 37,750,000 shares issued and outstanding at December 31, 2014 and 2013, respectively; liquidation preference of \$0 and \$40,663 at December 31, 2014 and 2013, respectively		37,709
Stockholders' equity (deficit):		
Preferred stock, \$0.0001 par value; 5,000,000 and no shares authorized at December 31, 2014 and 2013, respectively; no shares issued or outstanding at December 31, 2014 and 2013, respectively		
Common stock, \$0.0001 par value; 120,000,000 and 66,000,000 shares authorized at December 31, 2014 and 2013, respectively; 25,621,791 and 1,622,761 shares issued and outstanding at December 31, 2014 and 2013, respectively	3	
Additional paid-in capital	188,727	139
Accumulated deficit	(66,845)	(31,675)
Total stockholders' equity (deficit)	121,885	(31,536)
Total liabilities, redeemable convertible preferred stock and stockholders' equity (deficit)	\$ 129,665	\$ 8,532

The accompanying notes are an integral part of these consolidated financial statements.

Table of Contents**Sage Therapeutics, Inc. and Subsidiary****Consolidated Statements of Operations and Comprehensive Loss**

(in thousands, except share and per share data)

	2014	2013	2012
Operating expenses:			
Research and development	\$ 24,100	\$ 14,357	\$ 7,229
General and administrative	9,710	3,922	2,402
Total operating expenses	33,810	18,279	9,631
Loss from operations	(33,810)	(18,279)	(9,631)
Interest income (expense), net	8	1	
Other income (expense), net	(9)	(3)	(1)
Net loss and comprehensive loss	(33,811)	(18,281)	(9,632)
Accretion of redeemable convertible preferred stock to redemption value	(2,294)	(7)	(4)
Net loss attributable to common stockholders	\$ (36,105)	\$ (18,288)	\$ (9,636)
Net loss per share attributable to common stockholders basic and diluted	\$ (1.67)	\$ (12.26)	\$ (8.62)
Weighted average number of common shares used in net loss per share attributable to common stockholders basic and diluted	21,574,347	1,492,288	1,118,288

The accompanying notes are an integral part of these consolidated financial statements.

Table of Contents**Sage Therapeutics, Inc. and Subsidiary****Consolidated Statements of Changes in Redeemable Convertible Preferred Stock and Stockholders Equity (Deficit)**

(in thousands, except share data)

	Series A, B and C Redeemable Convertible Preferred Stock		Common Stock		Additional Paid-in Capital	Accumulated Deficit	Total Stockholders Equity (Deficit)
	Shares	Amount	Shares	Amount			
Balances at December 31, 2011	6,000,000	\$ 5,972	680,560	\$	\$	\$ (3,761)	\$ (3,761)
Issuance of Series A Preferred Stock, net of issuance costs of \$6	9,000,000	8,994					
Issuance of common stock from exercise of stock options			5,555				
Vesting of restricted stock			709,158		3		3
Accretion of Series A Preferred Stock issuance costs		4			(3)	(1)	(4)
Net loss						(9,632)	(9,632)
Balances at December 31, 2012	15,000,000	14,970	1,395,273			(13,394)	(13,394)
Issuance of Series A Preferred Stock, net of issuance costs of \$18	22,750,000	22,732					
Issuance of common stock from exercise of stock options			3,174		1		1
Vesting of restricted stock			176,695		20		20
Accretion of Series A Preferred Stock issuance costs		7			(7)		(7)
Issuance of common stock in payment of licensing fees			47,619		64		64
Stock-based compensation expense					61		61
Net loss						(18,281)	(18,281)
Balances at December 31, 2013	37,750,000	37,709	1,622,761		139	(31,675)	(31,536)
Issuance of Series B Preferred Stock, net of issuance costs of \$30	9,999,999	14,970					
Issuance of Series C Preferred Stock, net of issuance costs of \$110	8,973,905	37,890					
Issuance of common stock from exercise of stock options			87,475		40		40
Vesting of restricted stock			138,108		14		14
Issuance of common stock in payment of consultant fees			15,872		127		127
Stock-based compensation expense					2,512		2,512
Accretion of redeemable convertible preferred stock to redemption value		2,294			(935)	(1,359)	(2,294)
Conversion of redeemable convertible preferred stock to common stock	(56,723,904)	(92,863)	18,007,575	2	92,861		92,863
Initial public offering of common stock, net of offering costs			5,750,000	1	93,969		93,970
Net loss						(33,811)	(33,811)
Balances at December 31, 2014		\$	25,621,791	\$ 3	\$ 188,727	\$ (66,845)	\$ 121,885

The accompanying notes are an integral part of these consolidated financial statements.

Table of Contents**Sage Therapeutics, Inc. and Subsidiary****Consolidated Statements of Cash Flows**

(in thousands)

	Year Ended December 31,		
	2014	2013	2012
Cash flows from operating activities			
Net loss	\$ (33,811)	\$ (18,281)	\$ (9,632)
Adjustments to reconcile net loss to net cash used in operating activities:			
Stock-based compensation expense	2,512	61	
Licensing or consulting fees paid in common stock	127	64	
Depreciation and amortization	51	47	44
Changes in operating assets and liabilities:			
Prepaid expenses and other current assets	(715)	(317)	71
Accounts payable	441	674	609
Accrued expenses	4,374	236	(18)
Other liabilities	(21)		
Net cash used in operating activities	(27,042)	(17,516)	(8,926)
Cash flows from investing activities			
Purchase of property and equipment	(128)	(3)	(111)
Net cash used in investing activities	(128)	(3)	(111)
Cash flows from financing activities			
Proceeds from the issuance of Series A preferred stock, net of issuance costs		22,732	8,994
Proceeds from the issuance of Series B preferred stock, net of issuance costs	14,970		
Proceeds from the issuance of Series C preferred stock, net of issuance costs	37,890		
Proceeds from the issuance of common stock and restricted stock, net	40	51	3
Proceeds from initial public offering of common stock, net of commissions and underwriting discounts	96,255		
Payment of offering costs	(2,285)		
Net cash provided by financing activities	146,870	22,783	8,997
Net increase (decrease) in cash and cash equivalents	119,700	5,264	(40)
Cash and cash equivalents at beginning of period	8,066	2,802	2,842
Cash and cash equivalents at end of period	\$ 127,766	\$ 8,066	\$ 2,802
Supplemental disclosure of non-cash investing and financing activities			
Accretion of redeemable convertible preferred stock to redemption value	\$ 2,294	\$ 7	\$ 4
Conversion of redeemable convertible preferred stock to common stock	\$ 92,863	\$	\$

The accompanying notes are an integral part of these consolidated financial statements.

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

1. Nature of the Business

Sage Therapeutics, Inc. (Sage or the Company) is a biopharmaceutical company committed to developing and commercializing novel medicines to treat life-threatening, rare central nervous system (CNS) disorders, where there are inadequate or no approved existing therapies. The Company is targeting CNS indications where patient populations are easily identified, acute treatment is typically initiated in the hospital setting, clinical endpoints are well-defined, and development pathways are feasible. This focus allows the Company to make highly informed decisions when advancing its product candidates through the development process. The Company s initial product candidates are aimed at treating different stages of status epilepticus, a life-threatening condition in which the brain is in a state of persistent seizure.

The Company was incorporated under the laws of the state of Delaware on April 16, 2010 and commenced operations on January 19, 2011 as Sterogen Biopharma, Inc. On September 13, 2011, the Company changed its name to Sage Therapeutics, Inc. under its second Amended and Restated Certificate of Incorporation.

The Company is subject to risks and uncertainties common to early-stage companies in the biotech industry, including, but not limited to, development by competitors of new technological innovations, dependence on key personnel protection of proprietary technology, compliance with government regulations, and ability to secure additional capital to fund operations.

We have incurred losses and negative cash flows from operations since our inception. As of December 31, 2014, we had an accumulated deficit of \$66.9 million. From our inception through December 31, 2014, we raised aggregate net proceeds of \$90.6 million from the issuance of Series A, Series B and Series C redeemable convertible preferred stock. In July 2014, we raised gross proceeds of \$96.3 million from the sale of common stock in our initial public offering. We believe our cash balance of \$127.8 million as of December 31, 2014 will be sufficient to fund our anticipated level of operations for at least the next 12 months. The future viability of the Company is largely dependent on its ability to raise additional capital to finance its operations. Management expects that future sources of funding may include new or expanded partnering arrangements and sales of equity or debt securities. Adequate additional funding may not be available to the Company on acceptable terms or at all. The failure to raise capital as and when needed could have a negative impact on the Company s financial condition and ability to pursue business strategies. The Company may be required to delay, reduce the scope of or eliminate research and development programs, or obtain funds through arrangements with collaborators or others that may require the Company to relinquish rights to certain product candidates that the Company might otherwise seek to develop or commercialize independently.

