Vanda Pharmaceuticals Inc. Form S-1/A January 09, 2007

## As filed with the Securities and Exchange Commission on January 9, 2007 Registration No. 333-139485

# SECURITIES AND EXCHANGE COMMISSION Washington, D.C. 20549

#### AMENDMENT NO. 1 TO FORM S-1 REGISTRATION STATEMENT UNDER THE SECURITIES ACT OF 1933

#### Vanda Pharmaceuticals Inc.

(Exact Name of Registrant as Specified in its Charter)

Delaware283403-0491827(State or Other Jurisdiction of<br/>Incorporation or Organization)(Primary Standard Industrial<br/>Classification Code Number)(I.R.S. Employer<br/>Identification Number)

9605 Medical Center Drive Suite 300 Rockville, Maryland 20850 (240) 599-4500

(Address, including zip code, and telephone number, including area code, of registrant s principal executive offices)

Mihael H. Polymeropoulos, M.D. Chief Executive Officer 9605 Medical Center Drive Suite 300 Rockville, Maryland 20850 (240) 599-4500

(Name, address, including zip code, and telephone number, including area code, of agent for service)

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**Approximate date of proposed sale to the public:** From time to time or at one time after this registration statement becomes effective in light of market conditions and other factors.

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, as amended, check the following box. o

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. o \_\_\_\_\_\_\_

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. o \_\_\_\_\_\_

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. o \_\_\_\_\_\_

#### **CALCULATION OF REGISTRATION FEE**

		Proposed Maximum	Proposed Maximum	Amount of
Title of Each Class of	Amount to be	Offering Price	Aggregate Offering	Registration
Securities to be Registered	Registered(1)	Per Share(2)	Price(2)	Fee(2)(3)
Common stock, \$0.001 par				
value	4,025,000	\$24.78	\$99,739,500	\$193.80

- (1) Includes 525,000 shares of common stock that may be purchased by the underwriters to cover over-allotments, if any.
- (2) Estimated solely for the purpose of calculating the registration fee pursuant to Rule 457(c) promulgated under the Securities Act of 1933, as amended, by taking the average of the high and low sales price of the common stock on The Nasdaq Global Market on January 3, 2007.
- (3) A fee of \$10,478.33 was paid at the time of the initial filing of this Registration Statement.

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 this Registration Statement shall become effective on such date as the Commission, acting pursuant to said Section 8(a), may determine.

The information in this 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 prospectus is not an offer to sell these securities, and we are not soliciting an offer to buy these securities in any state where the offer or sale is not permitted.

#### Subject to completion, dated January 9, 2007

us

3,500,000 shares

Common stock

We are offering 3,500,000 shares of our common stock.

Our common stock is quoted on The Nasdaq Global Market under the symbol VNDA. The last reported sale price for our common stock on January 3, 2007 was \$24.83 per share.

	Per share	Total
Public offering price	\$	\$
Underwriting discounts and commissions	\$	\$
Proceeds to us, before expenses	\$	\$

We have granted the underwriters an option for a period of 30 days to purchase up to 525,000 additional shares of common stock to cover over-allotments, if any.

Investing in our common stock involves a high degree of risk. See Risk factors beginning on page 8.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or passed on the adequacy or accuracy of this prospectus. Any representation to the contrary is a criminal offense.

JPMorgan Morgan Stanley

**Banc of America Securities LLC** 

Natexis Bleichroeder Inc.

, 2007

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You should rely only on the information contained in this prospectus. We have not authorized anyone to provide you with information different from that contained in this prospectus. We are offering to sell, and are seeking offers to buy, shares of common stock only in jurisdictions where offers and sales are permitted. The information contained in this prospectus is accurate only as of the date of this prospectus, regardless of the time of delivery of this prospectus or of any sale of the common stock.

For investors outside the United States: Neither we nor any of the underwriters have 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 in the United States. You are required to inform yourselves about and to observe any restrictions relating to this offering and the distribution of this prospectus.

Vanda is a trademark of Vanda Pharmaceuticals Inc. This prospectus may also include other registered and unregistered trademarks of Vanda Pharmaceuticals Inc. and other persons.

Unless the context otherwise requires, we use the terms Vanda, the Company, we, us and our in this prospectus refer to Vanda Pharmaceuticals Inc.

#### **Prospectus summary**

This summary highlights the most important features of this offering and the information contained elsewhere in this prospectus. This summary is not complete and does not contain all of the information that you should consider before investing in our common stock. You should also read the entire prospectus carefully, especially the risks of investing in our common stock discussed under Risk factors and our consolidated financial statements and related notes included in this prospectus.

#### Vanda Pharmaceuticals Inc.

We are a biopharmaceutical company focused on the development and commercialization of our portfolio of clinical-stage product candidates for central nervous system disorders. We believe that each of our product candidates will address a large market with significant unmet medical needs by offering advantages over currently available therapies. Our product portfolio includes:

iloperidone, a compound for the treatment of schizophrenia and bipolar disorder, which has demonstrated positive top-line results from a recently completed Phase III trial in schizophrenia. We expect to file a New Drug Application (NDA) for iloperidone in schizophrenia with the United States Food and Drug Administration (FDA) by the end of 2007

VEC-162, a compound for the treatment of sleep and mood disorders, which has demonstrated positive top-line results from a recently completed Phase III trial in transient insomnia. We expect to initiate at least one additional Phase III trial in chronic sleep disorders in the second half of 2007

VSF-173, a compound for the treatment of excessive sleepiness, for which we expect to begin a Phase II trial in mid-2007

We hold exclusive, worldwide rights to these compounds and plan to develop a focused U.S. sales force for the commercialization of iloperidone and VSF-173. We plan to seek partners for commercialization of these compounds outside of the United States. Given the large size of the prescribing physician base for sleep and mood disorders, we plan to partner with a global pharmaceutical company for the development and commercialization of VEC-162 worldwide, although we have not yet identified such a partner.

Our founder and Chief Executive Officer, Mihael H. Polymeropoulos, M.D., started our operations early in 2003 after establishing and leading the Pharmacogenetics Department at Novartis AG. In acquiring and developing our compounds we have relied upon our deep expertise in the scientific disciplines of pharmacogenetics and pharmacogenomics. These scientific disciplines examine both genetic variations among people that influence response to a particular drug, and the multiple pathways through which drugs affect people. We believe that the combination of our expertise in these disciplines and our drug development expertise may provide us with preferential access to compounds discovered by other pharmaceutical companies, and will allow us to identify new uses for these compounds. These capabilities should also enable us to shorten the time it takes to commercialize a drug when compared to traditional approaches.

*Iloperidone for Schizophrenia and Bipolar Disorder*. We are developing iloperidone for the treatment of schizophrenia and bipolar disorder. Today, schizophrenia patients are treated primarily with drugs known as atypical antipsychotics. These drugs have been called atypical because they are regarded as being safer and more effective than drugs known as typical antipsychotics, which have been prescribed since the 1950s. Atypical antipsychotics achieved worldwide sales in excess of \$12 billion in 2005. However, despite their commercial success, atypical antipsychotics

offer only modest and unpredictable efficacy and induce serious side

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effects, resulting in poor patient compliance. Consequently, there remains a high degree of dissatisfaction with atypical antipsychotics among patients and physicians. A recent study conducted by the National Institute of Mental Health and published in *The New England Journal of Medicine* found that 74% of patients taking antipsychotics discontinued treatment within 18 months. Given the safety and efficacy shortcomings of current drugs, we believe that iloperidone may be an attractive alternative therapy.

Iloperidone may offer several advantages over existing therapies. In multiple Phase III trials of more than 2,000 patients, iloperidone showed a reduced risk of the side effects most associated with atypical antipsychotics, including weight gain, diabetes induction, involuntary body movements, elevated levels of the hormone prolactin, and sleepiness. The application of our pharmacogenetics and pharmacogenomics expertise may provide additional differentiation for iloperidone by identifying genetic markers of iloperidone is efficacy and safety. Our market research indicates that physicians treating schizophrenia patients would welcome a test that leads to improved patient outcomes by using genetic information to customize drug therapy. We also plan to distinguish iloperidone through the development of an extended-release injectable formulation of the compound that is administered only once every four weeks. We believe this formulation will help address the patient compliance and discontinuation problems commonly associated with atypical antipsychotics. Our extended-release injectable formulation has successfully completed a Phase I/IIa trial. We believe we will need to complete one Phase III trial with this formulation to be able to file for FDA approval.

In December 2006 we announced positive top-line results from our Phase III trial of iloperidone in schizophrenia. This Phase III trial was a randomized, double-blind, placebo-controlled, multi-center, four-week inpatient study that enrolled 604 patients, and examined the effects of a 12-mg oral formulation of iloperidone dosed twice-daily (or 24 mg each day). The primary endpoint of the trial was efficacy versus placebo on the Positive and Negative Symptoms Scale (PANSS), for which iloperidone demonstrated statistically significant improvement. Iloperidone also demonstrated statistically significant improvement versus placebo in several other measures of efficacy. The drug also appeared to be safe and well-tolerated in the trial. Based on discussions with the FDA, we believe that our data and documentation on oral iloperidone will be sufficient to support the filing of an NDA with the FDA by the end of 2007. We expect to meet with the FDA in the first quarter of 2007 regarding this filing.

The trial results also validated the pharmacogenetics work undertaken by the Company. Patients in the trial with a common genetic mutation, estimated to occur in approximately 70% of the population, experienced significantly better treatment results with iloperidone than the general treatment population. We also demonstrated in the trial that patients with an uncommon genetic attribute may experience longer QTc intervals (a measurement of specific electrical activity in the heart as captured on an electrocardiogram, corrected for heart rate) while taking iloperidone. The Company has developed a single blood test with these markers and may seek to commercialize this test alongside iloperidone.

In addition to schizophrenia, we believe iloperidone may be effective in treating bipolar disorder. All of the approved atypical antipsychotics have received approval for bipolar disorder subsequent to commercialization for the treatment of schizophrenia. Iloperidone is ready for an initial Phase III trial in bipolar disorder.

We expect to build our own sales force to market iloperidone directly to psychiatrists and other target physicians in the U.S. This medical community is relatively small and we believe that we can cost-effectively develop such a sales force. Outside of the U.S., we expect to find commercial partners for iloperidone.

VEC-162 for Sleep and Mood Disorders. We are developing VEC-162 for the treatment of sleep and mood disorders. The markets for both sleep disorder drugs and for mood disorder drugs are large and growing. Insomnia drugs enjoyed worldwide sales of approximately \$4.5 billion in 2005, even though industry sources indicate that the majority of people suffering from insomnia do not receive any treatment at all for their condition. In addition, antidepressant drugs achieved worldwide sales in excess of \$19 billion in 2005.

We believe VEC-162 may offer several benefits when compared to currently approved insomnia therapies. Unlike many approved therapies, VEC-162 works by directly targeting the melatonin receptors in the brain which govern the body s natural sleep/wake cycle. Because it appears to modulate the sleep/wake cycle, we believe that VEC-162 may be the first drug to address the underlying cause of sleeplessness in circadian rhythm sleep disorders, which, according to research conducted by LEK Consulting, LLC, a leading consulting firm, represent a significant portion of the insomnia market. Circadian rhythm sleep disorders are those, such as jet lag, where the circadian rhythm, or the rhythmic output of the human biological clock governed by melatonin and other hormones, is out of alignment with a person s daily activities or lifestyle. VEC-162 also appears to be safe, with no significant side effects or effects on next-day performance. As demonstrated in our recently completed Phase III trial, VEC-162 provides a benefit in both sleep onset, or time to fall asleep, and sleep maintenance, or ability to stay asleep. Based on these trial results, we believe that VEC-162 will compare favorably to efficacy achieved by currently approved insomnia drugs, not only for circadian rhythm sleep disorders but also for other types of insomnia. We also believe that VEC-162 is unlikely to be classified as a Schedule IV controlled substance by the United States Drug Enforcement Agency (DEA) because a recently approved compound with a similar mechanism of action has been shown not to have potential for abuse.

In November 2006 we announced positive top-line results from our Phase III clinical trial evaluating VEC-162 in transient insomnia. VEC-162 demonstrated statistically significant improvements at all three tested doses compared to placebo in the primary endpoint of the trial, latency to persistent sleep, a measure of sleep onset. VEC-162 also produced statistically significant improvements relative to placebo in latency to non-awake, another measure of sleep onset, wake after sleep onset, a measure of sleep maintenance, and total sleep time. VEC-162 was also demonstrated to be safe and well-tolerated. We believe that we will need to conduct additional Phase III trials in chronic sleep disorders to receive FDA approval of VEC-162 for the treatment of insomnia.

In addition to insomnia, we believe that VEC-162 may be effective in treating depression. VEC-162 has properties similar to Novartis agomelatine, an older compound with a similar mechanism of action, which in a Phase III trial demonstrated more rapid efficacy and reduced side effects when compared to a market-leading antidepressant. VEC-162 is ready for Phase II trials in depression, having demonstrated an antidepressant effect in animal models and having completed several Phase I trials.

VSF-173 for Excessive Sleepiness. VSF-173 is an oral compound that has demonstrated effects on animal sleep/wake patterns and gene expression suggestive of a stimulant effect. As a result of these observations and safety data from previous human trials, we are planning to initiate a Phase II trial of VSF-173 in excessive sleepiness in mid-2007. Excessive sleepiness is a rapidly growing market which generated worldwide sales of approximately \$500 million in 2005 and is currently treated primarily by stimulants.

#### **Strategy**

Our goal is to create a leading biopharmaceutical company focused on developing and commercializing products that address critical unmet medical needs through the application of our drug development and pharmacogenemics and pharmacogenetics expertise. The key elements of our strategy to accomplish this goal are to:

pursue the clinical development and regulatory approval of our current product candidates

enter into partnerships to extend our commercial reach

develop a focused commercialization capability in the United States

apply our pharmacogenomics and pharmacogenetics expertise to differentiate our product candidates from other available therapies

expand our product portfolio through the identification and acquisition of additional compounds

#### **Recent developments**

We intend to engage an investment bank to provide financial and strategic advisory services to the Company, which may lead to one or more possible transactions, including the acquisition, sale or licensing by the Company of businesses or product candidates, the sale or licensing to a third party of one or more of our own product candidates, or the acquisition of the Company. We cannot assure you that we will complete any acquisitions, sales or licenses, or that, if completed, any acquisition, sale or license will be successful or on attractive terms.

#### Risks associated with our business

Our business is subject to numerous risks, as more fully described in the section entitled Risk factors. We may be unable, for many reasons, including those that are beyond our control, to implement our current business strategy. Those reasons could include delays in obtaining, or a failure to obtain, regulatory approval for our product candidates, a failure to maintain and to protect our intellectual property, our failure to meet certain development and commercialization milestones in our sublicense agreement with Novartis AG, which could cause our rights to iloperidone to be terminated, the exercise by Bristol-Myers Squibb Company of its option to reacquire our rights to VEC-162 at the end of our Phase III program (if we have not entered into a commercialization agreement with a third party covering significant markets by that time) and the exercise by Novartis of its option to reacquire rights to VSF-173 at the end of our Phase II trials or at the end of our Phase III trials. We have a limited operating history and have incurred net losses from our inception. We expect to continue to generate operating losses for the next several years. We will need to obtain additional capital to fund our continuing research and development activities. All of our product candidates are in development and none have been approved by the FDA for commercial sale. Even if we succeed in developing and commercializing one or more of our product candidates, we may never generate sufficient revenue to achieve and then sustain profitability.

#### **Corporate information**

We were incorporated in Delaware in November 2002. Our principal executive offices are located at 9605 Medical Center Drive, Suite 300, Rockville, Maryland, 20850 and our telephone number is (240) 599-4500. Our website address is www.vandapharma.com. The information on, or that can be accessed through, our website is not part of this prospectus.

#### The offering

Common stock we are offering: 3,500,000 shares

Common stock to be outstanding after this offering: 25,628,534 shares

#### Use of proceeds

We expect to use the net proceeds of this offering for working capital and for other general corporate purposes, including the funding of our NDA filing for iloperidone and our clinical development efforts. See Use of Proceeds.

Nasdaq Global Market symbol: VNDA

The number of shares of common stock to be outstanding after the offering is based on 22,128,534 shares of common stock outstanding as of December 31, 2006. Except where we state otherwise, the number of shares of common stock to be outstanding after this offering does not take into account:

1,347,205 shares of common stock issuable upon the exercise of stock options outstanding as of December 31, 2006 under our Second Amended and Restated Management Equity Plan and agreements entered into under such plan, with a weighted-average exercise price of \$1.69 per share

359,527 shares of common stock issuable upon the exercise of stock options outstanding as of December 31, 2006 under our 2006 Equity Incentive Plan and agreements entered into pursuant to such plan, with a weighted-average exercise price of \$20.21 per share

an additional 1,140,470 shares reserved for issuance under the 2006 Equity Incentive Plan as of December 31, 2006 for future stock option grants and purchases (see note 4 of Notes to condensed consolidated financial statements )

Finally, except where we state otherwise, the information we present in this prospectus reflects no exercise of the underwriter s over-allotment option.