2. Summary of Significant Accounting Policies

The following is a summary of significant accounting policies followed in the preparation of these financial statements.

Basis of Presentation

The accompanying consolidated financial statements include those of the Company and its subsidiary, Sage Securities Corporation, after elimination of all intercompany accounts and

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

transactions. The accompanying consolidated financial statements have been prepared in conformity with accounting principles generally accepted in the United States of America (GAAP).

Use of Estimates

The preparation of consolidated financial statements in conformity with GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and the disclosure of contingent assets and liabilities at the date of the financial statements and the reported amounts of expenses during the reporting period. Actual results could differ from those estimates.

Cash and Cash Equivalents

The Company considers all highly liquid investments with an original maturity of three months or less at the date of purchase to be cash equivalents.

Restricted Cash

A deposit of \$39 was restricted from withdrawal as of December 31, 2014 and 2013. The restriction is related to securing the Company's facility lease and expires in 2017 in accordance with the operating lease agreement. This balance is included in restricted cash on the accompanying balance sheets.

Property and Equipment

Property and equipment are recorded at cost and depreciated over their estimated useful lives using the straight-line method. Upon retirement or sale, the cost of assets disposed of and the related accumulated depreciation are removed from the accounts and any resulting gain or loss is credited or charged to income. Repairs and maintenance costs are expensed as incurred.

Impairment of Long-Lived Assets

Long-lived assets consist of property and equipment. Long-lived assets to be held and used are tested for recoverability whenever events or changes in business circumstances indicate that the carrying amount of the assets may not be fully recoverable. Factors that the Company considers in deciding when to perform an impairment review include significant underperformance of the business in relation to expectations, significant negative industry or economic trends, and significant changes or planned changes in the use of the assets. If an impairment review is performed to evaluate a long-lived asset for recoverability, the Company compares forecasts of undiscounted cash flows expected to result from the use and eventual disposition of the long-lived asset to its carrying value. An impairment loss would be recognized when estimated undiscounted future cash flows expected to result from the use of an asset are less than its carrying amount. The impairment loss would be based on the excess of the carrying value of the impaired asset over its fair value, determined based on discounted cash flows. To date, the Company has not recorded any impairment losses on long-lived assets.

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

Research and Development

Research and development expenses are comprised of costs incurred in performing research and development activities, including salaries and benefits, overhead costs, depreciation, contract services and other related costs. Research and development costs are expensed to operations as the related obligation is incurred.

Research Contract Costs and Accruals

The Company has entered into various research and development contracts with research institutions and other companies both inside and outside of the United States. These agreements are generally cancelable, and related payments are recorded as research and development expenses as incurred. The Company records accruals for estimated ongoing research costs. When evaluating the adequacy of the accrued liabilities, the Company analyzes progress of the studies, including the phase or completion of events, invoices received and contracted costs. Significant judgments and estimates may be made in determining the accrued balances at the end of any reporting period. Actual results could differ from the Company's estimates. The Company's historical accrual estimates have not been materially different from the actual costs.

Patent Costs

The Company expenses patent costs as incurred and classifies such costs as general and administrative expenses in the accompanying statements of operations and comprehensive loss.

Stock-Based Compensation

The Company recognizes compensation expense for all stock-based awards made to employees and nonemployee directors, including grants of stock options and restricted stock, based on estimated fair value on date of grant, over the requisite service period.

For stock options and restricted stock issued to nonemployee consultants, the Company recognizes the fair value of such instruments as an expense over the period in which the related services are received. The fair value of the awards and measurement of related stock-based compensation is subject to periodic adjustments as the underlying equity instruments vest.

The fair value of each option grant is estimated using the Black-Scholes option-pricing model. Through July 2014, the Company was a private company and lacks sufficient Company-specific historical and implied volatility information. Therefore, the Company estimates expected volatility based on the historical volatility of publicly traded peer companies and expects to continue to do so until such time as it has adequate historical data regarding the volatility of its traded stock price. The expected term of the Company's options has been determined utilizing the simplified method for awards that qualify as plain-vanilla options, while the expected term of its options granted to consultants and nonemployees has been determined based on the contractual term of the options. The risk-free interest rate is determined by reference to the U.S. Treasury yield curve in effect at the time of grant of the award for time periods approximately equal to the expected term of the award. Expected dividend yield is based on the fact that the Company never paid cash dividends and does not expect to pay any cash dividends in the foreseeable future.

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

The Company also applies a forfeiture rate in order to calculate stock-based compensation expense. To the extent actual forfeitures differ from the estimates, the difference will be recorded as a cumulative adjustment in the period the estimates are revised. The Company recognizes compensation expense for only the portion of awards that are expected to vest. Expected forfeitures are based on the Company's historical experience and management's expectations of future forfeitures.

Basic and Diluted Net Income (Loss) Per Share

Upon the closing of the Company's IPO in July 2014, all of the Company's outstanding redeemable convertible preferred shares were converted into shares of common stock. Prior to this conversion, the Company followed the two-class method when computing net income (loss) per share as the Company had issued shares that meet the definition of participating securities. The two-class method determines net income (loss) per share for each class of common and participating securities according to dividends declared or accumulated and participation rights in undistributed earnings. The two-class method requires income available to common shareholders for the period to be allocated between common and participating securities based upon their respective rights to receive dividends as if all income for the period had been distributed. The Company's redeemable convertible preferred shares contractually entitled the holders of such shares to participate in dividends, but did not contractually require the holders of such shares to participate in losses of the Company. Accordingly, the two-class method did not apply for periods in which the Company reported a net loss or a net loss attributable to common shareholders resulting from dividends or accretion related to its redeemable convertible preferred shares.

Basic net income (loss) per share attributable to common shareholders is computed by dividing the net income (loss) attributable to common shareholders by the weighted average number of common shares outstanding for the period. Diluted net income (loss) per share attributable to common shareholders is computed by dividing the diluted net income (loss) attributable to common shareholders by the weighted average number of common shares outstanding for the period, including potential dilutive common shares assuming the dilutive effect of outstanding stock options and unvested restricted common shares, as determined using the treasury stock method. For periods in which the Company has reported net losses, diluted net loss per common share attributable to common shareholders is the same as basic net loss per common share attributable to common shareholders, since dilutive common shares are not assumed to have been issued if their effect is antidilutive.

The Company reported a net loss attributable to common stockholders for the years ended December 31, 2014, 2013 and 2012.

Risks and Uncertainties

The product candidates developed by the Company require approvals from the U.S. Food and Drug Administration or foreign regulatory agencies prior to commercial sales. There can be no assurance that the Company's current and future product candidates will receive the necessary approvals. If the Company is denied approval or approval is delayed, it may have a material adverse impact on the Company's business and its financial statements.

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

The Company is subject to risks common to companies in the development stage including, but not limited to, dependency on the clinical and commercial success of its product candidates, ability to obtain regulatory approval of its product candidates, the need for substantial additional financing to achieve its goals, uncertainty of broad adoption of its approved products, if any, by physicians and consumers, significant competition and untested manufacturing capabilities.

Concentration of Credit Risk and of Significant Suppliers

Financial instruments that potentially expose the Company to concentrations of credit risk consist primarily of cash and cash equivalents. The Company has all cash and cash equivalents balances at two accredited financial institutions, in amounts that exceed federally insured limits. The Company does not believe that it is subject to unusual credit risk beyond the normal credit risk associated with commercial banking relationships.

The Company is dependent on third-party manufacturers to supply products for research and development activities in its programs. In particular, the Company relies and expects to continue to rely on a small number of manufacturers to supply it with its requirements for the active pharmaceutical ingredients and formulated drugs related to these programs. These programs could be adversely affected by a significant interruption in the supply of active pharmaceutical ingredients and formulated drugs.

Income Taxes

The Company accounts for income taxes under the asset and liability method. Under this method, deferred tax assets and liabilities are recognized for the estimated future tax consequences attributable to differences between financial statement carrying amounts of existing assets and liabilities and their respective tax bases. Deferred tax assets and liabilities are measured using enacted rates in effect for the year in which these temporary differences are expected to be recovered or settled. Valuation allowances are provided if based on the weight of available evidence, it is more likely than not that some or all of the deferred tax assets will not be realized.

Fair Value Measurements

Fair value is the price that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants at the measurement date. Financial assets and liabilities carried at fair value are to be classified and disclosed in one of the following three categories:

Level 1	Quoted market prices in active markets for identical assets or liabilities. At December 31, 2014 and 2013, the Company's Level 1 assets consisted of money market funds totaling \$127,766 and \$8,066, respectively.
Level 2	Observable inputs other than Level 1 prices, such as quoted prices for similar assets or liabilities; quoted prices in markets that are not active; or other inputs that are observable or can be corroborated by observable market data for substantially the full term of the assets or liabilities. At December 31, 2014 and 2013, the Company had no Level 2 assets or liabilities.
Level 3	Unobservable inputs that are supported by little or no market activity and that are significant to the fair value of the assets or liabilities. At December 31, 2014 and 2013, the Company had no Level 3 assets or liabilities.

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The Company's financial instruments generally consist of cash equivalents, accounts payable and accrued expenses. The carrying amounts for the applicable financial instruments reported in the balance sheets approximate their fair values at December 31, 2014 and 2013, respectively.

Deferred Offering Costs

The Company capitalizes certain legal, accounting and other third-party fees that are directly associated with in-process equity financings as other assets until such financings are consummated. After consummation of the equity financing in July 2014, \$2,285 of these costs were recorded in stockholders' equity (deficit) as a reduction of additional paid-in capital generated as a result of the initial public offering.

Segment Data

The Company manages its operations as a single segment for the purposes of assessing performance and making operating decisions. The Company's singular focus is on advancing medicines to treat central nervous system disorders, where there are inadequate or no approved existing therapies, including status epilepticus. All tangible assets are held within the U.S.

Comprehensive Loss

Comprehensive loss includes net loss as well as other changes in stockholders' equity (deficit) that result from transactions and economic events other than those with stockholders. For the years ended December 31, 2014, 2013 and 2012, there was no difference between net loss and comprehensive loss.

Government Grants

The Company records amounts received under grants as a reduction to research and development expense in the period it has incurred the expenditures in compliance with the specific restrictions of the grant. The Company recorded \$129, \$96 and \$0 for the years ended December 31, 2014, 2013 and 2012, respectively.

Initial Public Offering

On July 23, 2014, the Company completed the sale of 5,750,000 shares of its common stock in its initial public offering (the IPO), at a price to the public of \$18.00 per share, resulting in net proceeds to the Company of \$94.0 million after deducting underwriting discounts and commissions and offering costs paid by the Company. The shares began trading on Nasdaq Global Market on July 18, 2014.

In connection with preparing for the IPO, the Company's board of directors and stockholders approved a 1-for-3.15 reverse stock split of the Company's common stock effective July 2, 2014. All share and per share amounts in the financial statements contained herein and notes thereto have been retroactively adjusted, where necessary, to give effect to this reverse stock split. In connection with the closing of the IPO, all of the Company's outstanding redeemable convertible preferred stock automatically converted into shares of common stock as of July 23, 2014, resulting in the issuance by

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the Company of an additional 18,007,575 shares of common stock. The significant increase in common stock outstanding in July 2014 will impact the year-over-year comparability of the Company's net loss per share calculations over the next year.

Recently Issued Accounting Pronouncements

In May 2014, the Financial Accounting Standards Board, (FASB), issued Accounting Standards Update (ASU) 2014-09, Revenue from Contracts with Customers (Topic 606), (ASU 2014-09). ASU 2014-09 outlines a new, single comprehensive model for entities to use in accounting for revenue arising from contracts with customers and supersedes most current revenue recognition guidance, including industry-specific guidance. This new revenue recognition model provides a five-step analysis in determining when and how revenue is recognized. The new model will require revenue recognition to depict the transfer of promised goods or services to customers in an amount that reflects the consideration a company expects to receive in exchange for those goods or services. ASU 2014-09 is effective for public entities for annual reporting periods beginning after December 15, 2016 and interim periods within those periods. Early adoption is not permitted. Companies may use either a full retrospective or a modified retrospective approach to adopt ASU 2014-09. The Company is currently assessing the method of adoption and the impact this new accounting guidance will have on its financial statements and footnote disclosures.

In June 2014, the Financial Accounting Standard Board, or FASB, issued amended accounting guidance for development stage entities. The amendment eliminates certain financial reporting requirements for development stage entities, specifically, the presentation of inception-to-date information, the development stage entity label on the financial statements, the description of the activities in which the entity is engaged, and disclosure in the first year that the entity is no longer a development stage entity that it had been in prior years. The amendment is effective retrospectively for annual reporting periods beginning after December 15, 2014, and interim periods therein with early adoption permitted. The Company elected to early adopt this standard in the period ended June 30, 2014. Other than a change in presentation, the adoption of this guidance did not have an impact on the Company's financial statements.

In August 2014, the FASB issued ASU 2014-15, Presentation of Financial Statements - Going Concern (Subtopic 205-40). The new guidance addresses management's responsibility to evaluate whether there is substantial doubt about an entity's ability to continue as a going concern and to provide related footnote disclosures. Management's evaluation should be based on relevant conditions and events that are known and reasonably knowable at the date that the financial statements are issued. The standard will be effective for the first interim period within annual reporting periods beginning after December 15, 2016. Early adoption is permitted. The Company is evaluating the effect that this guidance will have on its financial statements.

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3. Balance Sheet Components***Property and Equipment, net***

Property and equipment, net consists of the following:

	Useful Life (Years)	December 31,	
		2014	2013
Computer and office equipment	3	\$ 206	\$ 78
Furniture and equipment	5	103	103
		309	181
Less: Accumulated depreciation		(146)	(95)
Property and equipment, net		\$ 163	\$ 86

Depreciation expense for the years ended December 31, 2014, 2013 and 2012 was \$51, \$47, and \$44, respectively.

Accrued Expenses

Accrued expenses consist of the following:

	December 31,	
	2014	2013
Employee related expenses	\$ 1,279	\$ 49
Development costs	2,788	57
Professional services	574	190
Other accrued expenses	46	31
	\$ 4,687	\$ 327

4. Commitments and Contingencies***Operating Leases***

The Company rents its 6,500 square foot office space under an operating lease that was executed in 2011 and expires in 2017. In March 2013, the Company entered into a second lease for an additional 4,100 square feet. The second lease, which commenced on August 26, 2013, has a term of 42 months and rent expense of \$9 per month. Also in March 2013, the Company signed a sublease agreement to sublet 1,900 square feet. The sublease is for a term of 42 months and rental income is \$4 per month.

Rent expense, net of sublease income, for the years ended December 31, 2014, 2013, and 2012 was \$302, \$274 and \$194, respectively.

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Future minimum lease payments, net of sublease income, under non-cancelable operating leases are as follows at December 31, 2014:

Years Ending December 31,	
2015	\$ 262
2016	268
2017	45
	\$ 575

License Agreements***CyDex License Agreement***

In October 2011, the Company entered into a research and development license with CyDex Pharmaceuticals, Inc. (CyDex) for the development of drug product using licensed technology for a period of one year. Under the terms of the license agreement, the Company paid an initial licensing fee of \$200 and an additional fee of \$100 for CyDex to perform research and development services to evaluate the licensed technology for formulation with the Company's developmental product.

The \$200 payment was recorded as research and development expense as the acquired technology was in-process research and development, and the \$100 payment was recorded to research and development expense in 2011 and 2012 as services were performed.

In December 2012, the Company exercised its option to enter into a commercial license and supply agreement for CyDex's proprietary technology and paid \$100 for the perpetual license, which was recorded as research and development expense.

In August 2013, the Company entered into a commercial license agreement as a result of which the December 2012 license was terminated and the December 2012 supply agreement was amended. Specifically, CyDex granted the Company an exclusive license to the CyDex technology for use in the fields of status epilepticus and traumatic brain injury. In exchange, the Company is required to pay upfront, milestone and royalty-based compensation. In addition, CyDex granted the Company a research license to Captisol for allopregnanolone for use in proof of concept studies. The August 2013 agreement will continue in effect unless and until terminated. In consideration for the amended license rights, the Company paid \$300. The Company is obligated to make milestone payments based on achievement of clinical development and regulatory milestones of \$900 and \$3,750, respectively. Also under this agreement, the Company is required to pay royalties in the low single digits based on levels of net sales.

Under the amended supply agreement with CyDex, the Company is required to purchase all of its supply of Captisol from CyDex and CyDex is required to supply the Company with Captisol, subject to certain limitations.

In April 2014, the Company amended its commercial license and supply agreements with CyDex to expand the fields of use to include the treatment, prevention or diagnosis of any disease or symptom in humans or animals. In consideration for the amended terms, the Company paid \$200 upfront and is

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obligated to make milestone payments, once per field, based on the achievement of clinical development and regulatory milestones for the development of SAGE-547 in the fields of status epilepticus and traumatic brain injury of \$750 and \$3,750, respectively. For the development in two additional fields, the Company is obligated to make milestone payments, once per field, based on the achievement of clinical development and regulatory milestones of \$1,250 and \$8,500, respectively.

As of December 31, 2014, the Company had not made any milestone or royalty payments.