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#### Summary consolidated financial data

The following tables summarize our consolidated financial data. The summary consolidated financial data are derived from our audited financial statements for the period from March 13, 2003 (inception of our operations) through December 31, 2003, and for the years ended December 31, 2004 and 2005. Data are also included from our unaudited financial statements for the nine months ended September 30, 2005 and 2006. This data should be read together with our financial statements and related notes, Selected Consolidated Financial Data, and Management s Discussion and Analysis of Financial Condition and Results of Operations included elsewhere in this prospectus. The as-adjusted balance sheet data contained in the following tables reflects our unaudited consolidated balance sheet data at September 30, 2006, adjusted for the sale of shares of common stock in this offering at an assumed public offering price of \$24.83 (the last reported sale price of our common stock on January 3, 2007), after deducting the estimated underwriting discounts, commissions and offering expenses payable by us.

Period from

	March 13, 2003 (inception) to December 31, 2003		Year ended December 31, 2004 2005				Nine months ended September 30, 2005 2006		
Statements of operations data Revenue	\$	47,565	\$	33,980	\$		\$	\$	
Operating expenses: Research and development General and	2	2,010,532	,	7,442,983	1	16,890,615	11,641,565	44,130,78	38
administrative	-	1,052,659	2	2,119,394		7,396,038	5,587,147	9,170,43	39
Total operating expenses	3	3,063,191	Ç	9,562,377	2	24,286,653	17,228,712	53,301,22	27
Loss from operations Interest and other	(3	3,015,626)	(9	9,528,397)	(2	24,286,653)	(17,228,712)	) (53,301,22	27)
income, net		44,805		59,060		410,001	188,288	1,681,53	34
Net loss before tax provision Tax provision	(2	2,970,821)	(9	9,469,337) 4,949	(2	23,876,652) 7,649	(17,040,424)	) (51,619,69	93)

Net loss Beneficial conversion feature deemed dividend to preferred stockholders(1)	(	2,970,821)	(9	9,474,286)		23,884,301) 33,486,623)	·	17,040,424) 18,500,005)	(	51,619,693)
Net loss attributable to common stockholders	\$ (	2,970,821)	\$ (9	9,474,286)	\$ (	57,370,924)	\$ (	35,540,429)	\$ (	51,619,693)
Net loss per share applicable to common stockholders, basic and diluted Weighted-average number of shares used in computing net loss per share, basic and	\$	(983.72)	\$	(3,137.18)	\$	(3,374.33)	\$	(3,094.51)	\$	(3.72)
diluted		3,020		3,020		17,002		11,485		13,862,613

<sup>(1)</sup> In September and December of 2005, we completed the sale of an additional 27,235,783 shares of Series B Preferred Stock for net proceeds of approximately \$33.5 million. After evaluating the fair value of the common stock obtainable upon conversion by the stockholders, we determined that the issuance of the Series B Preferred Stock sold in 2005 resulted in a beneficial conversion feature which was fully accreted in 2005 and is recorded as a deemed dividend to preferred stockholders of approximately \$33.5 million and approximately \$18.5 million for the year ended December 31, 2005 and the nine months ended September 30, 2005, respectively.

As of September 30, 2006	Actual	As adjusted
Balance sheet data		
Cash and cash equivalents and restricted cash	\$ 32,330,209	\$ 112,770,909
Short-term investments	11,096,506	11,096,506
Working capital	34,735,547	115,176,247
Total assets	47,282,498	127,723,198
Total liabilities	10,330,866	10,330,866
Deficit accumulated during the development stage	(87,949,101)	(87,949,101)
Total stockholders equity	36,951,632	117,392,332

#### **Risk factors**

Investing in our common stock involves a high degree of risk. You should consider carefully the risks and uncertainties described below, together with all of the other information in this prospectus, including the consolidated financial statements and the related notes appearing at the end of this prospectus, before deciding to invest in shares of our common stock. If any of the following risks actually occurs, our business, financial condition, results of operations and future prospects would likely be materially and adversely affected. In that event, the market price of our common stock could decline and you could lose all or part of your investment.

#### Risks related to our business and industry

Our success is dependent on the success of our three product candidates in clinical development: iloperidone, VEC-162 and VSF-173. If any of these product candidates are determined to be unsafe or ineffective in humans, whether in clinical trials or commercially, our business will be materially harmed.

Despite the positive results of our recently completed Phase III trials, we are uncertain whether any of our current product candidates in clinical development will ultimately prove to be effective and safe in humans. Frequently, product candidates that have shown promising results in clinical trials have suffered significant setbacks in later clinical trials or even after they are approved for commercial sale. Future uses of any of our product candidates, whether in clinical trials or commercially, may reveal that the product candidate is ineffective, unacceptably toxic, has other undesirable side effects or is otherwise not fit for further use. If we are unable to discover and develop products that are safe and effective, our business will be materially harmed.

#### Any failure or delay in completing clinical trials for our product candidates could severely harm our business.

Pre-clinical studies and clinical trials required to demonstrate the safety and efficacy of our product candidates are time-consuming and expensive and together take several years to complete. The completion of clinical trials for our product candidates may be delayed by many factors, including:

our inability to manufacture or obtain from third parties materials sufficient for use in pre-clinical studies and clinical trials

delays in patient enrollment and variability in the number and types of patients available for clinical trials

difficulty in maintaining contact with patients after treatment, resulting in incomplete data

poor effectiveness of product candidates during clinical trials

unforeseen safety issues or side effects

governmental or regulatory delays and changes in regulatory requirements and guidelines

If we fail to complete successfully one or more clinical trials for any of our product candidates, we may not receive the regulatory approvals needed to market that product candidate. Therefore, any failure or delay in commencing or completing these clinical trials would harm our business materially.

#### We face heavy government regulation, and FDA regulatory approval of our products is uncertain.

The research, testing, manufacturing and marketing of drug products such as those that we are developing are subject to extensive regulation by federal, state and local government authorities, including the FDA. To obtain regulatory approval of a product, we must demonstrate to the satisfaction of the applicable regulatory agency that, among other things, the product is safe and effective for its intended use. In addition, we must show that the manufacturing facilities used to produce the products are in compliance with current Good Manufacturing Practices regulations, or cGMP.

The process of obtaining FDA and other required regulatory approvals and clearances will require us to expend substantial time and capital. Despite the time and expense expended, regulatory approval is never guaranteed. The number of pre-clinical and clinical tests that will be required for FDA approval varies depending on the drug candidate, the disease or condition that the drug candidate is in development for, and the regulations applicable to that particular drug candidate. The FDA can delay, limit or deny approval of a drug candidate for many reasons, including that:

a drug candidate may not be safe or effective they may interpret data from pre-clinical and clinical testing in different ways than we do they may not approve our manufacturing process they may change their approval policies or adopt new regulations

For example, if certain of our methods for analyzing our trial data are not approved by the FDA, we may fail to obtain regulatory approval for our product candidates.

Moreover, if and when our products do obtain such approval or clearances, the marketing, distribution and manufacture of such products would remain subject to extensive ongoing regulatory requirements. Failure to comply with applicable regulatory requirements could result in:

warning letters
fines
civil penalties
injunctions
recall or seizure of products
total or partial suspension of production
refusal of the government to grant approvals
withdrawal of approvals
criminal prosecution

Any delay or failure by us to obtain regulatory approvals for our product candidates could diminish competitive advantages that we may attain and would adversely affect the marketing of our products. We have not received regulatory approval to market any of our product candidates in any jurisdiction.

Even if we do receive regulatory approval for our drug candidates, the FDA may impose limitations on the indicated uses for which our products may be marketed, subsequently withdraw approval or take other actions against us or our products that are adverse to our business. The FDA generally approves products for particular indications. An approval for a more limited indication reduces the size of the potential market for the product. Product approvals, once granted, may be withdrawn if problems occur after initial marketing.

We also are subject to numerous federal, state and local laws, regulations and recommendations relating to safe working conditions, laboratory and manufacturing practices, the environment and the use and disposal of hazardous substances used in connection with our discovery, research and development work. In addition, we cannot predict the extent of government regulations or the impact of new governmental regulations that might significantly harm the discovery, development, production and marketing of our products. We may be required to incur significant costs to comply with current or future laws or regulations, and we may be adversely affected by the cost of such compliance.

## We intend to seek regulatory approvals for our products in foreign jurisdictions, but we may not obtain any such approvals.

We intend to market our products outside the United States, either alone or with a commercial partner. In order to market our products in foreign jurisdictions, we may be required to obtain separate regulatory approvals and comply with numerous and varying regulatory requirements. The approval procedure varies among countries and jurisdictions and can involve additional testing, and the time required to obtain approval may differ from that required to obtain FDA approval. We have no experience with obtaining any such foreign approvals. Additionally, the foreign regulatory approval process may include all of the risks associated with obtaining FDA approval. For all of these reasons, we may not obtain foreign regulatory approvals on a timely basis, if at all. Approval by the FDA does not ensure approval by regulatory authorities in other countries or jurisdictions, and approval by one foreign regulatory authority does not ensure approval by regulatory authorities in other foreign countries or jurisdictions or by the FDA. We may not be able to file for regulatory approvals and may not receive necessary approvals to commercialize our products in any market. The failure to obtain these approvals could harm our business materially.

## Our product candidates may cause undesirable side effects or have other properties that could delay or prevent their regulatory approval or limit their marketability.

Undesirable side effects caused by our product candidates could interrupt, delay or halt clinical trials and could result in the denial of regulatory approval by the FDA or other regulatory authorities for any or all targeted indications, and in turn prevent us from commercializing our product candidates and generating revenues from their sale. For example, like many other drugs in its class, iloperidone is associated with a prolongation of the heart s QTc interval, which is a measurement of specific electrical activity in the heart as captured on an electrocardiogram, corrected for heart rate. A QTc interval that is significantly prolonged may result in an abnormal heart rhythm with adverse consequences including fainting, dizziness, loss of consciousness and death. No patient in the controlled portion of any of iloperidone s clinical trials was observed to have an interval that exceeded a 500-millisecond threshold of particular concern to the FDA. Two patients experienced a prolongation of 500 milliseconds or more during the open-label extension of one trial. We will continue to assess the side effect profile of iloperidone and our other product candidates in our ongoing clinical development program.

In addition, if any of our product candidates receive marketing approval and we or others later identify undesirable side effects caused by the product, we could face one or more of the following:

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

regulatory authorities may withdraw their approval of the product

we may be required to change the way the product is administered, conduct additional clinical trials or change the labeling of the product

our reputation may suffer

Any of these events could prevent us from achieving or maintaining market acceptance of the affected product or could substantially increase the costs and expenses of commercializing the product candidate, which in turn could delay or prevent us from generating significant revenues from its sale.

#### Our product candidates may never achieve market acceptance even if we obtain regulatory approvals.

Even if we receive regulatory approvals for the sale of our product candidates, the commercial success of these products will depend, among other things, on their acceptance by physicians, patients, third-party payors and other members of the medical community as a therapeutic and cost-effective alternative to competing products and treatments. The degree of market acceptance of any of our product candidates will depend on a number of factors, including the demonstration of its safety and efficacy, its cost-effectiveness, its potential advantages over other therapies, the reimbursement policies of government and third-party payors with respect to the product candidate, and the effectiveness of our marketing and distribution capabilities. If our product candidates fail to gain market acceptance, we may be unable to earn sufficient revenue to continue our business. If our product candidates do not become widely accepted by physicians, patients, third-party payors and other members of the medical community, it is unlikely that we will ever become profitable.

If we fail to obtain the capital necessary to fund our research and development activities, we may be unable to continue operations or we may be forced to share our rights to commercialize our product candidates with third parties on terms that may not be attractive to us.

Based on our current operating plans, and assuming the sale of 3,500,000 shares of our common stock in this offering at \$24.83 per share (the last reported sale price of our common stock on The Nasdaq Global Market on January 3, 2007), we believe that the proceeds from this offering, together with our existing cash, restricted cash, cash equivalents and short-term investments, will be sufficient to meet our anticipated operating needs through early 2008, and after that time we will require additional capital. In budgeting for our activities following this offering, we have relied on a number of assumptions, including assumptions that we will file an NDA for iloperidone in schizophrenia with the FDA by the end of 2007, that we will continue to expend funds in preparation of commercial launch of iloperidone, that we will expend funds on the extended-release injectable formulation of iloperidone, that we will initiate at least one additional VEC-162 Phase III trial for chronic sleep disorders in the second half of 2007 and that this trial will be conducted in accordance with our expectations, that we will initiate our VSF-173 Phase II trial for excessive sleepiness in mid-2007 and that this trial will be conducted in accordance with our expectations, that we will not engage in further in-licensing activities, that we will not receive any proceeds from potential partnerships, that we will not expend funds on the bipolar indication for iloperidone or on a Phase II trial of VEC-162 for depression, that we will continue to evaluate pre-clinical compounds for potential development, that we will be able to continue the manufacturing of our product candidates at commercially reasonable prices, that we will be able to retain our key personnel, and that we will not incur any significant contingent liabilities. We may need to raise additional funds more quickly if one or more of our assumptions proves to be incorrect or if we choose to expand our product development efforts more rapidly than presently anticipated or seek to acquire additional

product candidates, and we may also decide to raise additional funds even before they are needed if the conditions for raising capital are favorable.

We may seek to sell additional equity or debt securities or obtain a bank credit facility. The sale of additional equity or debt securities, if convertible, could result in dilution to our stockholders. The incurrence of indebtedness would result in increased fixed obligations and could also result in covenants that would restrict our operations.

We cannot assure you that additional funds will be available when we need them on terms that are acceptable to us, or at all. The unavailability of financing may require us to delay, scale back or eliminate expenditures for our research, development and marketing activities necessary to commercialize our potential biopharmaceutical products. If we are unable to secure sufficient capital to fund our research and development activities, we may not be able to continue operations or we may have to enter into collaboration agreements that could require us to share commercial rights to our products to a greater extent or at earlier stages in the drug development process than we currently intend. Collaborations that are consummated by us prior to proof-of-efficacy and safety of a product candidate could impair our ability to realize value from that product candidate.

## We have incurred operating losses in each year since our inception and expect to continue to incur substantial and increasing losses for the foreseeable future.

We have a limited operating history. We have not generated any revenue from product sales to date and we cannot estimate with precision the extent of our future losses. We do not currently have any products that have been approved for commercial sale and we may never generate revenue from selling products or achieve profitability. We expect to continue to incur substantial and increasing losses for the foreseeable future, particularly as we increase our research, clinical development and administrative activities. As a result, we are uncertain when or if we will achieve profitability and, if so, whether we will be able to sustain it. We have been engaged in identifying and developing compounds and product candidates since March 2003. As of September 30, 2006, we have accumulated net losses of approximately \$87.9 million. Our ability to achieve revenue and profitability is dependent on our ability to complete the development of our product candidates, obtain necessary regulatory approvals, and have our products manufactured and marketed. We cannot assure you that we will be profitable even if we successfully commercialize our products. Failure to become and remain profitable may adversely affect the market price of our common stock and our ability to raise capital and continue operations.

# If our contract research organizations do not successfully carry out their duties or if we lose our relationships with contract research organizations, our drug development efforts could be delayed.

We are dependent on contract research organizations, third-party vendors and investigators for pre-clinical testing and clinical trials related to our drug discovery and development efforts and we will likely continue to depend on them to assist in our future discovery and development efforts. These parties are not our employees and we cannot control the amount or timing of resources that they devote to our programs. If they fail to devote sufficient time and resources to our drug development programs or if their performance is substandard, it will delay the development and commercialization of our product candidates. The parties with which we contract for execution of our clinical trials play a significant role in the conduct of the trials and the subsequent collection and analysis of data. Their failure to meet their obligations could adversely affect clinical development of our product candidates. Moreover, these parties may

also have relationships with other commercial entities, some of which may compete with us. If they assist our competitors, it could harm our competitive position.

If we lose our relationship with any one or more of these parties, we could experience a significant delay in both identifying another comparable provider and then contracting for its services. We may be unable to retain an alternative provider on reasonable terms, if at all. Even if we locate an alternative provider, it is likely that this provider may need additional time to respond to our needs and may not provide the same type or level of service as the original provider. In addition, any provider that we retain will be subject to current Good Laboratory Practices, or cGLP, and similar foreign standards and we do not have control over compliance with these regulations by these providers. Consequently, if these practices and standards are not adhered to by these providers, the development and commercialization of our product candidates could be delayed.

### If our CTD contractors do not successfully carry out their duties or if we lose our relations with our CTD contractors, our NDA for iloperidone could be delayed.

We are dependent on third-party vendors for the preparation of the Common Technical Dossier (CTD) related to the NDA we expect to file for iloperidone by the end of 2007. These parties are not our employees and we cannot control the amount or timing of resources that they devote to our program. If they fail to devote sufficient time and resources to our NDA preparation or if their performance is substandard, it will delay the approval of iloperidone.

If we lose our relationship with any one or more of these third parties, we could experience a significant delay in both identifying another comparable provider and then contracting for its services. We may be unable to retain an alternative provider on reasonable terms, if at all. Even if we locate an alternative provider, it is likely that this provider may need additional time to respond to our needs and may not provide the same type or level of service as the original provider. Consequently, the NDA and commercialization of iloperidone could be delayed.

### We rely on a limited number of manufacturers for our product candidates and our business will be seriously harmed if these manufacturers are not able to satisfy our demand and alternative sources are not available.

We do not have an in-house manufacturing capability and depend completely on a small number of third-party manufacturers and active pharmaceutical ingredient formulators for the manufacture of our product candidates. We do not have long-term agreements with any of these third parties, and if they are unable or unwilling to perform for any reason, we may not be able to locate alternative acceptable manufacturers or formulators or enter into favorable agreements with them. Any inability to acquire sufficient quantities of our product candidates in a timely manner from these third parties could delay clinical trials and prevent us from developing our product candidates in a cost-effective manner or on a timely basis. In addition, manufacturers of our product candidates are subject to cGMP and similar foreign standards and we do not have control over compliance with these regulations by our manufacturers. If one of our contract manufacturers fails to maintain compliance, the production of our product candidates could be interrupted, resulting in delays and additional costs. In addition, if the facilities of such manufacturers do not pass a pre-approval plant inspection, the FDA will not grant pre-market approval of our products.

Our manufacturing strategy presents the following additional risks:

the manufacturing process for VSF-173 has not been tested in quantities needed for continued clinical trials or commercial sales, and delays in scale-up to commercial quantities of VEC-162

and VSF-173 could delay clinical trials, regulatory submissions and commercialization of these product candidates

because most of our third-party manufacturers and formulators are located outside of the United States, there may be difficulties in importing our compounds or their components into the United States as a result of, among other things, FDA import inspections, incomplete or inaccurate import documentation or defective packaging

because of the complex nature of our compounds, our manufacturers may not be able to successfully manufacture our compounds in a cost-effective and/or timely manner

Materials necessary to manufacture our product candidates may not be available on commercially reasonable terms, or at all, which may delay the development, regulatory approval and commercialization of our product candidates.

We rely on our manufacturers to purchase from third-party suppliers the materials necessary to produce our product candidates for our clinical trials. Suppliers may not sell these materials to our manufacturers at the times we need them or on commercially reasonable terms. We do not have any control over the process or timing of the acquisition of these materials by our manufacturers. Moreover, we currently do not have any agreements for the commercial production of these materials. If our manufacturers are unable to obtain these materials for our clinical trials, product testing and potential regulatory approval of our product candidates could be delayed, significantly affecting our ability to develop our product candidates. If our manufacturers or we are unable to purchase these materials after regulatory approval has been obtained for our product candidates, the commercial launch of our product candidates would be delayed or there would be a shortage in supply, which would materially affect our ability to generate revenues from the sale of our product candidates.

We face substantial competition which may result in others developing or commercializing products before or more successfully than we do.

Our future success will depend on our ability to demonstrate and maintain a competitive advantage with respect to our product candidates and our ability to identify and develop additional product candidates through the application of our pharmacogenetics and pharmacogenomics expertise. Large, fully integrated pharmaceutical companies, either alone or together with collaborative partners, have substantially greater financial resources and have significantly greater experience than we do in:

developing products

undertaking pre-clinical testing and clinical trials

obtaining FDA and other regulatory approvals of products

manufacturing and marketing products

These companies may invest heavily and quickly to discover and develop novel products that could make our product candidates obsolete. Accordingly, our competitors may succeed in obtaining patent protection, receiving FDA approval or commercializing superior products or other competing products before we do.

We believe the primary competitors for each of our product candidates are as follows:

For iloperidone in the treatment of schizophrenia, the atypical antipsychotics Risperdal® (risperidone) by Johnson & Johnson (including the depot formulation Risperdal® Consta®), Zyprexa® (olanzapine) by Eli Lilly and Company,

 $Seroquel @ \ (quetiapine) \ by \ AstraZeneca \ PLC, \ Abilify @ \ (aripiprazole) \ by \ Bristol-Myers \ Squibb \ Company/Otsuka \ Pharmaceutical \ Co., \ Ltd.,$ 

Geodon® (ziprasidone) by Pfizer Inc., Invega® (paliperidone) by Johnson & Johnson, and generic clozapine, as well as the typical antipsychotics haloperidol, chlorpromazine, thioridazine, and sulpiride (all of which are generic). In addition to the approved products, compounds in Phase III trials (or for which an NDA has been recently filed) for the treatment of schizophrenia include bifeprunox (Wyeth/Solvay S.A./Lundbeck A/S), and asenapine (Organon International).

For VEC-162 in the treatment of insomnia, Rozerem<sup>tm</sup> (ramelteon) by Takeda Pharmaceuticals Company Limited, hypnotics such as Ambien<sup>®</sup> (zolpidem) by Sanofi-Aventis (including Ambien CR<sup>®</sup>), Lunesta<sup>®</sup> (eszopiclone) by Sepracor Inc. and Sonata<sup>®</sup> (zaleplon) by King Pharmaceuticals, Inc., generic compounds such as trazodone and doxepin, and over-the-counter remedies such as Benadryl<sup>®</sup> and Tylenol PM<sup>®</sup>. In addition to the approved products, compounds in Phase III trials for insomnia include indiplon (Neurocrine Biosciences, Inc.) gaboxadol (Merck & Co., Inc./Lundbeck A/S), and low-dose doxepin (Silenor<sup>tm</sup>, Somaxon Pharmaceuticals, Inc.).

For VEC-162 in the treatment of depression, antidepressants such as Paxil® (paroxetine) by GlaxoSmithKline (GSK), Zoloft® (sertraline) by Pfizer, Prozac® (fluoxetine) by Eli Lilly, and Lexapro (escitalopram) by Lundbeck A/S /Forest Pharmaceuticals Inc., Effexor® (venlafaxine) by Wyeth as well as other compounds such as Wellbutrin® (buproprion) by GSK and Cymbalta® (duloxetine) by Eli Lilly. In addition to the approved products, compounds in Phase III trials for depression include agomelatine (Novartis and Les Laboratoires Servier).

For VSF-173 in the treatment of excessive sleepiness, Provigil® (modafinil) and NuVigil® (armodafinil) by Cephalon Inc., and Xyrem® (sodium oxybate) by Jazz Pharmaceuticals, Inc.

#### We have no experience selling, marketing or distributing products and no internal capability to do so.

At present, we have limited marketing and no sales personnel. In order to commercialize any of our product candidates, we must either acquire or internally develop sales, marketing and distribution capabilities, or enter into collaborations with partners to perform these services for us. We may not be able to establish sales and distribution partnerships on acceptable terms or at all, and if we do enter into a distribution arrangement, our success will be dependent upon the performance of our partner. In the event that we attempt to acquire or develop our own in-house sales, marketing and distribution capabilities, factors that may inhibit our efforts to commercialize our products without partners or licensees include:

our inability to recruit and retain adequate numbers of effective sales and marketing personnel

the inability of sales personnel to obtain access to or persuade adequate numbers of physicians to prescribe our product

the lack of complementary products to be offered by our sales personnel, which may put us at a competitive disadvantage against companies with broader product lines

unforeseen costs associated with creating our own sales and marketing team or with entering into a partnering agreement with an independent sales and marketing organization

#### We will need to increase the size of our organization, and we may experience difficulties in managing our growth.

As of December 31, 2006, we had 44 full-time employees. We will need to continue to expand our managerial, operational, financial and other resources in order to manage and fund our operations, continue our development activities and commercialize our product candidates. Our

current personnel, systems and facilities are not adequate to support this future growth. To manage our growth, we must:

manage our clinical trials effectively

manage our internal development efforts effectively

improve our operational, financial, accounting and management controls, reporting systems and procedures

attract and retain sufficient numbers of talented employees

We may be unable to successfully implement these tasks on a larger scale and, accordingly, may not achieve our development and commercialization goals.

If we cannot identify, or enter into licensing arrangements for, new product candidates, our ability to develop a diverse product portfolio may be limited.

A component of our business strategy is acquiring rights to develop and commercialize compounds discovered or developed by other pharmaceutical and biotechnology companies for which we may find effective uses and markets by using our unique pharmacogenetics and pharmacogenomics expertise. Competition for the acquisition of these compounds is intense. If we are not able to identify opportunities to acquire rights to commercialize additional products, we may not be able to develop a diverse portfolio of products and our business may be harmed. Additionally, it may take substantial human and financial resources to secure commercial rights to promising product candidates. Moreover, if other firms develop pharmacogenetics and pharmacogenomics capabilities, we may face increased competition in identifying and acquiring additional product candidates.

If we lose key scientists or management personnel, or if we fail to recruit additional highly skilled personnel, it will impair our ability to identify, develop and commercialize product candidates.

We are highly dependent on principal members of our management team and scientific staff, including our Chief Executive Officer, Mihael H. Polymeropoulos, M.D. These executives each have significant pharmaceutical industry experience. The loss of any such executives, including Dr. Polymeropoulos, or any other principal member of our management team or scientific staff, would impair our ability to identify, develop and market new products.

Product liability lawsuits could divert our resources, result in substantial liabilities and reduce the commercial potential of our products.

The risk that we may be sued on product liability claims is inherent in the development of pharmaceutical products. For example, we face a risk of product liability exposure related to the testing of our product candidates in clinical trials and will face even greater risks upon any commercialization by us of our product candidates. We believe that we may be at a greater risk of product liability claims relative to other pharmaceutical companies because our compounds are intended to treat behavioral disorders, and it is possible that we may be held liable for the behavior and actions of patients who use our compounds. These lawsuits may divert our management from pursuing our business strategy and may be costly to defend. In addition, if we are held liable in any of these lawsuits, we may incur substantial liabilities and may be forced to limit or forego further commercialization of one or more of our products. Although we maintain general liability and product liability insurance, our aggregate coverage limit under this insurance is \$10,000,000, and while we believe this amount of insurance is sufficient to cover our product liability exposure, these limits may not be high enough to fully cover

potential liabilities. In addition, product liability insurance is becoming increasingly expensive, and we may not be able to obtain or maintain sufficient insurance coverage at an acceptable cost or otherwise to protect against potential product liability claims, which could prevent or inhibit the commercial production and sale of our products.

Legislative or regulatory reform of the healthcare system in the U.S. and foreign jurisdictions may affect our ability to sell our products profitably.

The continuing efforts of the U.S. and foreign governments, insurance companies, managed care organizations and other payors of health care services to contain or reduce health care costs may adversely affect our ability to set prices for our products which we believe are fair, and our ability to generate revenues and achieve and maintain profitability.

Specifically, in both the United States and some foreign jurisdictions there have been a number of legislative and regulatory proposals to change the healthcare system in ways that could affect our ability to sell our products profitably. In the United States, the Medicare Prescription Drug Improvement and Modernization Act of 2003 reforms the way Medicare will cover and reimburse for pharmaceutical products. This legislation could decrease the coverage and price that we may receive for our products. Other third-party payors are increasingly challenging the prices charged for medical products and services. It will be time-consuming and expensive for us to go through the process of seeking reimbursement from Medicare and private payors. Our products may not be considered cost effective, and coverage and reimbursement may not be available or sufficient to allow us to sell our products on a competitive and profitable basis. Further federal and state proposals and healthcare reforms are likely which could limit the prices that can be charged for the drugs we develop and may further limit our commercial opportunity. Our results of operations could be materially adversely affected by the Medicare prescription drug coverage legislation, by the possible effect of this legislation on amounts that private insurers will pay and by other healthcare reforms that may be enacted or adopted in the future.

In some foreign countries, including major markets in the European Union and Japan, the pricing of prescription pharmaceuticals is subject to governmental control. In these countries, pricing negotiations with governmental authorities can take six to twelve months or longer after the receipt of regulatory marketing approval for a product. To obtain reimbursement or pricing approval in some countries, we may be required to conduct a clinical trial that compares the cost-effectiveness of our product candidate to other available therapies. Our business could be materially harmed if reimbursement of our products is unavailable or limited in scope or amount or if pricing is set at unsatisfactory levels.

#### Our quarterly operating results may fluctuate significantly.

We expect our operating results to be subject to quarterly fluctuations. The revenues we generate, if any, and our operating results will be affected by numerous factors, including:

our addition or termination of development programs

variations in the level of expenses related to our existing three product candidates or future development programs

our execution of collaborative, licensing or other arrangements, and the timing of payments we may make or receive under these arrangements

any intellectual property infringement lawsuit in which we may become involved

regulatory developments affecting our product candidates or those of our competitors

If our quarterly operating results fall below the expectations of investors or securities analysts, the price of our common stock could decline substantially. Furthermore, any quarterly fluctuations in our operating results may, in turn, cause the price of our stock to fluctuate substantially. We believe that quarterly comparisons of our financial results are not necessarily meaningful and should not be relied upon as an indication of our future performance.

#### Risks related to intellectual property and other legal matters

Our rights to develop and commercialize our product candidates are subject in part to the terms and conditions of licenses or sublicenses granted to us by other pharmaceutical companies. With respect to VEC-162 and VSF-173, these terms and conditions include options in favor of these pharmaceutical companies to reacquire rights to commercialize and develop these product candidates in certain circumstances.

Iloperidone is based in part on patents and other intellectual property owned by Sanofi-Aventis and Novartis. Titan Pharmaceuticals, Inc. (Titan) holds an exclusive license from Sanofi-Aventis to the intellectual property owned by Sanofi-Aventis, and Titan has sublicensed its rights under such license on an exclusive basis to Novartis. We have acquired exclusive rights to this and other intellectual property through a further sublicense from Novartis. Our rights with respect to the intellectual property to develop and commercialize iloperidone may terminate, in whole or in part, if we fail to meet certain milestones contained in our sublicense agreement with Novartis relating to the time it takes for us to launch iloperidone commercially following regulatory approval, and the time it takes for us to receive regulatory approval following our submission of an NDA or equivalent foreign filing. We may also lose our rights to develop and commercialize iloperidone if we fail to pay royalties to Novartis, if we fail to comply with certain requirements in the sublicense regarding our financial condition, or if we fail to comply with certain restrictions regarding our other development activities. Finally, our rights to develop and commercialize iloperidone may be impaired if we do not cure breaches by Novartis and Titan of similar obligations contained in these sublicense and license agreements, although we are not aware of any such breach by Titan or Novartis. In the event of an early termination of our sublicense agreement, all rights licensed and developed by us under this agreement may be extinguished, which would have a material adverse effect on our business.

VEC-162 is based in part on patents that we have licensed on an exclusive basis and other intellectual property licensed from Bristol-Myers Squibb Company (BMS). Following the completion of the entire Phase III program for VEC-162, which may consist of several Phase III trials, and in the event that we have not entered into one or more development and commercialization agreements with one or more third parties covering certain significant markets, BMS has retained an option to reacquire the rights it has licensed to us to exclusively develop and commercialize VEC-162 on pre-determined financial terms, including the payment of royalties and milestone payments to us. BMS may terminate our license if we fail to meet certain milestones or if we otherwise breach our royalty or other obligations in the agreement. In the event that we terminate our license, or if BMS terminates our license due to our breach, all of our rights to VEC-162 (including any intellectual property we develop with respect to VEC-162) will revert back to BMS or otherwise be licensed back to BMS on an exclusive basis. Any termination or reversion of our rights to develop or commercialize VEC-162, including any reacquisition by BMS of our rights, may have a material adverse effect on our business.

VSF-173 is based in part on patents and other intellectual property that we have licensed on an exclusive basis from Novartis. Novartis has the option to reacquire rights to co-develop and exclusively commercialize VSF-173 following the completion of the Phase II trials, and an additional option to reacquire co-development rights and exclusive commercialization rights

following the completion of the Phase III clinical trials, subject in each case to Novartis payment of pre-determined royalties and other payments to us. In the event that Novartis chooses not to exercise either of these options and we decide to enter into a partnering arrangement to help us commercialize VSF-173, Novartis has a right of first refusal to negotiate such an agreement with us, as well as a right to submit a last matching counteroffer regarding such an agreement. In addition, our rights with respect to VSF-173 may terminate, in whole or in part, if we fail to meet certain development and commercialization milestones described in our license agreement relating to the time it takes us to complete our development work on VSF-173. These rights may also terminate in whole or in part if we fail to make royalty or milestone payments or if we do not comply with requirements in our license agreement regarding our financial condition. In the event of an early termination of our license agreement, all rights licensed and developed by us under this agreement may revert back to Novartis. Any termination or reversion of our rights to develop or commercialize VSF-173, including any reacquisition by Novartis of our rights, may have a material adverse effect on our business.

If our efforts to protect the proprietary nature of the intellectual property related to our products are not adequate, we may not be able to compete effectively in our markets.

In addition to the rights we have licensed from Novartis and BMS relating to our product candidates, we rely upon intellectual property we own relating to our products, including patents, patent applications and trade secrets. As of December 31, 2006, we owned 15 pending provisional patent applications in the United States and three pending Patent Cooperation Treaty applications, which permit the pursuit of patents outside of the United States, relating to our product candidates in clinical development. Our patent applications may be challenged or fail to result in issued patents and our existing or future patents may be too narrow to prevent third parties from developing or designing around these patents. In addition, we rely on trade secret protection and confidentiality agreements to protect certain proprietary know-how that is not patentable, for processes for which patents are difficult to enforce and for any other elements of our drug development processes that involve proprietary know-how, information and technology that is not covered by patent applications. While we require all of our employees, consultants, advisors and any third parties who have access to our proprietary know-how, information and technology to enter into confidentiality agreements, we cannot be certain that this know-how, information and technology will not be disclosed or that competitors will not otherwise gain access to our trade secrets or independently develop substantially equivalent information and techniques. Further, the laws of some foreign countries do not protect proprietary rights to the same extent as the laws of the United States. As a result, we may encounter significant problems in protecting and defending our intellectual property both in the United States and abroad. If we are unable to protect or defend the intellectual property related to our technologies, we will not be able to establish or maintain a competitive advantage in our market.