Washington University License Agreement

In November 2013, the Company entered into a license agreement with Washington University whereby the Company was granted exclusive, worldwide rights to develop and commercialize a novel set of neuroactive steroids developed by Washington University. In exchange for development and commercialization rights, the Company paid an upfront, non-refundable payment of \$50 and is required to pay an annual license maintenance fee of \$15 on each subsequent anniversary date, until the first Phase 2 clinical study for a licensed product is initiated. The Company is obligated to make milestone payments based on achievement of clinical development and regulatory milestones of up to \$650 and \$500, respectively. Additionally, the Company fulfilled its obligation to issue Washington University 47,619 shares of common stock on December 13, 2013. The fair values of these shares totaling \$64 were recorded as research and development expense in 2013.

The Company is obligated to pay royalties of low single digits on net sales for licensed products covered under patent rights and royalties of low single digits on net sales for licensed products not covered under patent rights. Additionally, the Company has the right to sublicense and is required to make payments at varying percentages of sublicensing revenue received, initially in the mid-teens and descending to the mid-single digits over time.

As of December 31, 2014, the Company had not made any milestone or royalty payments.

University of California License Agreement

In October 2013, the Company entered into a non-exclusive license agreement with The Regents of the University of California whereby the Company was granted a non-exclusive license to certain clinical data and clinical material for use in the development and commercialization of biopharmaceutical products in the licensed field, including status epilepticus and post-partum depression. In May 2014, the license agreement was amended to add the treatment of essential tremor to the licensed field of use, materials and milestone fee provisions of the agreement.

The Company will be required to pay clinical development milestones of up to \$100 and pay royalties of less than 1% on net sales for a period of fifteen years following the sale of the first commercial product.

The license will terminate on the earlier to occur of (i) 27 years after the effective date or (ii) 15 years after the last-derived product is first commercially sold.

As of December 31, 2014, the Company had not made any milestone or royalty payments.

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Consulting Agreement

In January 2014, the Company entered into a consulting agreement with a nonemployee advisor whereby the Company is obligated to make cash payments of up to \$2,000 and to issue up to 126,984 shares of common stock upon attainment of certain clinical development and regulatory milestones.

In January and March 2014, the first milestone for each of two programs included in the consulting agreement were met. Accordingly, the Company made cash payments of \$50 and issued 15,872 shares of the Company's common stock. In connection with the shares of common stock issued, the Company recorded \$127 as research and development expense for the year ended December 31, 2014.

5. Redeemable Convertible Preferred Stock

As of December 31, 2014 and 2013, the Company's Certificate of Incorporation, as amended and restated, authorizes the Company to issue no shares and 37,750,000, respectively, shares of \$0.0001 par value preferred stock. In July 2014, all issued and outstanding redeemable convertible preferred stock was converted to common stock, see Note 2.

The Company had issued Series A, Series B and Series C redeemable convertible preferred stock (collectively, the Redeemable Preferred Stock). The Redeemable Preferred Stock was classified outside of stockholders' equity (deficit) as of December 31, 2013 because the shares contain redemption features that are not solely within the control of the Company.

On March 18, 2013, the Company issued an additional 5,000,000 shares in the second funding of the second tranche of Series A Preferred Stock at \$1.00 per share, resulting in net proceeds of \$4,996.

On July 1, 2013, the Company issued an additional 5,000,000 shares in the third funding of the second tranche of Series A Preferred Stock at \$1.00 per share, resulting in net proceeds of \$4,999.

On September 12, 2013, the Company issued an additional 12,500,000 shares in the third tranche of Series A Preferred Stock at \$1.00 per share, resulting in net proceeds of \$12,487.

On October 18, 2013, the Company issued 250,000 shares of Series A Preferred Stock at \$1.00 per share, resulting in net proceeds of \$250.

The Company incurred issuance costs of \$18 and \$6 in 2013 and 2012, respectively, with the issuance of the Series A Preferred Stock which were recorded as a reduction of the proceeds received. These costs were accreted on a straight-line basis to the carrying value of preferred stock, beginning with the date of issue to the date of earliest redemption.

On October 15, 2013, the Company entered into a Stock Purchase Agreement whereby the Company would issue up to \$20,000 of Series B redeemable convertible preferred stock (Series B Preferred Stock) at \$1.50 per share. The initial purchase and sale in the amount of \$10,000 could have occurred once certain development milestones had been successfully achieved. The second tranche of \$10,000 could be issued after the initial closing and at the discretion of the Board of Directors. In November 2013, the Company met the development milestones to issue the first tranche of the Series B Preferred Stock.

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On January 7, 2014, the Company issued 6,666,666 shares of Series B Preferred Stock at \$1.50 per share, resulting in net proceeds of \$9,995.

On February 12, 2014, the Company issued 3,333,333 shares of Series B Preferred Stock at \$1.50 in a second closing, resulting in net proceeds of \$5,000. At that time, the Company decided not to draw on the remaining \$5,000 of the second tranche of the Series B Preferred Stock.

The Company incurred issuance costs of \$30 in 2014 with the issuance of the Series B Preferred Stock which were recorded as a reduction of the proceeds received.

On March 11, 2014, the Company entered into a Stock Purchase Agreement whereby the Company issued 8,973,905 shares of Series C redeemable convertible preferred stock (Series C Preferred Stock) at \$4.2345 per share for net proceeds of \$37,981.

The Company incurred issuance costs of \$110 in 2014 with the issuance of the Series C Preferred Stock which were recorded as a reduction of the proceeds received.

The holders of the Redeemable Preferred Stock had the following rights and preferences:

Voting Rights

The holders of Redeemable Preferred Stock were entitled to vote, together with the holders of common stock, on all matters submitted to stockholders for a vote. Each preferred stockholder was entitled to the number of votes equal to the number of shares of common stock into which each share of Redeemable Preferred Stock was convertible at the time of such vote.

Dividends

The holders of Series A, Series B and Series C Preferred Stock were entitled to receive dividends in preference to any dividend on common stock at the rate of \$0.08, \$0.12 and \$0.34, respectively, per share per annum. Dividends were payable only when, as, and if declared by the Board of Directors. As of December 31, 2014 and 2013, no dividends had been declared or paid by the Company.

Liquidation

In the event of any liquidation, dissolution or winding up of the affairs of the Company, the holders of the Series A, Series B and Series C Preferred Stock were entitled to receive an amount per share equal to the original issue price of \$1.00, \$1.50 and \$4.2345, respectively, per share (the Original Issue Price), plus all accruing dividends, whether or not declared, payable in preference and priority to any payments made to the holders of the then outstanding common stock. In the event of a liquidation, dissolution or winding up of the affairs of the Company, holders of Series C Preferred Stock were to be paid their liquidation preference amounts prior to the payment to holders of Series A Preferred Stock and Series B Preferred Stock of their liquidation preference amounts on a pari passu basis. Series A Preferred Stock and Series B Preferred Stock were to be paid their liquidation preference amounts on a pari passu basis prior to the payment of any amounts to holders of common stock. If the liquidation proceeds exceeded the liquidation preferences, then holders of the Series A, Series B and Series C Preferred Stock were to participate in the excess on an as-if converted basis with the common shareholders up to \$2.50, \$3.75 and \$10.58, respectively, per share.

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Redemption Rights

Redeemable Preferred Stock was redeemable at the option of the preferred stockholders on or after September 30, 2020. If the holders of at least seventy-five percent of the then outstanding shares of Redeemable Preferred Stock exercised their redemption rights, the Company must notify all preferred stockholders of the election to exercise redemption rights. Under the terms of the Company's Certificate of Incorporation, as amended and restated on March 11, 2014, the holders of the Redeemable Preferred Stock who requested redemption of their Redeemable Preferred Stock were entitled to receive an amount per share equal to the Original Issue Price of \$1.00, \$1.50 or \$4.2345 for each share of Series A, Series B or Series C Preferred Stock, respectively, plus all accruing dividends, whether or not declared, and were to be paid in three annual installments commencing not more than sixty days after receipt of notification by the Company.

If the Company did not have sufficient funds legally available to redeem all shares of Redeemable Preferred Stock to be redeemed at the redemption date, the Company would redeem such shares ratably to the extent possible and would redeem the remaining shares as soon as sufficient funds are legally available.

Conversion

Each share of Redeemable Preferred Stock was convertible at any time at the option of the shareholder into fully paid and nonassessable shares of common stock determined by dividing the Original Issue Price by the Conversion Price in effect at the time of conversion. The initial Series A, Series B and Series C Conversion Price is \$3.15, \$4.725 and \$13.338675, respectively, per share (the Conversion Price).