If we do not obtain protection under the Hatch-Waxman Act and similar foreign legislation to extend our patents and to obtain market exclusivity for our product candidates, our business will be materially harmed.

The United States Drug Price Competition and Patent Term Restoration Act of 1984, more commonly known as the Hatch-Waxman Act, provides for an extension of patent protection for drug compounds for a period of up to five years to compensate for time spent in development. Assuming we gain a five-year extension for each of our current product candidates in clinical development, and that we continue to have rights under our sublicense and license agreements with respect to these product candidates, we would have exclusive rights to

iloperidone s United States new chemical entity patent (the primary patent covering the compound as a new composition of matter) until 2016, to VEC-162 s United States new chemical entity patent until 2022 and to VSF-173 s United States new chemical entity patent until 2019. In Europe, similar legislative enactments allow patent protection in the European Union to be extended for up to five years through the grant of a Supplementary Protection Certificate. Assuming we gain such a five-year extension for each of our current product candidates in clinical development, and that we continue to have rights under our sublicense and license agreements with respect to these product candidates, we would have exclusive rights to iloperidone s European new chemical entity patents until 2015, to VEC-162 s European new chemical entity patents until 2022 and to VSF-173 s European new chemical entity patents until 2017. Additionally, a recent directive in the European Union provides that companies who receive regulatory approval for a new compound will have a 10-year period of market exclusivity for that compound (with the possibility of a further one-year extension) in most EU countries, beginning on the date of such European regulatory approval, regardless of when the European new chemical entity patent covering such compound expires. A generic version of the approved drug may not be marketed or sold during such market exclusivity period. This directive may be of particular importance with respect to iloperidone, since the European new chemical entity patent for iloperidone will likely expire prior to the end of this 10-year period of market exclusivity. However, there is no assurance that we will receive the extensions of our patents or other exclusive rights available under the Hatch-Waxman Act or similar foreign legislation. If we fail to receive such extensions and exclusive rights, our ability to prevent competitors from manufacturing, marketing and selling generic versions of our products will be materially harmed.

### Litigation or third-party claims of intellectual property infringement could require us to divert resources and may prevent or delay our drug discovery and development efforts.

Our commercial success depends in part on our not infringing the patents and proprietary rights of third parties. Third parties may assert that we are employing their proprietary technology without authorization. In addition, third parties may obtain patents in the future and claim that use of our technologies infringes upon these patents. Furthermore, parties making claims against us may obtain injunctive or other equitable relief, which could effectively block our ability to develop and commercialize one or more of our product candidates. Defense of these claims, regardless of their merit, would divert substantial financial and employee resources from our business. In the event of a successful claim of infringement against us, we may have to pay substantial damages, obtain one or more licenses from third parties or pay royalties. In addition, even in the absence of litigation, we may need to obtain additional licenses from third parties to advance our research or allow commercialization of our product candidates. We may fail to obtain any of these licenses at a reasonable cost or on reasonable terms, if at all. In that event, we would be unable to develop and commercialize further one or more of our product candidates.

In addition, in the future we could be required to initiate litigation to enforce our proprietary rights against infringement by third parties. Prosecution of these claims to enforce our rights against others could divert substantial financial and employee resources from our business. If we fail to enforce our proprietary rights against others, our business will be harmed.

If we use hazardous and biological materials in a manner that causes injury or violates applicable law, we may be liable for damages.

Our research and development activities involve the controlled use of potentially hazardous substances, including toxic chemical and biological materials. We could be held liable for any

contamination, injury or other damages resulting from these hazardous substances. In addition, our operations produce hazardous waste products. While third parties are responsible for disposal of our hazardous waste, we could be liable under environmental laws for any required cleanup of sites at which our waste is disposed. Federal, state, foreign and local laws and regulations govern the use, manufacture, storage, handling and disposal of these hazardous materials. If we fail to comply with these laws and regulations at any time, or if they change, we may be subject to criminal sanctions and substantial civil liabilities, which may adversely affect our business.

Even if we continue to comply with all applicable laws and regulations regarding hazardous materials, we cannot eliminate the risk of accidental contamination or discharge and our resultant liability for any injuries or other damages caused by these accidents. Although we maintain pollution liability insurance, our coverage limit under this insurance is \$2,000,000, and while we believe this amount and type of insurance is sufficient to cover risks typically associated with our handling of materials, the insurance may not cover all environmental liabilities, and these limits may not be high enough to cover potential liabilities for these damages fully. The amount of uninsured liabilities may exceed our financial resources and materially harm our business.

### Risks related to this offering

Our stock price has been volatile and may be volatile in the future, and purchasers of our common stock could incur substantial losses.

The stock market has from time to time experienced significant price and volume fluctuations, and the market prices of the securities of life sciences companies without product revenues, such as ours, have historically been highly volatile. Since our initial public offering on April 12, 2006 and through January 3, 2007, our stock price has traded from a low of \$7.21 to a high of \$28.67.

The following factors, in addition to the other risk factors described in this section, may also have a significant impact on the market price of our common stock:

publicity regarding actual or potential testing or trial results or the outcome of regulatory review relating to products under development by us or our competitors

regulatory developments in the United States and foreign countries

developments concerning any collaboration or other strategic transaction we may undertake

announcements of patent issuances or denials, technological innovations or new commercial products by us or our competitors

actual or anticipated variations in our quarterly operating results

changes in estimates of our financial results or recommendations by securities analysts

additions or departures of key personnel or members of our board of directors

economic and other external factors beyond our control

As a result of these factors, holders of our common stock might be unable to sell their shares at or above the price they paid for such shares.

You will incur immediate and substantial dilution in the as-adjusted net tangible book value of the stock you purchase.

We estimate that the public offering price of our stock will be \$24.83 per share (the last reported sale price of our common stock on January 3, 2007). This amount is substantially higher

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than the as-adjusted net tangible book value that our outstanding common stock will have immediately after this offering. Accordingly, if you purchase shares of our common stock at the assumed public offering price, you will incur immediate and substantial dilution of \$20.21 per share (based on the number of shares of our common stock outstanding as of September 30, 2006). You may incur further dilution to the extent that holders of outstanding options exercise those options.

Our management will have broad discretion over the use of the proceeds we receive in this offering and might not apply the proceeds in ways that increase the value of your investment.

Our management will have broad discretion to use the net proceeds from this offering, and you will be relying on the judgment of our management regarding the application of these proceeds. Management might not apply the net proceeds of this offering in ways that increase the value of your investment. We expect to use the net proceeds from this offering for further clinical development of our current product candidates, possible investments in, or acquisitions of, new product candidates, working capital and other general corporate purposes. We have not allocated these net proceeds for any specific purposes. Our management might not be able to yield any return on the investment and use of these net proceeds. You will not have the opportunity to influence our decisions on how to use the proceeds.

#### If there are substantial sales of our common stock, our stock price could decline.

If our existing stockholders sell a large number of our common stock or the public market perceives that existing stockholders might sell shares of common stock, the market price of our common stock could decline significantly. As of December 31, 2006, we had 22,128,534 shares of our common stock outstanding. Of these shares, 6,185,534 shares are securities issued pursuant to registered transactions under the Securities Act and are freely tradable, unless purchased by an affiliate as that term is used in Rule 144 under the Securities Act of 1933, as amended (in which case any resale by such an affiliate will be subject to the restrictions imposed by Rule 144). An additional 15,914,391 of these shares were, as of December 31, 2006, tradable under Rule 144 or the Securities Act without registration, subject in some cases to volume limitations and holding periods under Rule 144 (including restrictions imposed on our affiliates). We expect that our directors and officers, as well as certain venture capital funds affiliated with certain of our directors (which funds held an aggregate of approximately 5,543,183 shares of our common stock as of December 31, 2006), will sign lock-up agreements pertaining to this offering, the restrictions of which will expire 30 days after this offering becomes effective.

Holders of approximately 6,103,579 shares of our outstanding and unregistered common stock as of December 31, 2006 have rights with respect to the registration of the sale of their shares of common stock with the SEC. These rights have been waived with respect to this offering.

In addition to our outstanding common stock, as of December 31, 2006 there are 1,347,205 shares of common stock that we have registered and that we are obligated to issue upon the exercise of currently outstanding options granted under our Second Amended and Restated Management Equity Plan. Upon the exercise of these options in accordance with their respective terms, these shares may be resold freely, subject to restrictions imposed on our affiliates under Rule 144.

We have also registered 1,500,000 shares of common stock that are authorized for issuance under our 2006 Equity Incentive Plan. The shares authorized for issuance under our 2006 Equity Incentive Plan can be freely sold in the public market upon issuance, subject to the restrictions imposed on our affiliates under Rule 144. We have granted options to purchase 359,527 shares under our 2006 Equity Incentive Plan as of December 31, 2006, none of which are vested.

If securities or industry analysts do not publish research or reports or publish unfavorable research about our business, our stock price and trading volume could decline.

The trading market for our common stock will depend in part on the research and reports that securities or industry analysts publish about us or our business. If one or more of the analysts who covers the Company downgrades our stock, our stock price would likely decline. If one or more of these analysts ceases to cover us or fails to publish regular reports on us, interest in the purchase of our stock could decrease, which could cause our stock price or trading volume to decline.

Anti-takeover provisions in our charter and bylaws, and in Delaware law, could prevent or delay a change in control of our company.

We are a Delaware corporation and the anti-takeover provisions of Section 203 of the Delaware General Corporation Law may discourage, delay or prevent a change in control by prohibiting us from engaging in a business combination with an interested stockholder for a period of three years after the person becomes an interested stockholder, even if a change of control would be beneficial to our existing stockholders. In addition, our amended and restated certificate of incorporation and bylaws may discourage, delay or prevent a change in our management or control over us that stockholders may consider favorable. Our amended and restated certificate of incorporation and bylaws:

authorize the issuance of blank check preferred stock that could be issued by our board of directors to thwart a takeover attempt

do not provide for cumulative voting in the election of directors, which would allow holders of less than a majority of the stock to elect some directors

establish a classified board of directors, as a result of which the successors to the directors whose terms have expired will be elected to serve from the time of election and qualification until the third annual meeting following their election

require that directors only be removed from office for cause

provide that vacancies on the board of directors, including newly-created directorships, may be filled only by a majority vote of directors then in office

limit who may call special meetings of stockholders

prohibit stockholder action by written consent, requiring all actions to be taken at a meeting of the stockholders

establish advance notice requirements for nominating candidates for election to the board of directors or for proposing matters that can be acted upon by stockholders at stockholder meetings

For information regarding these and other provisions, please see Description of capital stock.

## Forward-looking statements

This prospectus includes forward-looking statements, as defined by federal securities laws, with respect to our financial condition, results of operations and business, and our expectations or beliefs concerning future events, including increases in operating margins. Words such as, but not limited to, believe, expect, anticipate, estimate, intend, plan, targets, likely, will, would, could, and similar expressions or phrases identify forward-lookin statements.

All forward-looking statements involve risks and uncertainties. The occurrence of the events described, and the achievement of the expected results, depend on many events, some or all of which are not predictable or within our control. Actual results may differ materially from expected results. Factors that may cause actual results to differ from expected results include, among others:

- a failure of our product candidates to be demonstrably safe and effective
- a failure to obtain regulatory approval for our products or to comply with ongoing regulatory requirements
- a lack of acceptance of our product candidates in the marketplace, or a failure to become or remain profitable
- our inability to obtain the capital necessary to fund our research and development activities
- our failure to identify or obtain rights to new product candidates
- a failure to develop or obtain sales, marketing and distribution resources and expertise or to otherwise manage our growth
- a loss of any of our key scientists or management personnel
- losses incurred from product liability claims made against us
- a loss of rights to develop and commercialize our products under our license and sublicense agreements

All future written and verbal forward-looking statements attributable to us or any person acting on our behalf are expressly qualified in their entirety by the cautionary statements contained or referred to in this section. We undertake no obligation, and specifically decline any obligation, to update or revise publicly any forward-looking statements, whether as a result of new information, future events or otherwise. In light of these risks, uncertainties and assumptions, the forward-looking events discussed in this prospectus might not occur.

See the section entitled Risk factors for a more complete discussion of these and other risks and uncertainties. The risk factors described in this prospectus are not necessarily all of the important factors that could cause actual results to differ materially from those expressed in any of our forward-looking statements. Other unknown or unpredictable factors also could affect our results. Consequently, there can be no assurance that the actual results or developments anticipated by us will be realized or, even if substantially realized, that they will have the expected consequences to, or effects on, us. Given these uncertainties, prospective investors are cautioned not to place undue reliance on such forward-looking statements.

# Use of proceeds

We estimate the net proceeds to us from the sale of the 3,500,000 shares of common stock in this offering to be approximately \$80.4 million after deducting the underwriting discounts and commissions and estimated offering expenses. If the underwriters overallotment option is exercised in full, we estimate the net proceeds will be approximately \$92.7 million.

We currently intend to use the net proceeds of this offering for the continued clinical trials of our product candidates, the pursuit of regulatory approval and the development of a commercialization strategy for iloperidone, other research and development activities, and for working capital purposes. More specifically, we currently intend to use the net proceeds of this offering as follows:

Approximately \$46.0 million to prepare an NDA for iloperidone in schizophrenia, which we currently anticipate will be submitted by the end of 2007, to continue to develop the oral and extended-release injectable formulation of iloperidone in schizophrenia, to begin to fund the cost of carcinogenicity studies of inactive metabolites that we expect will be required after iloperidone is approved by the FDA, and to initiate the commercialization of iloperidone in schizophrenia, the commercial launch of which we currently anticipate in early 2009

Approximately \$22.0 million to initiate an additional Phase III trial for VEC-162 in chronic sleep disorders and related clinical manufacturing costs

Approximately \$5.0 million to initiate a Phase II trial for VSF-173 in excessive sleepiness

We anticipate that the balance of such net proceeds will be used for general research and development, business development and other corporate purposes as determined by our management, including for working capital, milestone payments under our existing license agreements, to the extent they become due. We may also use proceeds from this offering for the acquisition or licensing of businesses or product candidates that are complementary to our own. However, due to the uncertainties inherent in the clinical trial process and given that our product candidates have not completed their clinical development, we are unable to estimate precisely the total costs that will be associated with completing the above-mentioned clinical trials, and accordingly we cannot estimate precisely what proceeds will be available for general corporate purposes. The actual amounts could vary materially from our estimates. Currently, we have no specific plans or commitments with respect to any acquisition or license; however, we intend to engage an investment bank to provide financial and strategic advisory services to the Company, which may lead to one or more possible transactions, including the acquisition, sale or licensing by the Company of businesses or product candidates, the sale or licensing to a third party of one or more of our own product candidates, or the acquisition of the Company. We cannot assure you that we will complete any acquisitions, sales or licenses or that, if completed, any acquisition, sale or license will be successful or on attractive terms.

The amount and timing of our actual expenditures will depend on numerous factors, including the progress of our research and development activities and clinical trials, the number and breadth of our product development programs, our ability to establish and maintain corporate collaborations and other arrangements and the amount of cash, if any, generated by our operations.

We will retain broad discretion in the allocation and use of the remaining net proceeds of this offering. Pending application of the net proceeds, as described above, we intend to invest any remaining proceeds in short-term, investment-grade, interest-bearing securities.

# Price range of our common stock

Our common stock is quoted on The Nasdaq Global Market under the symbol VNDA. The following table sets forth, for the periods indicated, the range of high and low closing sale prices of our common stock as reported on The Nasdaq Global Market.

	High	Low
Second quarter 2006	\$ 11.26	\$ 7.99
Third quarter 2006	\$ 10.08	\$ 8.22
Fourth quarter 2006	\$ 26.17	\$ 9.06

The last reported sale price of our common stock on January 3, 2007 was \$24.83 per share.

As of December 31, 2006, there were 42 holders of record of our common stock.

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# **Dividend policy**

We have never declared or paid any dividends on our capital stock. We currently intend to retain any future earnings to finance our research and development efforts, the further development of our pharmacogenetics and pharmacogenemics expertise and the expansion of our business and do not intend to declare or pay cash dividends on our common stock in the foreseeable future. Any future determination to pay dividends will be at the discretion of our board of directors and will depend upon results of operations, financial condition, contractual restrictions, restrictions imposed by applicable law and other factors our board of directors deem relevant.

## Capitalization

The following table sets forth our actual capitalization as of September 30, 2006:

#### On an actual basis

On an as-adjusted basis to give effect to the sale of 3,500,000 shares of common stock that we are offering at \$24.83 per share, based upon the last reported sale price of our common stock on The Nasdaq Global Market on January 3, 2007, after deducting underwriting discounts and commissions and estimated offering expenses payable by us

You should read this table together with our 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.

The table excludes the following shares:

1,569,669 shares of common stock issuable upon the exercise of stock options outstanding as of September 30, 2006 under our Second Amended and Restated Management Equity Plan and agreements entered into pursuant to such plan

103,692 shares of common stock issuable upon the exercise of stock options outstanding as of September 30, 2006 under the 2006 Equity Incentive Plan and agreements entered into pursuant to such plan

an additional 1,396,308 shares reserved for issuance under the 2006 Equity Incentive Plan as of September 30, 2006 for future stock option grants and purchases under our equity compensation plans (see note 4 of Notes to condensed consolidated financial statements )

See Equity benefit plans, and Note 10 of Notes to consolidated financial statements for a description of our equity plans.

	Actual	As adjusted
Stockholders equity: Common stock, \$0.001 par value; 150,000,000 shares authorized, 21,907,188 shares issued and outstanding (actual); 25,407,188 shares issued and outstanding (as adjusted)	\$ 21,907	\$ 25,407
Additional paid-in capital Accumulated other comprehensive loss Deficit accumulated during the development stage	124,893,956 (15,130) (87,949,101)	205,331,156 (15,130) (87,949,101)
Total stockholders equity	36,951,632	117,392,332

#### **Dilution**

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

As of September 30, 2006, our net tangible book value was approximately \$36,951,632, or \$1.69 per share, based on 21,907,188 shares of our common stock outstanding as of September 30, 2006. Our net tangible book value per share represents the amount of our total tangible assets reduced by the amount of our total liabilities, divided by the number of shares of our common stock outstanding as of September 30, 2006.

After giving effect to our sale in this offering of 3,500,000 shares of our common stock at an assumed public offering price of \$24.83 per share, based on the last reported sale price of our common stock on The Nasdaq Global Market on January 3, 2007, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us, our as-adjusted net tangible book value as of September 30, 2006 would have been approximately \$117,392,332, or \$4.62 per share of our common stock. This represents an immediate increase of net tangible book value of \$2.93 per share to our existing stockholders and an immediate dilution of \$20.24 per share to investors purchasing shares in this offering.

The following table illustrates this per share dilution:

Assumed public offering price per share	\$ 24.83
Net tangible book value per share as of September 30, 2006	\$ 1.69
Increase in net tangible book value per share attributable to this offering	\$ 2.93
As-adjusted net tangible book value per share after giving effect to this offering	\$ 4.62
Dilution per share to new investors	\$ 20.21

If the underwriters exercise their option to purchase additional shares of our common stock in full in this offering, the as-adjusted net tangible book value per share after the offering would be \$5.00 per share, the increase in net tangible book value per share attributable to this offering would be \$3.31 per share, and the dilution to new investors purchasing shares in this offering would be \$19.83 per share.

#### The table above excludes:

1,569,669 shares of common stock issuable upon the exercise of stock options outstanding as of September 30, 2006 under our Second Amended and Restated Management Equity Plan and agreements entered into pursuant to such plan, with a weighted-average exercise price of \$1.50 per share

103,692 shares of common stock issuable upon the exercise of stock options outstanding as of September 30, 2006 under the 2006 Equity Incentive Plan and agreements entered into pursuant to such plan, with a weighted-average exercise price of \$8.73 per share

an additional 1,396,308 shares reserved for issuance under the 2006 Equity Incentive Plan as of September 30, 2006 for future stock option grants and purchases under our equity compensation plans (see note 4 of Notes to condensed consolidated financial statements )

#### Selected consolidated financial data

The consolidated statements of operations data for the period of March 13, 2003 (inception) to December 31, 2003 and the years ended December 31, 2004 and 2005 and the consolidated balance sheet data as of December 31, 2004 and 2005 are each derived from our audited consolidated financial statements included in this prospectus. The consolidated statements of operations data for the nine months ended September 30, 2005 and 2006 and the consolidated balance sheet data as of September 30, 2006 are each derived from our unaudited condensed consolidated financial statements included in this prospectus. The unaudited condensed consolidated financial data include, in the opinion of our management, all adjustments, consisting only of normal recurring adjustments, that are necessary for a fair presentation of our financial position and results of operations for these periods. Our historical results for any prior period are not necessarily indicative of results to be expected in any future period, and our results for any interim period are not necessarily indicative of results for a full fiscal year.

The following data should be read together with our consolidated financial statements and accompanying notes and the section entitled Management's discussion and analysis of financial condition and results of operations included in this prospectus.

	(inc	eriod from March 13, 2003 ception) to ember 31, 2003		Year ende 2004	ed De	ecember 31, 2005		ended 2005	Nine 1 I Septem	months ber 30, 2006
Statements of operations data Revenue Operating expenses:	\$	47,565	\$	33,980	\$		\$		\$	
Research and development General and administrative		2,010,532 1,052,659		7,442,983 2,119,394		16,890,615 7,396,038		,641,565 ,587,147		30,788
Total operating expenses		3,063,191		9,562,377		24,286,653	17	,228,712	53,3	301,227
Loss from operations Interest and other income, net		(3,015,626) 44,805	(	59,528,397)		(24,286,653) 410,001	(17	,228,712)		301,227) 581,534
Net loss before tax provision		(2,970,821)	(	(9,469,337)		(23,876,652)	(17	,040,424)	(51,6	519,693)

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Tax provision		4,949	7,649		
Net loss Beneficial conversion feature deemed dividend	(2,970,821)	(9,474,286)	(23,884,301)	(17,040,424)	(51,619,693)
to preferred stockholders(1)			(33,486,623)	(18,500,005)	
Net loss attributable to common stockholders	\$ (2,970,821)	\$ (9,474,286)	\$ (57,370,924)	\$ (35,540,429)	\$ (51,619,693)
Net loss per share applicable to common stockholders, basic and diluted Weighted average number of shares used in	\$ (983.72)	\$ (3,137.18)	\$ (3,374.33)	\$ (3,094.51)	\$ (3.72)
computing net loss per share, basic and diluted	3,020	3,020	17,002	11,485	13,862,613

<sup>(1)</sup> In September and December of 2005, we completed the sale of an additional 27,235,783 shares of Series B Preferred Stock for net proceeds of approximately \$33.5 million. After evaluating the fair value of the common stock obtainable upon conversion by the stockholders, we determined that the issuance of the Series B Preferred Stock sold in 2005 resulted in a beneficial conversion feature which was fully accreted in 2005 and is recorded as a deemed dividend to preferred stockholders of approximately \$33.5 million and approximately \$18.5 million for the year ended December 31, 2005 and the nine months ended September 30, 2005, respectively.

		As of December 31,	As of September 30,
	2004	2005	2006
Balance sheet data			
Cash and cash equivalents and restricted cash	\$ 16,259,770	\$ 21,443,045	\$ 32,330,209
Short-term investments		10,141,189	11,096,506
Working capital	14,827,621	28,308,434	34,735,547
Total assets	17,752,241	35,752,770	47,282,498
Total liabilities	1,808,654	5,087,963	10,330,866
Convertible preferred stock	28,308,564	61,795,187	
Deficit accumulated during the development			
stage	(12,445,107)	(36,329,408)	(87,949,101)
Total stockholders equity	15,943,587	30,664,807	36,951,632
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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 Selected Consolidated Financial Data and our consolidated financial statements and related notes appearing at the end of this prospectus. Some of the information contained in this discussion and analysis or set forth elsewhere in this prospectus include historical information and other information with respect to our plans and strategy for our business and contain forward-looking statements that involve risk, uncertainties and assumptions. Our actual results may differ materially from those anticipated in these forward-looking statements as a result of certain factors, including but not limited to those set forth under the Risk factors section of this prospectus and elsewhere in this prospectus.

#### Overview

Vanda was founded in November 2002 and commenced its operations on March 13, 2003. We are a biopharmaceutical company focused on the development and commercialization of clinical-stage product candidates for central nervous system disorders, with exclusive worldwide commercial rights to three product candidates in clinical development for various central nervous system disorders. Our lead product candidate, iloperidone, is a compound for the treatment of schizophrenia and bipolar disorder. In December 2006 we announced positive top-line results from our Phase III trial of iloperidone for schizophrenia. Our second product candidate, VEC-162, is a compound for the treatment of sleep and mood disorders. In November 2006 we announced positive top-line results from our Phase III trial of VEC-162 in transient insomnia. VEC-162 is also ready for Phase II trials for the treatment of depression. Our third product candidate, VSF-173, is a compound for the treatment of excessive sleepiness and is ready for a Phase II trial.

We expect to file a New Drug Application (NDA) for iloperidone in schizophrenia with the United States Food and Drug Administration (FDA) by the end of 2007. We expect to meet with the FDA in the first quarter of 2007 regarding this filing. We will have to conduct additional Phase III trials for VEC-162 in chronic sleep disorders prior to our filing of an NDA for VEC-162, and we expect to begin at least one of these additional trials in the second half of 2007. We also expect to begin a Phase II trial of VSF-173 for excessive sleepiness in mid-2007. Assuming successful outcomes of our clinical trials and approval by the FDA, we expect to commercialize iloperidone and VSF-173 with our own sales force in the U.S., and expect to commercialize VEC-162 through a partnership with a global pharmaceutical company, although we have not yet identified such a global partner.

We are a development-stage company and have accumulated net losses of approximately \$87.9 million since the inception of our operations through September 30, 2006. We have no product revenues to date and have no approved products for sale. Since we began our operations in March 2003, we have devoted substantially all of our resources to the in-licensing and clinical development of our product candidates. Our future operating results will depend largely on our ability to develop and commercialize our lead product candidate, iloperidone, successfully, and on the progress of other product candidates currently in our research and development pipeline. The results of our operations will vary significantly from year-to-year and quarter-to-quarter and depend on a number of factors, including risks related to our business, risks related to our industry, and other risks which are detailed in the Risk factors section of this prospectus.

Based on our current operating plans, and assuming the sale of 3,500,000 shares of our common stock in this offering at \$24.83 per share (the last reported sale price of our common stock on The Nasdaq Global Market on January 3, 2007), we believe that the proceeds from this offering, together with our existing cash, restricted cash, cash equivalents and short-term investments, will be sufficient to meet our anticipated operating needs through early 2008, and after that time we will require additional capital. In budgeting for our activities following this offering, we have relied on a number of assumptions, including assumptions that we will file an NDA for iloperidone in schizophrenia with the FDA by the end of 2007, that we will continue to expend funds in preparation of the commercial launch of iloperidone, that we will initiate at least one additional VEC-162 Phase III trial in chronic sleep disorders in the second half of 2007 and that this trial will be conducted in accordance with our expectations, that we will initiate our VSF-173 Phase II trial for excessive sleepiness in mid-2007 and that this trial will be conducted in accordance with our expectations, that we will expend funds on the extended-release injectable formulation of iloperidone, that we will not engage in further in-licensing activities, that we will not receive any proceeds from potential partnerships, that we will not expend funds on the bipolar indication for iloperidone or on a Phase II trial of VEC-162 for depression, that we will continue to evaluate pre-clinical compounds for potential development, that we will be able to continue the manufacturing of our product candidates at commercially reasonable prices, that we will be able to retain our key personnel, and that we will not incur any significant contingent liabilities. We may need to raise additional funds more quickly if one or more of our assumptions proves to be incorrect or if we choose to expand our product development efforts more rapidly than presently anticipated or seek to acquire additional product candidates, and we may also decide to raise additional funds even before they are needed if the conditions for raising capital are favorable. We may seek to sell additional equity or debt securities or obtain a bank credit facility. The sale of additional equity or debt securities, if convertible, could result in dilution to our stockholders. The incurrence of indebtedness would result in increased fixed obligations and could also result in covenants that would restrict our operations.

On April 18, 2006 we consummated our initial public offering, consisting of 5,750,000 shares of common stock. On April 21, 2006 the underwriters exercised an over-allotment option to purchase additional 214,188 shares of our common stock. Including the over-allotment shares, the offering totaled 5,964,188 shares of common stock at a public offering price of \$10.00, resulting in net proceeds to the Company of approximately \$53.3 million (after deducting underwriters discounts and commissions as well as offering expenses).

In connection with the initial public offering, the Company effected a 1-for-3.309755 reverse stock split of the issued and outstanding common stock. Information in this prospectus relating to common stock and common stock-equivalents (including the share numbers in the preceding paragraph) has been restated to reflect this split for all periods presented. Upon completion of the initial public offering, all shares of the Company s Series A Preferred Stock and Series B Preferred Stock were converted into an aggregate of 15,794,632 shares of common stock.

Phase III trial for iloperidone. We reported positive top-line results from our Phase III trial of iloperidone in schizophrenia in December 2006. The primary endpoint of the trial was efficacy versus placebo on the Positive and Negative Symptoms Scale (PANSS), for which iloperidone demonstrated statistically significant improvement. Iloperidone also demonstrated statistically significant improvement versus placebo in several other measures of efficacy. Iloperidone also appeared to be safe and well-tolerated in the trial, which reinforced the results of three short-term and three long-term clinical trials of iloperidone comprising a total of over 2,000 patients, in which iloperidone differentiated itself from currently available atypical antipsychotics by offering a number of reduced side effects.

Prior to September 30, 2006 we incurred approximately \$30.2 million in clinical costs related to this trial. We currently expect that between October 1, 2006 and December 31, 2006 we will incur approximately \$2.0 million in additional clinical costs related to the trial. In 2007, we expect that we will incur approximately \$2.0 million to \$3.0 million in costs related to the trial and for services rendered to us in connection with the analysis of trial data and the preparation of regulatory filings. We expect to make a New Drug Application filing for iloperidone by the end of 2007 and we would then expect to launch iloperidone commercially in early 2009. However, the time it takes to receive cash inflows from the sale of iloperidone are highly dependent on facts and circumstances that we may not be able to control and are subject to a number of risks. For example, delays in the approval process and subsequent commercial launch of iloperidone following our filing may occur if the FDA fails to attend to our filing in a timely manner or requires further data to approve iloperidone. Please see the Risk factors section of this prospectus for a more detailed discussion of these and other risks.

Phase III trial for VEC-162 in insomnia. In November 2006 we announced positive top-line results from our Phase III trial of VEC-162 in the treatment of transient insomnia. VEC-162 demonstrated statistically significant improvement in several parameters used to measure the efficacy of insomnia therapies, including reduced duration of wake after sleep onset, improved sleep efficiency and shortened time to persistent sleep. In addition, VEC-162 also appeared to be safe and well-tolerated in the trial.

Prior to September 30, 2006 we incurred approximately \$6.0 million in clinical costs related to this trial. We expect that between October 1, 2006 and December 31, 2006 we will incur approximately \$1.0 million in clinical costs related to the trial, related administrative services and for services rendered to us in connection with the analysis of trial data. In 2007, we expect that we will incur less than \$0.5 million in costs related to the trial. We believe that we will have to conduct additional Phase III trials in chronic sleep disorders to receive FDA approval of VEC-162 for the treatment of insomnia. We expect to begin at least one of these additional trials in the second half of 2007.

Revenues. We generated some revenue during the period from March 13, 2003 (inception) to December 31, 2003 and during the year ended December 31, 2004 under research and development contracts that were derived principally from consulting agreements we entered into during our start-up phase to defray research costs. We completed our obligations during those periods under these agreements and no longer seek such arrangements.

We have not generated any other operating revenue since our inception. Any revenue that we may receive in the near future is expected to consist primarily of license fees, milestone payments and research and development reimbursement payments to be received from partners. If our development efforts result in clinical success, regulatory approval and successful commercialization of our products, we could generate revenue from sales of our products and from receipt of royalties on sales of licensed products.

Research and development expenses. The Company s research and development expenses consist primarily of fees paid to third-party professional service providers in connection with the services they provide for our clinical trials, costs of contract manufacturing services, costs of materials used in clinical trials and research and development, depreciation of capital resources used to develop our products, and all related facilities costs. We expense research and development costs as incurred, including payments made to date under our license agreements. We believe that significant investment in product development is a competitive necessity and plan to continue these investments in order to realize the potential of our product candidates and pharmacogenetics and pharmacogenomics expertise. From inception through September 30,

2006 we incurred research and development expenses in the aggregate of approximately \$70.5 million, including stock-based compensation expenses of approximately \$1.3 million. We expect our research and development expenses to increase as we continue to develop our product candidates and we also expect to incur licensing costs in the future that could be substantial, as we continue our efforts to evaluate potential in-license product candidates.

The following table summarizes our product development initiatives for the period from March 13, 2003 (inception) to December 31, 2003, for the years ended December 31, 2004 and December 31, 2005, for the nine months ended September 30, 2005 and September 30, 2006, and for the period from March 13, 2003 (inception) to September 30, 2006. Included in this table are the research and development expenses recognized in connection with our product candidates in clinical development. Included in Other product candidates are the costs directly related to research initiatives for all other product candidates.