In addition to the above optional conversion feature, the Redeemable Preferred Stock included a mandatory conversion feature whereby upon either of the following events, all outstanding shares of Redeemable Preferred Stock would automatically be converted into shares of common stock at the then-effective conversion ratio: (i) Initial Public Offering resulting in a closing price of at least \$14.18 per share that results in at least \$30,000 in gross proceeds to the Company, or (ii) the date and time, or the occurrence of an event, specified by vote or written consent of the holders of at least 75% of the then outstanding shares of Redeemable Preferred Stock. All shares that are required to be surrendered per the provisions above will be deemed to have been retired and canceled and may not be reissued as shares of Redeemable Preferred Stock.

The Company has newly authorized preferred stock amounting to 5,000,000 shares as of December 31, 2014. The newly authorized preferred stock was classified under stockholders' equity (deficit) as of December 31, 2014.

6. Common Stock

As of December 31, 2014 and 2013, the Company has authorized 120,000,000 and 66,000,000 shares, respectively, of common stock with a par value of \$0.0001 per share.

Each share of common stock entitles the holder to one vote on all matters submitted to a vote of the Company's stockholders. Common stockholders are entitled to receive dividends, as may be

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declared by the Board of Directors, if any, subject to the preferential dividend rights of the Series A, Series B and Series C preferred stockholders. As of December 31, 2014 and 2013, no dividends have been declared.

7. Stock-Based Compensation***2014 Stock Option Plan***

On July 2, 2014, the Company's stockholders approved the 2014 Stock Option and Incentive Plan (the "2014 Stock Option Plan"), which became effective upon the completion of the IPO. The 2014 Stock Option Plan provides for the grant of restricted stock awards, incentive stock options and non-statutory stock options. The 2014 Stock Option Plan replaced the Company's 2011 Stock Option and Grant Plan (the "2011 Stock Option Plan"). The Company will grant no further stock options or other awards under the 2011 Stock Option Plan. Any options or awards outstanding under the 2011 Stock Option Plan remained outstanding and effective. As of December 31, 2014, the total number of shares reserved under all equity plans is 3,505,868, and the Company had 1,509,253 shares available for future issuance under such plans.

The 2014 Stock Option Plan provides for an annual increase, to be added on the first day of each fiscal year, by up to 4% of the Company's issued and outstanding shares of common stock on the immediately preceding December 31.

2014 Employee Stock Purchase Plan

On July 2, 2014, the Company's stockholders approved the 2014 Employee Stock Purchase Plan. A total of 282,000 shares of common stock were initially authorized for issuance under this plan. The 2014 Employee Stock Purchase Plan became effective upon the completion of the IPO. As of December 31, 2014, no shares have been issued under this plan.

Terms of restricted stock awards and stock option agreements, including vesting requirements, are determined by the Board of Directors, subject to the provisions of the applicable stock option plan. Options and restricted stock awards granted by the Company generally vest based on the grantee's continued service with the Company during a specified period following grant. Awards generally vest ratably over four years, with a 25% cliff vesting at the one year anniversary for new employee awards. During 2013, the Company also granted a pool of option awards which vest ratably over one year. All awards are exercisable from the date of grant for a period of ten years.

The stock-based compensation expense recognized during the years ended December 31, 2014, 2013, and 2012 was as follows:

	Year Ended December 31,		
	2014	2013	2012
Stock compensation expense:			
Research and development	\$ 1,093	\$ 38	\$
General and administrative	1,419	23	
	\$ 2,512	\$ 61	\$

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During the year ended 2012, the Company did not record stock-based compensation expense as the amounts were inconsequential.

Prior to December 31, 2012, the estimated fair market value of the Company's common stock was determined solely by the Board of Directors on the date of grant. From December 31, 2012 until its IPO, the Company secured a third-party valuation to assist the Board of Directors in the determination of the estimated fair market value of the Company's common stock.

For stock option awards, the fair value of the options is estimated at the grant date using the Black-Scholes option-pricing model, taking into account the terms and conditions upon which options are granted. The fair value of the options is amortized on a straight-line basis over the requisite service period of the awards. The weighted average grant date fair value per share relating to outstanding stock options granted under the Company's stock option plans during the years ended December 31, 2014 and 2013 was \$14.33 and \$0.38, respectively.

The fair value of each option granted to employees and directors during the years ended December 31, 2014, 2013, and 2012 under the Company's stock option plans has been calculated on the date of grant using the following weighted average assumptions:

	Year Ended December 31,		
	2014	2013	2012
Expected dividend yield	0.00%	0.00%	0.00%
Expected volatility	98.86%	99.89%	0.00%
Risk free interest rate	1.95%	1.66%	0.00%
Expected term	6.38 years	6.04 years	

Expected dividend yield: The Company has not paid and does not anticipate paying any dividends in the foreseeable future.

Risk-free interest rate: The Company determined the risk-free interest rate by using a weighted average equivalent to the expected term based on the U.S. Treasury yield curve in effect as of the date of grant.

Expected volatility: As the Company has only been a public company since July 2014, there is not sufficient historical volatility for the expected term of the options. Therefore, the Company used an average historical share price volatility based on an analysis of reported data for a peer group of comparable companies.

Expected term (in years): Expected term represents the period that the Company's share option grants are expected to be outstanding. As the Company has only been a public company since July 2014, there is not sufficient historical term data to calculate the expected term of the options. Therefore, the Company elected to utilize the simplified method to estimate the expected term of option grants issued to employees. Under this approach, the weighted average expected life is presumed to be the average of the vesting term and the contractual term of the option.

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Forfeitures are estimated at the time of grant and revised, if necessary, in subsequent periods if actual forfeitures differ from estimates. The Company estimates forfeitures based on historical termination behavior. For the years ended December 31, 2014, 2013 and 2012, a forfeiture rate of 10% was applied.

For options granted to nonemployees, the expected life of the option used is ten years, which is the contractual term of each such option. All other assumptions used to calculate the grant date fair value are generally consistent with the assumptions used for options granted to employees.

The table below summarizes activity related to stock options:

	Shares	Weighted Average Exercise Price \$	Weighted Average Remaining Life (in years)	Aggregate Intrinsic Value \$
Outstanding as of December 31, 2011		\$		
Granted	5,555	0.04		
Exercised	(5,555)	0.04		
Forfeited				
Outstanding as of December 31, 2012		\$		
Granted	1,206,655	0.51		
Exercised	(3,174)	0.45		
Forfeited				
Outstanding as of December 31, 2013	1,203,481	\$ 0.51	9.60	\$ 1,038
Granted	942,513	14.34		
Exercised	(87,476)	0.46		
Forfeited	(61,903)	1.36		
Outstanding as of December 31, 2014	1,996,615	\$ 7.01	8.98	\$ 59,362
Vested or expected to vest as of December 31, 2014	1,782,257	\$ 6.61	8.96	\$ 53,670
Exercisable as of December 31, 2014	383,032	\$ 0.82	8.66	\$ 13,705

As of December 31, 2014, the Company had unrecognized stock-based compensation expense related to its unvested stock option awards of \$9,705, which is expected to be recognized over the remaining weighted average vesting period of 3.24 years. The total fair value of shares vested for the years ended December 31, 2014, 2013 and 2012 was \$988, \$9, and \$1, respectively. During the year ended December 31, 2014, stock option exercises resulted in proceeds of \$40, and during the years ended December 31, 2013 and 2012 stock option exercises resulted in the proceeds of less than \$1. The intrinsic value of stock options exercised during the year ended December 31, 2014 was \$2,446 and the intrinsic value of stock options exercised during the years ended December 31, 2013 and 2012 was zero.

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Restricted Stock Awards

During the years ended December 31, 2014, 2013 and 2012, the Company granted restricted stock awards to certain officers, employees, directors, and consultants of the Company. During the years ended December 31, 2014 and 2013, the Company recorded \$213 and \$11, respectively, of stock-based compensation expense related to its restricted stock. There was no stock-based compensation expense for the year ended December 31, 2012. The table below summarizes activity relating to restricted stock:

	Shares	Weighted Average Grant Date Fair Value Per Share
Outstanding as of December 31, 2011	613,086	
Issued	561,104	\$ 0.04
Vested	(709,158)	
Forfeited		
Repurchased	(102,777)	
Outstanding as of December 31, 2012	362,255	
Issued	130,158	\$ 0.45
Vested	(176,695)	
Forfeited		
Repurchased		
Outstanding as of December 31, 2013	315,718	
Issued		
Vested	(138,108)	
Forfeited		
Repurchased	(6,778)	
Outstanding as of December 31, 2014	170,832	

As of December 31, 2014, the Company had unrecognized stock-based compensation expense related to its unvested restricted stock awards of \$340, which is expected to be recognized over the remaining weighted average vesting period of 1.15 years.

During the year ended December 31, 2014, no shares of restricted stock were issued. During the years ended December 31, 2013, and 2012, current and former employees of the Company purchased a total of 130,158 and 561,104 shares of restricted stock, respectively, resulting in the proceeds of \$50 and \$19, respectively.