	March 13, 2003 (inception)			Nine months	Nine months	Period from March 13, 2003
	to	Year ended December 31, 2004	Year ended December 31, 2005	ended September 30, 2005	ended September 30, 2006	(inception) to September 30, 2006
Direct project costs(1)				<b>.</b>	<b>.</b>	
Iloperidone		\$ 1,123,000	\$ 7,798,000	\$ 4,423,000	\$ 31,478,000	\$ 40,398,000
VEC-162		3,221,000	6,133,000	5,057,000	9,559,000	18,912,000
VSF-173		568,000	943,000	707,000	849,000	2,360,000
Other product candidates		1,037,000	899,000	608,000	873,000	2,810,000
Total direct product costs	\$	5,949,000	15,773,000	10,795,000	42,759,000	64,480,000
	φ	3,949,000	13,773,000	10,793,000	42,739,000	04,460,000
Indirect project costs(1)						
Facility(3)		259,000	247,000	185,000	447,000	952,000
Depreciation	69,000	345,000	375,000	281,000	350,000	1,139,000
Other indirect	,	,	- · · · <b>,</b> · · · ·	,,,,,,	,	,,
overhead	1,941,000	890,000	496,000	380,000	575,000	3,904,000
Total indirect						
expenses	2,010,000	1,494,000	1,118,000	846,000	1,372,000	5,995,000
Total research & development						
expenses	\$ 2,010,000	\$ 7,443,000	\$ 16,891,000	\$ 11,641,000	\$ 44,131,000	\$ 70,475,000

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- (1) Many of our research and development costs are not attributable to any individual project because we share resources across several development projects. We record direct costs, including personnel costs and related benefits and stock-based compensation, on a project-by-project basis. We record indirect costs that support a number of our research and development activities in the aggregate.
- (2) In 2003, there were no active development programs in process for our product candidates listed in the table.
- (3) In 2003, all facility-related costs were allocated to general and administrative expenses.

General and administrative expenses. General and administrative expenses consist primarily of salaries and other related costs for personnel serving executive, finance, accounting, information technology, marketing and human resource functions. Other costs include facility costs not otherwise included in research and development expense and professional fees for legal and accounting services. We expect that our general and administrative expenses will increase as we add personnel and fulfill our reporting obligations applicable to public companies, including the compliance with Section 404 of the Sarbanes-Oxley Act. From inception through September 30, 2006, we incurred general and administrative expenses in the aggregate of approximately \$19.7 million, including stock-based compensation expenses of approximately \$8.4 million.

Stock-based compensation. We adopted SFAS 123(R), Share Based Payment, on January 1, 2006 using the modified prospective method of implementation and adopted the accelerated vesting method. Prior to January 1, 2006 we followed APB Opinion No. 25, Accounting for Stock Issued to Employees (APB 25), and related interpretations, in accounting for our stock-based compensation plans, rather than the alternative fair value accounting method provided for under SFAS No. 123, Accounting for Stock-Based Compensation. In the notes to our financial statements we provide pro forma disclosures in accordance with SFAS No. 123 and related pronouncements for the periods prior to adoption of SFAS 123(R).

Factors which affect charges or credits to operations related to stock-based compensation are the fair value of the common stock underlying stock options for which stock-based compensation is recorded, the volatility of such fair value, and risk-free rate, expected dividend yield and expected life of the option used in the calculation of the fair value of the stock option. The stock-based compensation expense for a period is also affected by expected forfeiture rate for the respective option grants. If our estimates of the fair value of these equity instruments are too high or too low, it would have the effect of overstating or understating expenses.

On April 12, 2006 our common stock began trading on The Nasdaq Global Market. Prior to April 12, 2006, given the absence of an active market for our common stock, the exercise price of our stock options on the date of grant was determined by our board of directors using several factors, including progress and milestones achieved in our business development and performance, the price per share of our convertible preferred stock offerings, the perspectives provided by our underwriters regarding estimates of a potential price per share in an initial public offering of our common stock and general industry and economic trends. In establishing our estimates of fair value, we considered the guidance set forth in the AICPA Practice Guide, *Valuation of Privately-Held-Company Equity Securities Issued as Compensation* and made a retrospective determination of fair value. The exercise price for employee options granted after April 12, 2006 is based on the market price of our common stock.

Stock-based compensation expense recognized in accordance to APB 25 prior to January 1, 2006 related to employee stock options granted below fair market value and modifications of employee stock option awards. We recorded stock-based compensation expense of approximately \$23,000 and approximately \$1.3 million in respect of the options granted below fair value for the years ended December 31, 2004 and 2005, respectively.

In August 2004 we approved a modification to an employee s stock option award at the time of employment termination. The modification was to accelerate a portion of the unvested stock options so the shares could be immediately exercisable. According to FASB Interpretation No. 44, *Accounting for Certain Transactions involving Stock Compensation* (FIN 44), the result of such a modification is to remeasure the stock options that were modified. The remeasurement of the stock options resulted in an immediate charge of approximately \$15,000, which was included in general and administrative expense for the year ended December 31, 2004.

In February 2005 the board of directors approved a modification to all outstanding granted stock option awards, repricing the options from their original exercise price of \$1.32 to \$0.33. According to FIN 44, the result of such a modification is to account for the modified stock option awards as variable from the date of the modification to the date the awards are exercised, forfeited, or cancelled. For the year ended December 31, 2005, we remeasured approximately 335,000 outstanding stock options, resulting in initial deferred stock compensation of approximately \$1.7 million. Compensation expense relating to the remeasurement of modified stock options was approximately \$3.8 million for the year ended December 31, 2005,

which includes approximately \$3.1 million of immediate stock compensation charges for vested shares at the time of remeasurement for the year ended December 31, 2005.

According to EITF 00-23, Issues Related to the Accounting for Stock Compensation under APB Opinion No. 25 and FASB Interpretation No. 44, FASB Interpretation No. 28, Accounting for Stock Appreciation Rights and Other Variable Stock Option or Award Plans and interpretation of APB Opinions No. 15 and 25 (FIN 28), is required for variable awards. FIN 28 specifies that compensation should be measured at the end of each period as the amount by which the quoted market value of the shares of the enterprise s stock covered by the grant exceeds the option price or value specified under the plan and that amount should be accrued as a charge to expense over the periods the employee performs the related services.

Stock-based compensation expense recognized after January 1, 2006 is based on the value of the portion of stock-based payment awards that is ultimately expected to vest during the period and includes:

compensation expense for stock-based payment awards granted prior to, but not yet vested as of, December 31, 2005 based on the grant date fair value estimated in accordance with the pro forma provisions of SFAS 123

compensation expense for stock-based payment awards granted subsequent to December 31, 2005 based on the grant date fair value estimated in accordance with SFAS 123(R)

Total stock-based compensation expense, related to all of the Company s stock-based awards, recognized under SFAS 123(R) and APB 25, respectively, was comprised of the following:

	March 13, 2003 (inception) to December 31, 2003	Year ended December 31, 2004	Year ended December 31, 2005	Nine months  ended September 30, 2005	Nine months ended September 30, 2006
Research and development General and administrative	\$	\$ 2,000	\$ 789,000 4,313,000	\$ 659,000 3,431,000	\$ 476,000 4,013,000
Total stock-based compensation expense	e \$	\$ 38,000	\$ 5,102,000	\$ 4,090,000	\$ 4,489,000

Beneficial conversion feature. In September 2005 we completed the sale of an additional 15,040,654 shares of Series B Preferred Stock for proceeds of approximately \$18.5 million. After evaluating the fair value of our common stock obtainable upon conversion by the stockholders, we determined that the issuance of the Series B Preferred Stock sold in September 2005 resulted in a beneficial conversion feature calculated in accordance with EITF Issue No. 98-5, Accounting for Convertible Securities with Beneficial Conversion Features or Contingently Adjustable Conversion Ratios, as interpreted by EITF Issue No. 00-27, Application of Issue No. 98-5 to Certain Convertible Instruments, of approximately \$18.5 million which was fully accreted in September 2005 and is recorded as a deemed dividend to

preferred stockholders for the year ended December 31, 2005. Likewise, in December 2005, we completed the sale of an additional 12,195,129 shares of Series B Preferred Stock for additional proceeds of approximately \$15.0 million. After evaluating the fair value of our common stock obtainable upon conversion by the stockholders, we determined that the issuance of the Series B Preferred Stock sold in December 2005 resulted in a beneficial conversion feature calculated in accordance with EITF Issue No. 98-5,

as interpreted by EITF Issue No. 00-27, approximately \$15.0 million of which was fully accreted in December 2005 and is recorded as a deemed dividend to preferred stockholders for the year ended December 31, 2005.

Interest and other income, net. Interest income consists of interest earned on our cash, restricted cash and cash equivalents and short-term investments. Interest expense consists of interest incurred on equipment debt. Other expense, net, consists of foreign currency loss related to our wholly-owned foreign subsidiary located in Singapore.

Operations. We have a limited history of operations. We anticipate that our quarterly results of operations will fluctuate for the foreseeable future due to several factors, including any possible payments made or received pursuant to licensing or collaboration agreements, progress of our research and development efforts, and the timing and outcome of clinical trials and related possible regulatory approvals. Our limited operating history makes predictions of future operations difficult or impossible. Since our inception, we have incurred significant losses. As of September 30, 2006, we had a deficit accumulated during the development stage of approximately \$87.9 million. We anticipate incurring additional losses, which may increase, for the foreseeable future.

## **Results of operations**

# Nine months ended September 30, 2006 compared to nine months ended September 30, 2005

Research and development expenses. Research and development expenses increased by approximately \$32.5 million, or 279%, to approximately \$44.1 million for the nine months ended September 30, 2006 compared to approximately \$11.6 million for the nine months ended September 30, 2005. Research and development expense consists of direct costs which include salaries and related costs of research and development personnel, stock-based compensation, the costs of consultants, materials and supplies associated with research and development projects, as well as clinical activities. Indirect research and development costs include facilities, depreciation, and other indirect overhead costs.

The following table discloses the components of research and development expenses reflecting all of our project expenses:

	1	Nine months ended September 30,
Research and development expenses	2005	2006
Direct project costs:		
Clinical trials	\$ 4,101,000	\$ 33,055,000
Contract research and development, consulting, materials and other costs	4,496,000	6,855,000
Salaries, benefits and related costs	1,539,000	2,373,000
Stock-based compensation	659,000	476,000
Total direct costs	10,795,000	42,759,000
Indirect project costs	846,000	1,372,000
Total	\$ 11,641,000	\$ 44,131,000

Direct costs increased approximately \$32.0 million primarily as a result of clinical development activities for iloperidone and VEC-162. Clinical trials expense increased approximately

\$29.0 million for the nine months ended September 30, 2006 primarily due to the cost incurred in our Phase III iloperidone and VEC-162 clinical trials that began in the fourth quarter of 2005 and in the first quarter of 2006, respectively. Contract research and development, consulting, materials and other costs increased approximately \$2.4 million for the nine months ended September 30, 2006, primarily as a result of a \$1.0 million milestone payment under our license agreement for VEC-162 with Bristol-Myers Squibb and due to increased regulatory and manufacturing-related development costs incurred in connection with the manufacturing of clinical supply materials for the iloperidone and the VEC-162 clinical trial programs. Prior to FDA approval of our products, manufacturing-related costs are included in research and development expense. Salaries, benefits and related costs increased approximately \$834,000 for the nine months ended September 30, 2006 due to an increase in personnel to support the development and clinical trial activities for iloperidone and VEC-162. The stock-based compensation expense decreased approximately \$183,000 primarily as a result of the expense included in the modification of stock option awards recorded in 2005. Indirect project costs also increased by approximately \$526,000 for the nine months ended September 30, 2006 due primarily to the increase in facility rent expense.

We expect to continue to incur substantial research and development expenses due to our ongoing research and development efforts and as our existing and future product candidates proceed through clinical trials.

*General and administrative expenses*. General and administrative expenses increased approximately \$3.6 million, or 64%, to approximately \$9.2 million for the nine months ended September 30, 2006 from approximately \$5.6 million for the nine months ended September 30, 2005.

The following table discloses the components of our general and administrative expenses:

	Nine	e months ended September 30,
General and administrative expenses	2005	2006
Salaries, benefits and related costs	\$ 946,000	\$ 1,746,000
Stock-based compensation	3,431,000	4,013,000
Legal and consulting expenses	680,000	1,363,000
Other expenses	530,000	2,048,000
Total	\$ 5,587,000	\$ 9,170,000

General and administrative expenses consist of professional fees, salaries and related costs for executive and other administrative personnel and facility costs. Salaries, benefits and related costs increased approximately \$800,000 for the nine months ended September 30, 2006 due to an increase in personnel as we continued to develop the administrative, business development and other functions required to support the development and clinical trial activities for iloperidone, VEC-162 and our other product candidates. Stock-based compensation expense increased by approximately \$582,000 due to new option grants in late 2005 and in 2006.

Legal and consulting expenses increased approximately \$683,000 for the nine months ended September 30, 2006 due primarily to a higher level of consulting activity in 2006 in support of business development and market research activities related to our lead product candidates as well as an increase in legal, accounting and other professional expenses associated with being a public company. Other expenses increased approximately \$1.5 million for the nine

months ended September 30, 2006, due to an increase in facilities expenses of approximately \$429,000,

which includes expenses relating to abandonment of our former office facilities of approximately \$267,000, an increase in insurance expenses of approximately \$515,000, primarily due to an increase in directors and officers and clinical trial insurance, and an increase in other general and administrative expenses.

We expect our general and administrative expenses to continue to increase as we support our discovery and development efforts, continue with our commercial development activities and fulfill our reporting and other regulatory obligations applicable to public companies, including the compliance with Section 404 of the Sarbanes-Oxley Act.

Interest income, net. Net interest income in the nine months ended September 30, 2006 was approximately \$1.7 million compared to net interest income of approximately \$188,000 in the nine months ended September 30, 2005. Interest income was higher in 2006 due to higher average cash balances for the period and higher short-term interest rates which generated substantially higher interest income than it did in 2005.

Our interest income and expense for the nine months ended September 30, 2006 and September 30, 2005 are as follows:

	Nine months ended September 30,		
	2005	2006	
Interest income	\$ 209,000	\$ 1,686,000	
Interest expense	(21,000)	(5,000)	
Total, net	\$ 188,000	\$ 1,681,000	

#### Year ended December 31, 2005 compared to year ended December 31, 2004

*Revenues*. Revenues decreased approximately \$34,000 for the year ended December 31, 2005 to zero. Revenue earned in 2004 was derived principally from consulting agreements we entered into during our start-up phase under research and development contracts. We have completed our obligations under these agreements and no longer seek such arrangements.

Research and development expenses. Research and development expenses increased by approximately \$9.5 million, or 128%, to approximately \$16.9 million for the year ended December 31, 2005 compared to approximately \$7.4 million for the year ended December 31, 2004. Research and development expenses consist of direct costs which include salaries and related costs of research and development personnel, stock-based compensation, and the costs of consultants, materials and supplies associated with research and development projects, as well as clinical activities. Indirect research and development costs include facilities, depreciation, and other indirect overhead costs.

The following table discloses the components of research and development expenses reflecting all of our project expenses:

		Year ended December 31,
Research and development expenses	2004	2005
Direct project costs:		
Clinical trials	\$ 916,000	\$ 6,305,000
Contract research and development, consulting, materials and other costs	3,876,000	6,747,000
Salaries, benefits and related costs	1,155,000	1,962,000
Stock-based compensation	2,000	789,000
Total direct costs	5,949,000	15,803,000
Indirect project costs	1,494,000	1,088,000
Total	\$ 7,443,000	\$ 16,891,000

Direct costs increased approximately \$9.9 million primarily as a result of approximate increases of \$6.7 million, \$2.9 million and \$375,000, relating to clinical development activities for iloperidone, VEC-162 and VSF-173, respectively. During the year ended December 31, 2005, we conducted additional clinical development and manufacturing work on iloperidone as we prepared for and commenced its Phase III trial. We also conducted a Phase II clinical trial for VEC-162. Salaries, benefits and related costs increased approximately \$807,000 for the year ended December 31, 2005 due to an increase in personnel to support the development and clinical trial activities for iloperidone and VEC-162.

Contract research and development, consulting, materials and other costs increased approximately \$2.9 million for the year ended December 31, 2005, primarily due to regulatory and manufacturing-related development costs of approximately \$2.9 million incurred in connection with the manufacturing of clinical supply materials for the iloperidone Phase III and the VEC-162 clinical trial programs. Prior to FDA approval of our products, manufacturing-related costs are included in research and development expense. Clinical trials expense increased approximately \$5.4 million for the year ended December 31, 2005 primarily due to the cost incurred as we prepared for and commenced our Phase III iloperidone clinical trial that began in the fourth quarter of 2005 and the costs related to the Phase II VEC-162 trial that was conducted in 2005. Stock-based compensation expense increased by approximately \$787,000 due to expenses relating to employee stock options granted below fair market value and modifications of employee stock option awards. Indirect project costs decreased by approximately \$406,000 for the year ended December 31, 2005 due primarily to the elimination of contract manufacturing activities we previously conducted.

In 2006 and thereafter we expect research and development expenses to continue to increase substantially as we increase our research and development efforts and as our existing and future product candidates proceed through clinical trials.

*General and administrative expenses.* General and administrative expenses increased approximately \$5.3 million, or 249%, to approximately \$7.4 million for the year ended December 31, 2005 from approximately \$2.1 million for the year ended December 31, 2004.

The following table discloses the components of our general and administrative expenses:

	Year ended December 31,		
General and administrative expenses		2004	2005
Salaries, benefits and related costs	\$	906,000	\$ 1,411,000
Stock-based compensation		36,000	4,313,000
Legal and consulting expenses		690,000	899,000
Other expenses		487,000	773,000
Total	\$ 2	2,119,000	\$ 7,396,000

General and administrative expenses consist of professional fees, salaries and related costs for executive and other administrative personnel and facility costs. Salaries, benefits and related costs increased approximately \$505,000 for the year ended December 31, 2005 due to an increase in personnel as we continued to develop the administrative structure to support the development and clinical trial activities for iloperidone, VEC-162 and our other product candidates. Stock-based compensation expense increased by approximately \$4.3 million due to expenses relating to employee stock options granted below fair market value and modifications of employee stock option awards.

Legal and consulting expenses increased approximately \$209,000 for the year ended December 31, 2005 due primarily to a higher level of consulting activity in 2005 in support of business development and market research activities related to our lead product candidates. Other expenses increased approximately \$286,000 for the year ended December 31, 2005, primarily due to increased insurance and taxes.

In 2006 and thereafter we expect our general and administrative expenses to increase substantially. These increased expenses are expected to be necessary to support our discovery and development efforts and our commercial development activities and to fulfill our reporting and other regulatory obligations applicable to public companies.

*Interest income, net.* Net interest income in the year ended December 31, 2005 was approximately \$410,000 compared to net interest income of approximately \$59,000 in the year ended December 31, 2004. Interest income was higher in 2005 due to higher average cash balances for the year and higher short-term interest rates which generated substantially higher interest income than in 2004.

Our interest income and expense for the year ended December 31, 2004 and the year ended December 31, 2005 are as follows:

		Year ended December 31,		
	2004	2005		
Interest income	\$ 101,000	\$ 436,000		

Interest expense		(42,000)	(26,000)
Total, net		\$ 59,000	\$ 410,000
	42		

#### Year ended December 31, 2004 compared to period from March 13, 2003 (inception) to December 31, 2003

*Revenues*. We recorded revenues of approximately \$34,000 and approximately \$48,000 for 2004 and 2003, respectively. Revenue earned in 2004 and 2003 was derived principally from consulting agreements we entered into during our start-up phase under research and development contracts. We completed our obligations under these agreements and no longer seek such arrangements.

Research and development expenses. Research and development expenses increased approximately \$5.4 million, or 270%, to approximately \$7.4 million for the year ended December 31, 2004 compared to approximately \$2.0 million for the period from March 13, 2003 (inception) to December 31, 2003.

The following table discloses the components of research and development expenses reflecting all of our project expenses:

Research and development expenses	Period from March 13, 2003 (inception) to December 31, 2003	Year ended December 31, 2004
Direct project costs: Clinical trials Contract research and development, consulting, materials and other costs Salaries, benefits and related costs Stock-based compensation	\$	\$ 916,000 3,876,000 1,155,000 2,000
Total direct costs Indirect project costs	2,010,000	5,949,000 1,494,000
Total	\$ 2,010,000	\$ 7,443,000

Direct costs increased approximately \$5.9 million from zero as a result in the shift from contract development activities to the clinical development of iloperidone and VEC-162. Salaries, benefits and related costs increased approximately \$1.2 million in 2004 due to an increase in personnel to support the development and clinical trial activities for iloperidone and VEC-162. Personnel costs associated with contract development activities were charged to indirect project costs for the period from March 13, 2003 (inception) to December 31, 2003.

Contract research and development, consulting, materials and other direct costs increased approximately \$3.9 million primarily due to clinical manufacturing-related development costs incurred in connection with the manufacturing of clinical supply materials for iloperidone and VEC-162. Prior to FDA approval of our products, manufacturing-related costs are included in research and development expense. Clinical trials expense increased approximately \$916,000 due to the cost incurred for the VEC-162 Phase II clinical trial.

Indirect project costs decreased by approximately \$516,000, due primarily to the elimination of contract manufacturing activities we previously conducted.

*General and administrative expenses.* General and administrative expenses increased approximately \$1.0 million, or 101%, to approximately \$2.1 million for the year ended December 31,

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2004 compared to approximately \$1.1 million for the period from March 13, 2003 (inception) to December 31, 2003.

The following table discloses the components of our general and administrative expenses:

General and administrative expenses	Period from March 13, 2003 (inception) to Year end December 31, December 3 2003				
Salaries, benefits and related costs	\$ 21,000	\$ 906,000			
Stock-based compensation		36,000			
Legal and consulting expenses	620,000	690,000			
Other expenses	412,000	487,000			
Total	\$ 1,053,000	\$ 2,119,000			

General and administrative expenses consist of professional fees, salaries and related costs for executive and other administrative personnel, and facility costs. Salaries, benefits and related costs increased by approximately \$885,000 in 2004 due to an increase in personnel as we continued to develop the administrative structure to support the development and clinical trial activities of our product candidates.

Legal and consulting expenses increased by approximately \$70,000 due primarily to a higher level of consulting activity in 2004 in support of the business development and market research activities related to our lead product candidates.

Interest and other income, net. Net interest income for the year ended December 31, 2004 was approximately \$59,000 compared to net interest income of approximately \$45,000 for the period from March 13, 2003 (inception) to December 31, 2003. The increase in interest income was attributable to higher average cash balances for the year ended December 31, 2004, and partially offset by an increase in interest expense attributable to an increase in our equipment term loan obligations.

Our interest income and expenses for 2004 and for the period from March 13, 2003 (inception) to December 31, 2003 are as follows:

	Period from March 13, 2003 (inception) to Year en December 31, December 2003	
Interest income	\$ 53,000	\$ 101,000

Interest expense (8,000) (42,000)

Total, net \$ 45,000 \$ 59,000

# Liquidity and capital resources

We have funded our operations through September 30, 2006 principally with the net proceeds from private preferred stock offerings and our initial public offering, totaling approximately \$62.0 million and \$53.3 million, respectively.

At September 30, 2006, our total cash and cash equivalents, short-term investments and restricted cash were approximately \$43.4 million, compared to approximately \$31.6 million at December 31, 2005. Our cash and cash equivalents are highly liquid investments with a maturity of 90 days or less at date of purchase and consist of time deposits, investments in money market funds with commercial banks and financial institutions, and commercial paper of high-quality corporate issuers.

As of December 31, 2004 and 2005 and September 30, 2006 our liquidity resources are summarized as follows:

	As	As of September 30,	
	2004	2005	2006
Balance sheet data Cash and cash equivalents	\$ 16,260,000	\$ 21,013,000	\$ 31,900,000
U.S. government agencies securities U.S. corporate debt securities		6,055,000 4,086,000	6,765,000 4,332,000
Short-term investments Restricted cash		10,141,000 430,000	11,097,000 430,000
	\$ 16,260,000	\$ 31,584,000	\$ 43,427,000

As of September 30, 2006, we maintained all of our cash and cash equivalents in four financial institutions. Deposits held with these institutions may exceed the amount of insurance provided on such deposits, but do not anticipate any losses with respect to such deposits.

Our activities will necessitate significant uses of working capital throughout 2007 and beyond. We plan to continue financing our operations for the foreseeable future with cash received from financing activities. We believe that our current capital resources, together with the net proceeds from this offering, will be sufficient to meet our operating needs into early 2008, and after that time we will require additional capital.

In budgeting for our activities, we have relied on a number of assumptions, including assumptions that:

we will file an NDA for iloperidone in schizophrenia with the FDA by the end of 2007

we will continue to expend funds in preparation of the commercial launch of iloperidone

we will expend funds on the extended-release injectable formulation of iloperidone

we will initiate at least one additional VEC-162 Phase III trial for chronic sleep disorders in the second half of 2007 and that this trial will be conducted in accordance with our expectations

we will initiate our VSF-173 Phase II trial for excessive sleepiness in mid-2007 and that this trial will be conducted in accordance with our expectations

we will not engage in further in-licensing activities

we will not receive any proceeds from potential partnerships

we will not expend funds on the bipolar indication for iloperidone or on a Phase II trial of VEC-162 for depression

we will continue to evaluate pre-clinical compounds for potential development

we will be able to continue the manufacturing of our product candidates at commercially reasonable prices

we will be able to retain key personnel

we will not incur any significant contingent liabilities

We may need to raise additional funds more quickly if one or more of our assumptions proves to be incorrect, if we choose to expand our product development efforts more rapidly than presently anticipated or if we seek to acquire additional product candidates. We may decide to raise additional funds even before they are needed if the conditions for raising capital are favorable. However, we may not be able to raise additional funds on acceptable terms, or at all. If we are unable to secure sufficient capital to fund our research and development activities, we may not be able to continue operations, or we may have to enter into collaboration agreements that could require us to share commercial rights to our products to a greater extent or at earlier stages in the drug development process than is currently intended. These collaborations, if consummated prior to proof-of-efficacy or safety of a given product candidate, could impair our ability to realize value from that product candidate. In the absence of the ability to raise additional equity capital, we are also prepared and have the ability to curtail our existing clinical development commitments and extend them in such a manner so that we have operating funds through mid-2007.

In 2003, we entered into a \$515,147 credit facility to finance the purchase of specified equipment based on lender-approved schedules. The interest rate was fixed at 9.3% per annum. In September 2006 we settled this obligation in full. The total indebtedness relating to this credit facility was approximately \$142,000 as of December 31, 2005.

### Cash flow

The following table summarizes our cash flows for the period from March 13, 2003 (inception) to December 31, 2003, the years ended December 31, 2004 and 2005, and the nine months ended September 30, 2005 and 2006.

	March 13, 2003 (inception) to December 31, 2003	Year ended December 31, 2004	Year ended December 31, 2005	Nine months ended September 30, 2005	Nine months ended September 30, 2006
Net cash (used in) provided by					
Operating activities	\$ (2,108,000)	\$ (8,615,000)	\$ (17,714,000)	\$ (11,813,000)	\$ (40,503,000)
Investing activities	(1,162,000)	(415,000)	(10,818,000)	(511,000)	(1,843,000)
Financing activities	10,438,000	18,146,000	33,294,000	18,335,000	53,237,000
Effect of foreign					
currency translation	(2,000)	(22,000)	(9,000)	(6,000)	(4,000)
Net increase in cash	\$ 7166,000	¢ 0.004.000	\$ 4,753,000	\$ 6.005.000	¢ 10 007 000
and cash equivalents	\$ 7,166,000	\$ 9,094,000	\$ 4,753,000	\$ 6,005,000	\$ 10,887,000

## Nine months ended September 30, 2006 compared to nine months ended September 30, 2005

Net cash used in operations was approximately \$40.5 million and approximately \$11.8 million for the nine months ended September 30, 2006 and 2005, respectively. The net loss for the nine months ended September 30, 2006 of approximately \$51.6 million was offset primarily by non-cash charges for depreciation and amortization of approximately \$415,000, stock-based compensation of approximately \$4.5 million, an increase of accrued expenses of approximately \$5.3 million, principally related to clinical trial expenses, and other net changes in working capital. Net cash used in investing activities for the nine months ended September 30, 2006 was approximately \$1.8 million and consisted primarily of net purchases of short-term investments of

approximately \$0.6 million and purchases of property and equipment of approximately \$1.2 million. Net cash provided by financing activities for the nine months ended September 30, 2006 was approximately \$53.2 million, consisting primarily of net proceeds from the initial public offering of our common stock of \$53.3 million.

## Year ended December 31, 2005 compared to year ended December 31, 2004

Net cash used in operations was approximately \$17.7 million and approximately \$8.6 million for the years ended December 31, 2005 and 2004, respectively. The net loss for the year ended December 31, 2005 of approximately \$23.9 million was offset primarily by non-cash charges for depreciation and amortization of approximately \$424,000, stock-based compensation of approximately \$5.1 million, an increase in accrued expenses and accounts payable of approximately \$1.9 million and \$1.5 million, respectively, principally related to clinical trial expenses, and other net changes in working capital. Net cash used in investing activities for the year ended December 31, 2005 was approximately \$10.8 million and consisted primarily of net purchases of short-term investments of approximately \$10.1 million, equipment purchases of approximately \$292,000 and an investment of approximately \$430,000 in restricted cash for a security deposit on our new leased corporate research and development facility. Net cash provided by financing activities for the year ended December 31, 2005 was approximately \$33.3 million, consisting primarily of net proceeds from the issuance of Series B Preferred Stock of approximately \$33.5 million, offset primarily by payments of equipment debt financing obligations of approximately \$173,000.

## Year ended December 31, 2004 compared to period from March 13, 2003 (inception) to December 31, 2003

Net cash used in operations was approximately \$8.6 million and approximately \$2.1 million for the year ended December 31, 2004 and the period from March 13, 2003 (inception) to December 31, 2003, respectively. The net loss for 2004 of approximately \$9.5 million was partially offset by non-cash charges for depreciation and amortization of approximately \$377,000, an increase in accrued expenses of approximately \$416,000 and other net changes in working capital. Net cash used from investing activities for the year ended December 31, 2004 was approximately \$415,000 and consisted primarily of equipment purchases. Net cash from financing activities for 2004 was approximately \$18.1 million, which consists primarily of net proceeds from the issuance of Series B Preferred Stock of approximately \$18.3 million, offset by principal payments on notes payable and capital lease obligations of approximately \$200,000.

### **Contractual obligations and commitments**

The following table summarizes our long-term contractual cash obligations as of September 30, 2006:

						Ca	ash pa	yme	ents du	ie by	period
		October	•				- '				-
		to	)								
		December	•								After
(In thousands)	Total	2006	)	2007	2008		2009		2010		2010
Operating leases	\$ 4,848	\$ 127	\$	642	\$ 536	\$	427	\$	440	\$	2,676

Operating leases. Our commitments under operating leases shown above consist of payments relating to our real estate leases for our current and former headquarters located in Rockville, Maryland, expiring in 2016 and 2008, respectively, and for our research facility in Singapore that expired in December 2006. We intend to renew the

We vacated our previous headquarters in January 2006. According to Statement of Accounting Standards No. 146, *Accounting for Costs Associated with Exit or Disposal Activities* (SFAS 146), a liability for costs that will continue to be incurred under a lease for its remaining term without economic benefit to the company shall be recognized and measured when the company ceases using the right conveyed by the lease, reduced by estimated sublease rentals that could be reasonably obtained. In accordance with SFAS 146 we have recorded non-cash charges relating to the abandonment of our former office of approximately \$267,000 during the nine months ended September 30, 2006.

*Credit facility*. In 2003, we entered into a \$515,147 credit facility to finance the purchase of specified equipment based on lender-approved schedules. The facility was paid in full in September 2006.

Clinical research organization contracts and other contracts. We have entered into agreements with clinical research organizations responsible for conducting and monitoring our clinical trials for iloperidone and VEC-162, and have also entered into agreements with clinical supply manufacturing organizations and other outside contractors who will be responsible for additional services supporting our ongoing clinical development processes. These contractual obligations are not reflected in the table above because we may terminate them on no more than 60 days notice without incurring additional charges (other than charges for work completed but not paid for through the effective date of termination and other costs incurred by our contractors in closing out work in progress as of the effective date of termination).

We expect that we will incur approximately \$3.0 million in costs from October 1 to December 31, 2006, and approximately \$2.5 million to \$3.5 million in costs in 2007, for clinical trial services rendered in connection with our iloperidone and VEC-162 Phase III trials, primarily in connection with the analysis of trial data and the preparation of regulatory filings.

License agreements. In February 2004 and June 2004, we entered into separate licensing agreements with Bristol-Myers Squibb and Novartis, respectively, for the exclusive rights to develop and commercialize our three compounds in clinical development. In partial consideration for these rights, we paid a \$500,000 non-refundable fee for each compound. We are obligated to make additional payments under the conditions in the agreements upon the achievement of specified clinical, regulatory and commercial milestones. We met a clinical milestone earlier in 2006 under the VEC-162 agreement with Bristol-Myers Squibb and made an associated milestone payment and recorded an expense of \$1,000,000. We may meet other milestones in 2007 under our license agreements with Novartis for iloperidone and VSF-173, for which we would be obligated to make license payments of up to \$6,000,000. If the products are successfully commercialized we will be required to pay certain royalties based on net sales for each of the licensed products. Please see Note 11 to the condensed consolidated financial statements as of September 30, 2006 included with this prospectus for a more detailed description of these license agreements.

We have not included any contractual obligations relating to our license agreements in the above table, since the amount, timing and likelihood of these payments are unknown and will depend on the successful outcome of future clinical trials, regulatory filings, favorable FDA regulatory approvals and growth in product sales. For a more detailed description of the risks associated with the outcome of such clinical trials, regulatory filings, FDA approvals and product sales, please see the Risk factors section of this prospectus.

### Qualitative and quantitative disclosures about market risk

## Foreign exchange

We currently incur a portion of our operating expenses in Singapore. The reporting currency for our consolidated financial statements is U.S. Dollars. To date, we have determined that operating expenses incurred outside of the United States have not been significant. As a result, we have not been impacted materially by changes in exchange rates and do not expect to be impacted materially for the foreseeable future. However, if operating expenses incurred outside of the United States increase, our results of operations could be adversely impacted by changes in exchange rates. We do not currently hedge foreign currency fluctuations and do not intend to do so for the foreseeable future.