Unvested shares are subject to repurchase by the Company, at the issuance price, upon the employee's termination at the Company's sole discretion. In 2012, the Company repurchased 102,777 shares of restricted common stock issued to employees with a value of \$3 in conjunction with the employees' termination from the Company, and in the year ended December 31, 2014, the Company repurchased 6,778 shares of restricted common stock issued to employees at their \$0.04 original purchase price per share.

Table of Contents**SAGE THERAPEUTICS, INC. AND SUBSIDIARY****Notes to Consolidated Financial Statements**

(Amounts in thousands, except share and per share data)

8. Net Loss Per Share

Basic and diluted net income (loss) per share attributable to common stockholders was calculated as follows for the years ended December 31, 2014, 2013 and 2012:

	Year Ended December 31,		
	2014	2013	2012
Basic net income (loss) per share attributable to common stockholders:			
Numerator:			
Net loss	\$ (36,105)	\$ (18,288)	\$ (9,636)
Denominator:			
Weighted average common shares outstanding basic	21,574,347	1,492,288	1,118,288
Dilutive effect of common share equivalents resulting from common share options and preferred common shares (as converted)			
Weighted average common shares outstanding diluted	21,574,347	1,492,288	1,118,288
Net loss per share attributable to common stockholders basic and diluted	\$ (1.67)	\$ (12.26)	\$ (8.62)

The following common stock equivalents outstanding as of December 31, 2014 and 2013, were excluded from the computation of diluted net loss per share attributable to common stockholders for the periods presented because including them would have been anti-dilutive:

	Year Ended December 31,	
	2014	2013
Options to purchase common stock	1,621,906	766,156
Restricted stock	170,067	284,129
Redeemable convertible preferred stock (presented on a weighted average basis)		8,022,175
	1,791,973	9,072,460

9. Income Taxes

There is no provision for income taxes because the Company has historically incurred operating losses and maintains a full valuation allowance against its net deferred tax assets. The reported amount of income tax expense for the years differs from the amount that would result from applying domestic federal statutory tax rates to pretax losses primarily because of changes in valuation allowance.

Table of Contents**SAGE THERAPEUTICS, INC. AND SUBSIDIARY****Notes to Consolidated Financial Statements**

(Amounts in thousands, except share and per share data)

A reconciliation of U.S. statutory rate to the Company's effective tax rate is as follows:

	Year Ended December 31,		
	2014	2013	2012
Tax due at statutory rate	34.0%	34.0%	34.0%
State taxes, net of federal	4.5	5.2	5.2
Permanent items	(1.0)	(0.1)	(0.1)
Federal and state credits	8.5	1.5	0.8
Change in valuation allowance	(46.0)	(40.5)	(39.9)
Other		(0.1)	
	0.0%	0.0%	0.0%

Significant components of the Company's net deferred tax asset at December 31, 2014 and 2013 are as follows:

	December 31,	
	2014	2013
Net operating losses	\$ 21,907	\$ 9,401
Capitalized start-up costs	2,514	2,694
Accounting method change	(2,020)	
Tax credit carryforwards	4,433	369
Accrued expenses	494	15
Depreciation and amortization	332	239
Stock options	623	
Others	20	21
Total net deferred tax asset before valuation allowance	28,303	12,739
Valuation allowance	(28,303)	(12,739)
Net deferred tax asset	\$	\$

As of December 31, 2014, the Company had federal and state net operating loss carryforwards of \$55,821 and \$55,443, respectively, which begin to expire in 2031. As of December 31, 2014, the Company had federal and state research and development tax credits carryforwards of \$663 and \$326, respectively, which begin to expire in 2031 and 2027, respectively. As of December 31, 2014, the Company had federal orphan drug tax credit carry forwards of \$3,555, which begin to expire in 2034.

As of December 31, 2014, net deferred tax assets increased approximately \$15,564 primarily due to the operating loss and tax credits incurred during the year. This increase in net deferred tax assets was offset by a corresponding increase in the valuation allowance.

Management of the Company has evaluated the positive and negative evidence bearing upon the realizability of its deferred tax assets, which are comprised principally of net operating loss carryforwards and capitalized start-up costs. Under the applicable accounting standards, management has considered the Company's history of losses and concluded that it is more likely than not that the Company will not recognize the benefits of federal and state deferred tax assets. Accordingly, a full valuation allowance of \$28,303 and \$12,739 has been established at December 31, 2014

and 2013, respectively.

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Table of Contents**SAGE THERAPEUTICS, INC. AND SUBSIDIARY****Notes to Consolidated Financial Statements**

(Amounts in thousands, except share and per share data)

Pursuant to Section 382 of the Internal Revenue Code, certain substantial changes in the Company's ownership may result in a limitation on the amount of net operating loss carryforwards and tax carryforwards that may be used in future years. Utilization of the net operating loss (NOL) and tax credit carryforwards may be subject to a substantial annual limitation under Section 382 of the Internal Revenue Code of 1986 due to ownership change limitations that have occurred previously or that could occur in the future. These ownership changes may limit the amount of NOL and tax credit carryforwards that can be utilized annually to offset future taxable income and tax, respectively. The Company has not completed a study to assess whether an ownership change has occurred, or whether there have been multiple ownership changes since its formation, due to significant complexity and related costs associated with such a study. There could also be additional ownership changes in the future which may result in additional limitations on the utilization of NOL carryforwards and credits.

The Company applies the authoritative guidance on accounting for and disclosure of uncertainty in tax positions, which requires the Company to determine whether a tax position of the Company is more likely than not to be sustained upon examination, including resolution of any related appeals of litigation processes, based on the technical merits of the position. For tax positions meeting the more likely than not threshold, the tax amount recognized in the financial statements is reduced by the largest benefit that has a greater than fifty percent likelihood of being realized upon the ultimate settlement with the relevant taxing authority.

The following is a rollforward of the Company's unrecognized tax benefits:

	Year Ended December 31,		
	2014	2013	2012
Unrecognized tax benefits as of the beginning of the year	\$ 2,880	\$ 1,477	\$ 504
Gross increases current period tax positions		1,403	
Gross decreases tax positions of prior periods	(2,880)		973
Unrecognized tax benefits as of the end of the year	\$	\$ 2,880	\$ 1,477

During 2014, the Company filed an application for change in accounting method with the IRS to capitalize start-up costs that were historically deducted and included as part of the NOL carryforward through December 31, 2013. As a result, the Company's unrecognized tax benefits, which historically related to start-up costs, are zero at December 31, 2014.

The Company will recognize interest and penalties related to uncertain tax positions in income tax expense when in a taxable income position. As of December 31, 2014 and 2013, the Company had no accrued interest or penalties related to uncertain tax positions and no amounts have been recognized in the Company's statement of operations.

The Company files tax returns as prescribed by the tax laws of the jurisdictions in which it operates. In the normal course of business, the Company is subject to examination by federal and state jurisdictions, where applicable. There are currently no pending tax examinations, and the Company's tax returns are open under statute from 2011 to the present. The Company's policy is to record interest and penalties related to income taxes as part of the tax provision.

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SAGE THERAPEUTICS, INC. AND SUBSIDIARY

Notes to Consolidated Financial Statements

(Amounts in thousands, except share and per share data)

10. Employee Benefit Plan

The Company maintains a 401(k) profit sharing plan (the Plan) for its employees. Each participant in the Plan may elect to contribute a portion of his or her annual compensation to the Plan subject to annual limits established by the Internal Revenue Service. Effective November 1, 2014, the Company instituted an employer match of 50% of eligible contributions up to 6% of employee contributions. For the year ended December 31, 2014, the Company contributed \$15.

11. Related Party Transactions

Since inception, the Company has received consulting and management services from Third Rock Ventures LLC, which through its affiliates, has a controlling interest in the Company and owns 45.3% of common stock at December 31, 2014. The Company paid Third Rock Ventures LLC \$282, \$682 and \$994 for these services for the years ended December 31, 2014, 2013 and 2012, respectively. At December 31, 2014, the Company owed Third Rock Ventures LLC \$5, which is included in accrued expenses. At December 31, 2013 and 2012, the Company owed Third Rock Ventures LLC \$125 and \$209, respectively, which is included in accounts payable.

12. Subsequent Events

Effective January 2015, the Company signed an agreement to sublet approximately 2,700 feet of office space. The sublease term is through July 2017 and the Company will pay \$9 per month.

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\$100,000,000

Common Stock

Preliminary Prospectus

J.P. Morgan

Goldman, Sachs & Co.

Leerink Partners

Cowen & Company

, 2015

Until , 2015, all dealers that effect transactions in these securities, whether or not participating in this offering, may be required to deliver a prospectus. This is in addition to the dealers' obligation to deliver a prospectus when acting as underwriters and with respect to their unsold allotments or subscriptions.

Table of Contents**PART II****Information Not Required in Prospectus****Item 13. Other expenses of issuance and distribution**

The following table sets forth all expenses, other than the underwriting discounts and commissions, payable by Sage Therapeutics, Inc. (the Company or the Registrant) in connection with the sale of the common stock being registered. All the amounts shown are estimates except the SEC registration fee and the FINRA filing fee.

	Amount
SEC registration fee	\$ 13,363
FINRA filing fee	17,750
Blue sky qualification fees and expenses	10,000
Printing and engraving expenses	110,000
Legal fees and expenses	125,000
Accounting fees and expenses	150,000
Transfer agent and registrar fees	10,000
Miscellaneous	63,887
Total	\$ 500,000

Item 14. Indemnification of directors and officers

Section 145 of the Delaware General Corporation Law permits a corporation to include in its charter documents, and in agreements between the corporation and its directors and officers, provisions expanding the scope of indemnification beyond that specifically provided by the current law.

Section 145(a) of the Delaware General Corporation Law provides, in general, that a corporation may indemnify any person who was or is a party or is threatened to be made a party to any threatened, pending or completed action, suit or proceeding, whether civil, criminal, administrative or investigative (other than an action by or in the right of the corporation), because he or she is or was a director, officer, employee or agent of the corporation, or is or was serving at the request of the corporation as a director, officer, employee or agent of another corporation, partnership, joint venture, trust or other enterprise, against expenses (including attorneys' fees), judgments, fines and amounts paid in settlement actually and reasonably incurred by the person in connection with such action, suit or proceeding, if he or she acted in good faith and in a manner he or she reasonably believed to be in or not opposed to the best interests of the corporation and, with respect to any criminal action or proceeding, had no reasonable cause to believe his or her conduct was unlawful.

Section 145(b) of the Delaware General Corporation Law provides, in general, that a corporation may indemnify any person who was or is a party or is threatened to be made a party to any threatened, pending or completed action or suit by or in the right of the corporation to procure a judgment in its favor because the person is or was a director, officer, employee or agent of the corporation, or is or was serving at the request of the corporation as a director, officer, employee or agent of another corporation, partnership, joint venture, trust or other enterprise, against expenses (including attorneys' fees) actually and reasonably incurred by the person in connection with the defense or settlement of such action or suit if he or she acted in good faith and in a manner he or she reasonably believed to be in or not opposed to the best interests of the corporation, except that no indemnification shall be made with respect to any claim, issue or matter as to which he or she shall have been adjudged to be liable to the corporation unless and only to the extent that the Court of Chancery or other adjudicating court

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determines that, despite the adjudication of liability but in view of all of the circumstances of the case, he or she is fairly and reasonably entitled to indemnity for such expenses which the Court of Chancery or other adjudicating court shall deem proper.

Section 145(g) of the Delaware General Corporation Law provides, in general, that a corporation may purchase and maintain insurance on behalf of any person who is or was a director, officer, employee or agent of the corporation, or is or was serving at the request of the corporation as a director, officer, employee or agent of another corporation, partnership, joint venture, trust or other enterprise against any liability asserted against such person and incurred by such person in any such capacity, or arising out of his or her status as such, whether or not the corporation would have the power to indemnify the person against such liability under Section 145 of the Delaware General Corporation Law.

The Company's amended and restated certificate of incorporation provides for the indemnification of directors to the fullest extent permissible under Delaware law.

The Company's amended and restated by-laws provides for the indemnification of officers, directors and third parties acting on the Company's behalf if such persons act in good faith and in a manner reasonably believed to be in and not opposed to the Company's best interest, and, with respect to any criminal action or proceeding, such indemnified party had no reason to believe his or her conduct was unlawful.

The Company has entered into indemnification agreements with each of its directors and executive officers, in addition to the indemnification provisions provided for in its charter documents, and the Company intends to enter into indemnification agreements with any new directors and executive officers in the future. These agreements will provide that we will indemnify each of its directors and executive officers, and such entities to the fullest extent permitted by law.

The underwriting agreement (to be filed as Exhibit 1.1 hereto) will provide for indemnification by the underwriters of the Company, and its executive officers and directors, and indemnification of the underwriters by the Company for certain liabilities, including liabilities arising under the Securities Act of 1933, as amended, in connection with matters specifically provided in writing by the underwriters for inclusion in the registration statement.

The Company carries insurance on behalf of any person who is or was a director or officer against any loss arising from any claim asserted against him or her and incurred by him or her in that capacity, subject to certain exclusions and limits of the amount of coverage.

Item 15. Recent sales of unregistered securities

In the three years preceding the filing of this registration statement, we have issued the following securities that were not registered under the Securities Act:

Issuances of capital stock

On September 30, 2011, we issued 6,000,000 shares of our Series A redeemable convertible preferred stock to one investor for an aggregate consideration of \$6,000,000. On April 9, 2012, we issued 4,000,000 shares of our Series A redeemable convertible preferred stock to one investor for \$4,000,000. On November 12, 2012, we issued 5,000,000 shares of our Series A redeemable convertible preferred stock to one investor for \$5,000,000. On March 18, 2013, we issued 5,000,000 shares of our Series A redeemable convertible preferred stock to one investor for \$5,000,000. On July 1, 2013, we issued 5,000,000 shares of our Series A redeemable convertible preferred stock to

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one investor for \$5,000,000. On September 12, 2013, we issued 12,500,000 shares of our Series A redeemable convertible preferred stock to two investors for \$12,500,000. On October 18, 2013, we issued 250,000 shares of our Series A redeemable convertible preferred stock to one investor for \$250,000.

On December 13, 2013, we issued 47,619 shares of our common stock in connection with entering into a license agreement.

On January 7, 2014, we issued 6,666,666 shares of our Series B redeemable convertible preferred stock to two investors for aggregate consideration of \$10,000,000. On February 12, 2014, we issued an aggregate of 3,333,333 shares of our Series B redeemable convertible preferred stock to two investors for \$5,000,000.

On January 24, 2014, we issued 7,936 shares of our common stock to a nonemployee advisor upon attainment of certain clinical milestones.

On March 11, 2014, we issued 8,973,905 shares of our Series C redeemable convertible preferred stock to 13 investors for aggregate consideration of \$38,000,000.

On March 26, 2014, we issued 7,936 shares of our common stock to a nonemployee advisor upon attainment of certain clinical milestones.

No underwriters were used in the foregoing transactions. All sales of securities described above were made in reliance upon the exemption from registration provided by Section 4(2) of the Securities Act (and/or Regulation D promulgated thereunder) for transactions by an issuer not involving a public offering. All of the foregoing securities are deemed restricted securities for the purposes of the Securities Act.

Grants of stock options and restricted stock

Since January 1, 2012 through the filing of our registration statement on Form S-8 on July 18, 2014, we granted stock options to purchase an aggregate of 1,893,150 shares of our common stock, with exercise prices ranging from \$0.04 to \$8.92 per share, to employees, directors and consultants pursuant to our stock option plan. Since January 1, 2012 through the filing of our registration statement on Form S-8 on July 18, 2014, we have granted an aggregate of 691,262 shares of restricted stock. The issuances of these securities were exempt either pursuant to Rule 701, as a transaction pursuant to a compensatory benefit plan, or pursuant to Section 4(a)(2), as a transaction by an issuer not involving a public offering. Since July 18, 2014, we have not granted stock options or shares of restricted stock that were not registered under our registration statement on Form S-8 on July 18, 2014.

Item 16. Exhibits and financial statement schedules

(a) Exhibits.

The exhibits to the registration statement are listed in the Exhibit Index to this registration statement and are incorporated herein by reference.

(b) Financial statement schedules.

None.

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Item 17. Undertakings

Insofar as indemnification for liabilities arising under the Securities Act of 1933, as amended, may be permitted to directors, officers and controlling persons of the Registrant pursuant to the foregoing provisions, or otherwise, the Registrant has been advised that in the opinion of the Securities and Exchange Commission such indemnification is against public policy as expressed in the Securities Act and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the Registrant of expenses incurred or paid by a director, officer or controlling person of the Registrant in the successful defense of any action, suit or proceeding) is asserted by such director, officer or controlling person in connection with the securities being registered, the Registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question whether such indemnification by it is against public policy as expressed in the Securities Act of 1933 and will be governed by the final adjudication of such issue.

The undersigned Registrant hereby undertakes that:

- (a) The undersigned Registrant will provide to the underwriters at the closing specified in the underwriting agreement certificates in such denominations and registered in such names as required by the underwriters to permit prompt delivery to each purchaser.
- (b) For purposes of determining any liability under the Securities Act, the information omitted from the form of prospectus filed as part of this registration statement in reliance upon Rule 430A and contained in a form of prospectus filed by the Registrant pursuant to Rule 424(b)(1) or (4) or 497(h) under the Securities Act shall be deemed to be part of this registration statement as of the time it was declared effective.
- (c) For the purpose of determining any liability under the Securities Act, each post-effective amendment that contains a form of prospectus shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof.

Table of Contents**SIGNATURES**

Pursuant to the requirements of the Securities Act of 1933, as amended, the Registrant has duly caused this Registration Statement on Form S-1 to be signed on its behalf by the undersigned, thereunto duly authorized, in the City of Cambridge, Commonwealth of Massachusetts, on the 6th day of April, 2015.

SAGE THERAPEUTICS, INC.

By: /s/ Jeffrey M. Jonas
 Jeffrey M. Jonas, M.D.
Chief Executive Officer, President and Director
(Principal Executive Officer)

POWER OF ATTORNEY

Each person whose individual signature appears below hereby authorizes and appoints Jeffrey M. Jonas, M.D. and Kimi Iguchi and each of them, with full power of substitution and resubstitution and full power to act without the other, as his or her true and lawful attorney in fact and agent to act in his or her name, place and stead and to execute in the name and on behalf of each person, individually and in each capacity stated below, and to file any and all amendments to this Registration Statement, including any and all post effective amendments and amendments thereto, and any registration statement relating to the same offering as this Registration Statement that is to be effective upon filing pursuant to Rule 462(b) under the Securities Act of 1933, as amended, and to file the same, with all exhibits thereto, and other documents in connection therewith, with the Securities and Exchange Commission, granting unto said attorneys in fact and agents, and each of them, full power and authority to do and perform each and every act and thing, ratifying and confirming all that said attorneys in fact and agents or any of them or their or his substitute or substitutes may lawfully do or cause to be done by virtue thereof.

Pursuant to the requirements of the Securities Act of 1933, as amended, this Registration Statement has been signed by the following persons in the capacities indicated below on the 6th day of April, 2015.

Signature	Title	Date
/s/ Jeffrey M. Jonas Jeffrey M. Jonas, M.D.	Chief Executive Officer, President and Director (Principal Executive Officer)	April 6, 2015
/s/ Kimi Iguchi Kimi Iguchi	Chief Financial Officer (Principal Financial and Accounting Officer)	April 6, 2015
/s/ Robert T. Nelsen Robert T. Nelsen	Director	April 6, 2015
/s/ Steven Paul Steven Paul, M.D.	Director	April 6, 2015
/s/ Kevin P. Starr Kevin P. Starr	Director	April 6, 2015
/s/ Howard Pien	Director	April 6, 2015

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Howard Pien		
/s/ James Frates	Director	April 6, 2015
James Frates		
/s/ Michael F. Cola	Director	April 6, 2015
Michael F. Cola		

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Exhibit No.	Description
1.1*	Form of Underwriting Agreement
3.1	Fifth Amended and Restated Certificate of Incorporation of the Registrant, as currently in effect (incorporated by reference to Exhibit 3.2 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
3.2	By-laws of the Registrant and the amendments thereto, as currently in effect (incorporated by reference to Exhibit 3.4 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
4.1	Specimen Common Stock Certificate (incorporated by reference to Exhibit 4.1 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
4.2	Second Amended and Restated Investors' Rights Agreement by and among the Registrant and certain of its stockholders dated March 11, 2014 (incorporated by reference to Exhibit 4.2 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
5.1*	Opinion of Goodwin Procter LLP
10.1#	2014 Stock Option and Incentive Plan and forms of award agreements thereunder (incorporated by reference to Exhibit 10.1 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.2	Exclusive License Agreement by and between the Registrant and Washington University, dated November 11, 2013 (incorporated by reference to Exhibit 10.3 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.3	Commercial License Agreement by and between the Registrant and CyDex Pharmaceuticals, Inc., dated August 21, 2013, as amended April 30, 2014 (incorporated by reference to Exhibit 10.4 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.4	Non-Exclusive License Agreement by and between the Registrant and the Regents of University of California, dated October 23, 2013, as amended May 14, 2014 (incorporated by reference to Exhibit 10.5 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.5	Lease Agreement, by and between the Registrant and ARE-MA Region No. 38, LLC, dated December 11, 2011, as amended by First Amendment to Lease, by and between ARE-MA Region No. 38, LLC, dated October 26, 2012, and Second Amendment to Lease, by and between ARE-MA Region No. 38, LLC, dated May 9, 2013 (incorporated by reference to Exhibit 10.6 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.6	Offer letter by and between the Registrant and Jeffrey M. Jonas, dated July 18, 2013 (incorporated by reference to Exhibit 10.7 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.7	Offer letter by and between the Registrant and Albert J. Robichaud, dated September 25, 2011 (incorporated by reference to Exhibit 10.8 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.8	Offer letter by and between the Registrant and Stephen J. Kanes, dated May 21, 2013 (incorporated by reference to Exhibit 10.9 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)

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Exhibit No.	Description
10.9	Offer letter by and between the Registrant and Kimi Iguchi, dated February 7, 2013 (incorporated by reference to Exhibit 10.10 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.10	Non-Solicitation, Confidentiality and Assignment Agreement by and between the Registrant and Jeffrey M. Jonas, dated August 19, 2013 (incorporated by reference to Exhibit 10.11 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.11	Non-Solicitation, Confidentiality and Assignment Agreement by and between the Registrant and Albert J. Robichaud, dated November 7, 2011 (incorporated by reference to Exhibit 10.12 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.12	Non-Solicitation, Confidentiality and Assignment Agreement by and between the Registrant and Stephen J. Kaness, dated July 17, 2013 (incorporated by reference to Exhibit 10.13 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.13	Non-Solicitation, Confidentiality and Assignment Agreement by and between the Registrant and Kimi Iguchi, dated March 8, 2013 (incorporated by reference to Exhibit 10.14 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.14#	Senior Executive Cash Incentive Bonus Plan (incorporated by reference to Exhibit 10.15 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.15	Form of Indemnification Agreement to be entered into between the Registrant and its directors (incorporated by reference to Exhibit 10.16 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.16	Form of Indemnification Agreement to be entered into between the Registrant and its officers (incorporated by reference to Exhibit 10.17 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.17	Supply Agreement by and between the Registrant and CyDex Pharmaceuticals, Inc., dated December 13, 2012, as amended August 21, 2013 and April 30, 2014 (incorporated by reference to Exhibit 10.18 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.18#	2014 Employee Stock Purchase Plan (incorporated by reference to Exhibit 10.19 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.19	Offer Letter by and between the Registrant and Thomas D. Anderson, dated April 15, 2014 (incorporated by reference to Exhibit 10.20 of the Registrant's Registration Statement on Form S-1 (File No. 333-196849) filed on July 8, 2014)
10.20	Severance and Change In Control Agreement between the Registrant and Jeffrey M. Jonas, dated September 25, 2014 (incorporated by reference to Exhibit 10.20 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)
10.21	Severance and Change In Control Agreement between the Registrant and Kimi Iguchi, dated September 30, 2014 (incorporated by reference to Exhibit 10.21 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)
10.22	Severance and Change In Control Agreement between the Registrant and Stephen J. Kaness, dated September 30, 2014 (incorporated by reference to Exhibit 10.22 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)

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Exhibit No.	Description
10.23	Severance and Change In Control Agreement between the Registrant and Albert J. Robichaud, dated September 25, 2014 (incorporated by reference to Exhibit 10.23 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)
10.24	Severance and Change In Control Agreement between the Registrant and Thomas D. Anderson, dated September 26, 2014 (incorporated by reference to Exhibit 10.24 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)
21.1	Subsidiaries of the Registrant (incorporated by reference to Exhibit 21.1 of the Registrant's Annual Report on Form 10-K (File No. 001-36544) filed on March 6, 2015)
23.1	Consent of PricewaterhouseCoopers LLP, Independent Registered Public Accounting Firm
23.2*	Consent of Goodwin Procter LLP (included in Exhibit 5.1)
24.1	Power of Attorney (included on signature page)
101.INS	XBRL Instance Document
101.SCH	XBRL Taxonomy Extension Schema Document
101.CAL	XBRL Taxonomy Extension Calculation Document
101.DEF	XBRL Taxonomy Extension Definition Linkbase Document
101.LAB	XBRL Taxonomy Extension Labels Linkbase Document
101.PRE	XBRL Taxonomy Extension Presentation Link Document

* To be filed by amendment.

Application has been made to the Securities and Exchange Commission for confidential treatment of certain provisions. Omitted material for which confidential treatment has been requested has been filed separately with the Securities and Exchange Commission.

Represents management compensation plan.