#### **Interest rates**

Our exposure to market risk is currently confined to our cash and cash equivalents, restricted cash and short-term investments that have maturities of less than 12 months. We currently do not hedge interest rate exposure. We have not used derivative financial instruments for speculation or trading purposes. Because of the short-term maturities of our cash and cash equivalents, restricted cash and short-term investments, we do not believe that an increase in market rates would have any significant impact on the realized value of our investments, but may increase the interest expense associated with any long-term debt or long-term lease obligations.

### **Effects of inflation**

Our most liquid assets are cash, restricted cash and cash equivalents. Because of their liquidity, these assets are not directly affected by inflation. We also believe that we have intangible assets in the value of our intellectual property. In accordance with generally accepted accounting principles, we have not capitalized the value of this intellectual property on our balance sheet. Due to the nature of this intellectual property, we believe that these intangible assets are not affected by inflation. Because we intend to retain and continue to use our equipment, furniture and fixtures and leasehold improvements, we believe that the incremental inflation related to replacement costs of such items will not materially affect our operations. However, the rate of inflation affects our expenses, such as those for employee compensation and contract services, which could increase our level of expenses and the rate at which we use our resources.

## Off-balance sheet arrangements

We have no off-balance sheet arrangements, as defined in Item 303(a)(4) of the Securities and Exchange Commission s Regulation S-K.

## **Critical accounting policies**

The preparation of our consolidated financial statements requires us 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 our financial statements as well as the reported revenues and expenses during the reported periods. We base our estimates on historical experience and on various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying value of assets and liabilities that are not apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

Our significant accounting policies are described in the notes to our audited consolidated financial statements for the year ended December 31, 2005 included in this prospectus. However,

we believe that the following accounting policies are important to understanding and evaluating our reported financial results, and we have accordingly included them in this discussion.

Accrued expenses. As part of the process of preparing financial statements we are required to estimate accrued expenses. This process involves identifying services that have been performed on our behalf and estimating the level of service performed and the associated cost incurred for such service as of each balance sheet date in our financial statements. Examples of estimated accrued expenses include professional service fees, such as lawyers and accountants, and contract service fees, such as amounts paid to clinical monitors, data management organizations and investigators in conjunction with clinical trials, and fees paid to contract manufacturers in conjunction with the production of clinical materials. In connection with such service fees, our estimates are most affected by our understanding of the status and timing of services provided relative to the actual levels of services incurred by such service providers. The majority of our service providers invoice us monthly in arrears for services performed. In the event that we do not identify certain costs that have begun to be incurred or we under- or over-estimate the level of services performed or the costs of such services, our reported expenses for such period would be too low or too high. The date on which certain services commence, the level of services performed on or before a given date and the cost of such services are often subject to our judgment. We make these judgments based upon the facts and circumstances known to us in accordance with generally accepted accounting principles.

Stock-based compensation. On January 1, 2006, we began accounting for stock-based compensation under the provisions of Statement of Financial Accounting Standards (SFAS) No. 123(R), which requires the recognition of the fair value of stock-based compensation. Under the fair value recognition provisions of SFAS No. 123(R), stock-based compensation cost is estimated at the grant date based on the fair value of the awards expected to vest and recognized as expense ratably over the requisite service period of the award. We adopted SFAS No. 123(R) using the modified prospective method of implementation, which requires the application of the accounting standard with respect to all periods beginning after January 1, 2006. Our condensed consolidated financial statements as of and for the nine months ended September 30, 2006 reflect the impact of SFAS No. 123(R). In accordance with the modified prospective method, the consolidated financial statements for all periods prior to January 1, 2006 have not been restated to reflect, and do not include, the impact of SFAS No. 123(R).

Prior to January 1, 2006, we elected to follow APB Opinion No. 25, *Accounting for Stock Issued to Employees*, and related interpretations, in accounting for our stock-based compensation plans, rather than the alternative fair value accounting method provided for under SFAS No. 123, *Accounting for Stock-Based Compensation*. In the notes to our financial statements for periods ending prior to January 1, 2006, we have provided pro forma disclosures in accordance with SFAS No. 123 and related pronouncements. We accounted for transactions in which services are received in exchange for equity instruments based on the fair value of such services received from non-employees or of the equity instruments issued, whichever is more reliably measured, in accordance with SFAS No. 123 and EITF Issue No. 96-18, *Accounting for Equity Instruments that Are Issued to Other than Employees for Acquiring, or in Conjunction with Selling, Goods or Services*. The two factors which most affected charges or credits to operations related to stock-based compensation were the fair value of the common stock underlying stock options for which stock-based compensation is recorded and the volatility of such fair value. If our estimates of the fair value of these equity instruments are too high or too low, it would have had the effect of overstating or understating expenses.

Given the lack of an active public market prior to our initial public offering on April 12, 2006, our board of directors determined the fair value of our common stock for stock option awards.

The Company did not obtain any contemporaneous valuations of its common stock by an unrelated valuation specialist during the year 2004 and through November 2005 because the Company did not then have a reasonable expectation of conducting an initial public offering, and engaging an outside valuation firm to perform a valuation of the Company at the time of each option grant was not practical. Instead, we relied on our board of directors to determine fair value.

When discussions were initiated with the underwriters of our initial public offering in November 2005, our board of directors and management believed that the underwriters could provide us with additional perspective and points of reference which we could factor into our determination of the fair value of our common stock. We then engaged in retrospective valuations of our common stock for the years ended December 31, 2004 and December 31, 2005. In establishing our estimates of fair value in this retrospective analysis, we considered the guidance set forth in the AICPA Practice Guide, Valuation of Privately-Held-Company Equity Securities Issued as Compensation ( AICPA Practice Guide ). Our determinations of fair value were based on an approved valuation method under the AICPA Practice Guide the income method. We determined that this was an appropriate method to use based on the Company s development stage at the time the retrospective valuations were completed. The income method involves applying appropriate discount rates to estimated cash flows that are based on forecasts of revenue and costs. Our revenue forecasts and related cost of sales were based on information obtained from a third-party research consultant. Our revenue forecasts were based on expected annual growth rates ranging from approximately 50 percent following the first full year of commercial launch to approximately 7 percent beginning five years following commercial launch for our product candidates. Operating expenses were based on our own assumptions and estimates for growth, which were consistent with the information also obtained from our independent research consultant. We assumed that operating expenses would continue to increase through the development and commercialization of our product candidates and that the company would begin receiving revenues in 2009. There is inherent uncertainty in these estimates and the assumptions underlying our estimates, but the estimates that were used were consistent with our business plan. The forecast information used for our iloperidone and VEC-162 financial projections was evaluated, and these projections were discounted by 90% and 70%, respectively, in order to account for the uncertainties related to the future commercial launch of the products. In addition, the risks associated with achieving our forecasts were assessed when selecting the appropriate discount rates for the related discounted cash flow analysis, which ranged from 12% to 15%. The overall enterprise value of the Company was then allocated to the shares of preferred stock and common stock on a fully-diluted basis given the conversion of preferred stock to common stock upon the completion of our initial public offering. As set forth in the table below, we granted stock options with exercise prices ranging from \$0.33 to \$4.73 during the two years ended December 31, 2005.

Also as set forth in the table below, we retrospectively determined that the fair value of our common stock increased from \$3.21 to \$17.18 per share during that period. Based on the \$17.18 value per share (fully-diluted basis), we retrospectively assessed the fair value of common stock for each date during these two years on which stock options were granted. In assessing the value of the common stock at each grant date, management considered the factors listed above, including the achievement of success for the following key drivers: license agreements, clinical trials, strong management and infrastructure.

*License agreements:* Given the importance of our current license agreements to develop our iloperidone and VEC-162 compounds into drugs for commercial sale, the value for each license

agreement increased from the period the agreements were first entered through the end of 2005.

*Clinical trials:* We believe that our success in our clinical development programs for iloperidone and VEC-162 has created additional value. Our clinical trial development programs resulted in the increase in value of the Company for the period beginning June 2004 through the end of 2005.

Strong management and infrastructure: The collection of a team of expert scientists and the Chief Executive Officer, along with other key personnel, provided an increase in value to the Company at each hire date, beginning at the inception of the Company through the end of 2005.

As a result of assessing these drivers based on their importance to creating value for the Company, we determined that the fair value of our common stock on a fully-diluted basis steadily increased from \$3.21 per share at March 31, 2004 to \$17.18 per share at December 31, 2005. The reasons for the difference between the range of \$0.33 to \$4.73 per share and an estimated fair value of \$17.18 per share were as follows:

During the quarter ending June 30, 2004, the Company in-licensed its first product candidate, VEC-162 and formally commenced a Phase II clinical development program in insomnia.

During the quarter ending September 30, 2004, the Company in-licensed two additional product candidates; iloperidone for the treatment of schizophrenia and bipolar disorder, and VSF-173 for the treatment of excessive sleepiness. The Company also initiated a clinical development program for iloperidone in preparation for a Phase III clinical trial in schizophrenia. In addition, the Company completed its first closing of Series B Preferred Stock for \$18.5 million and added key executive management personnel.

During the quarter ending December 31, 2004, the Company conducted an initial guidance meeting with the FDA regarding its planned clinical trial for VEC-162 in transient insomnia. The Company also further defined its pharmacogenetic strategy for a future Phase III iloperidone clinical trial in schizophrenia.

During the quarter ending March 31, 2005, the Company developed additional insight regarding the previous clinical trials conducted by the licensor for its iloperidone product candidate. This review will result in improvements to the design and execution of the future Phase III iloperidone clinical trial in schizophrenia. In addition, the Company added key scientific staff and added to its executive management group.

During the quarter ending June 30, 2005, the Company conducted a guidance meeting with the FDA regarding its planned Phase III clinical trial for iloperidone in schizophrenia and the related pharmacogenetic elements of the study. The Company also completed a successful Phase II clinical trial for its VEC-162 product candidate in insomnia.

During the quarter ending September 30, 2005, the Company conducted a Phase II (b) and statistical guidance meeting with the FDA regarding its planned Phase III clinical trail for iloperidone in schizophrenia. In addition, the Company initiated clinical development activities in preparation for a Phase III clinical trial for VEC-162 in insomnia. The Company also completed the second closing of the Series B Preferred Stock financing for \$18.5 million.

During the quarter ending December 31, 2005, the Company began its Phase III clinical trial for iloperidone in schizophrenia. In addition, the Company added to its executive management group.

In March of 2006, the underwriters subsequently determined that the assumed initial public offering price would be \$10.00 per share. The difference between our prior estimated fair market value of \$17.18 and our initial public offering price was largely a result of the underwriters—view of current market conditions and other factors, including the latest available financial and market data from which our original projections and valuations were derived.

Information on stock option grants, net of forfeitures, during the previous two years ended December 31, 2005 and in the first quarter of 2006 (in which the last of our options were granted prior to our initial public offering on April 12, 2006) is summarized as follows:

				Fair market value	
		Number		estimate	
Date of		of options	Exercise	per common	Intrinsic value
issuance	Type of equity issuance	Granted	Price(1)	share	per share
06/15/04	Employee options	3,443	\$ 0.33	\$ 3.21	\$ 2.88
09/01/04	Employee options	91,668	0.33	4.07	3.74
12/06/04	Employee options	777	0.33	5.69	5.36
02/10/05	Employee options	209,893	0.33	10.52	10.19
04/05/05	Employee options	27,974	0.33	15.99	15.66
08/15/05	Employee options	15,559	0.33	16.85	16.52
09/28/05	Employee options	620,973	0.33	16.85	16.52
10/03/05	Employee options	906	0.33	17.18	16.85
11/14/05	Employee options	83,087	0.83	17.18	16.35
12/29/05	Employee options	358,847	4.73	17.18	12.45
01/26/06	Employee options	17,017	4.73	17.18	12.45
03/16/06	Employee options	17,372	7.45	13.00	6.00

(1) The board of directors approved a modification to all outstanding stock option awards that were granted prior to February 10, 2005, repricing the options from their original exercise price of \$1.32 to \$0.33. According to FIN 44, the result of such a modification is to account for the modified stock option awards as variable from the date of the modification to the date the awards are exercised, forfeited, or cancelled. We remeasured the modified awards that were outstanding at the end of each quarter during the year ended December 31, 2005 and the first quarter ended March 31, 2006.

Equity instruments issued to non-employees. We account for transactions in which services are received in exchange for equity instruments based on the fair value of such services received from non-employees or of the equity instruments issued, whichever is more reliably measured, in accordance with SFAS No. 123 and EITF Issue No. 96-18, Accounting for Equity Instruments that Are Issued to Other than Employees for Acquiring, or in Conjunction with Selling, Goods or Services. On January 19, 2006, the Company issued one of our consultants a grant to purchase 3,625 shares of our common stock with the exercise price of \$4.73, of which 2,190 were fully vested as of January 19, 2006 and the balance will vest ratably over 19 months. The option expires on January 19, 2016 and for the nine months ended September 30, 2006 we recorded a consulting expense of approximately \$36,000 relating to this option.

During the three months ended September 30, 2006 the Company entered into two consulting agreements that will require the Company to grant options to purchase up to 20,000 shares of common stock to these consultants subject to

certain performance criteria. The terms of the stock option grants will be finalized upon their issuance.

*Income taxes*. As part of the process of preparing our financial statements we are required to estimate our income taxes in each of the jurisdictions in which we operate. We account for income taxes by the liability method in accordance with the provisions of SFAS No. 109,

Accounting for Income Taxes. Under this method, deferred income taxes are recognized for tax consequences in future years of differences between the tax bases of assets and liabilities and their financial reporting amounts at each year-end, based on enacted laws and statutory tax rates applicable to the periods in which the differences are expected to affect taxable income. Valuation allowances are provided if, based upon the weight of available evidence, it is more likely than not that some of all of the deferred tax assets will not be realized. We have not recorded any tax provision or benefit for any period since our inception. We have provided a valuation allowance for the full amount of our net deferred tax assets since realization of any future benefit from deductible temporary differences and net operating loss carry-forwards cannot be sufficiently assured. As of December 31, 2004 and 2005, we had U.S. federal and state net operating loss carryforwards of approximately \$10.0 million and \$21.6 million, respectively, that will begin to expire in 2023.

New Accounting Standards. In July 2006, the Financial Accounting Standard Board (FASB) issued FASB Interpretation No. 48 (FIN 48) Accounting for Uncertainty in Income Taxes an interpretation of FASB Statement No. 109, to clarify certain aspects of accounting for uncertain tax positions, including issues related to the recognition and measurement of these tax positions. This interpretation is effective for fiscal years beginning after December 15, 2006. While we are currently evaluating FIN 48, this pronouncement is not currently expected to have significant impact on our results of operations and financial condition.

In September 2006, the FASB issued FASB Statement No. 157, *Fair Value Measurements* (FAS 157), which addresses how companies should measure fair value when they are required to use a fair value measure for recognition or disclosure purposes under generally accepted accounting principles (GAAP). FAS 157 outlines a common definition of fair value to be used throughout GAAP and the new standard intends to make the measurement of fair value more consistent and comparable and improve disclosures about those measures. Companies will need to adopt FAS 157 for financial statements issued for fiscal years beginning after November 15, 2007. We are currently evaluating the impact of FAS 157 on our results of operations and financial condition.

In September 2006, the Staff of the SEC issued Staff Accounting Bulletin No. 108, Considering the Effects of Prior Year Misstatements when Quantifying Misstatements in Current Year Financial Statements (SAB 108). SAB 108 provides guidance on the consideration of the effects of prior year misstatements in quantifying current year misstatements for the purpose of determining whether the current year s financial statements are materially misstated. SAB 108 will be effective for the Company in the fourth quarter of 2006. We are currently evaluating the requirements of SAB 108; however, we do not believe that its adoption will have a material effect on our financial statements.

### **Business**

### Overview

We are a biopharmaceutical company focused on the development and commercialization of clinical-stage drug candidates, with exclusive worldwide commercial rights to three product candidates in clinical development for various central nervous system disorders. Our lead product candidate, iloperidone, is a compound for the treatment of schizophrenia and bipolar disorder. In December 2006 we announced positive top-line results from our Phase III trial of iloperidone in schizophrenia. Iloperidone appeared to be safe and well-tolerated in the trial, and demonstrated statistically significant improvement in efficacy versus placebo on the Positive and Negative Symptoms Scale (PANSS), the trial s primary endpoint, as well as statistically significant improvements in other measures of efficacy. Our second product candidate, VEC-162, is a compound for the treatment of sleep and mood disorders. In November 2006 we announced positive top-line results from our Phase III trial of VEC-162 in transient insomnia. VEC-162 demonstrated statistically significant improvement in several parameters used to measure the efficacy of insomnia therapies, including reduced duration of wake after sleep onset, improved sleep efficiency and shortened time to persistent sleep. In addition, VEC-162 was found to be safe and well-tolerated. VEC-162 is also ready for Phase II trials for the treatment of depression. Our third product candidate, VSF-173, is a compound for the treatment of excessive sleepiness and is ready for a Phase II trial. Each of these product candidates benefits from strong new chemical entity patent protection and may offer substantial advantages over currently approved therapies.

We expect to file a New Drug Application (NDA) for iloperidone in schizophrenia with the United States Food and Drug Administration (FDA) by the end of 2007. We expect to meet with the FDA in the first quarter of 2007 regarding this filing. We will have to conduct additional Phase III trials for VEC-162 in chronic sleep disorders prior to our filing of an NDA for VEC-162, and we expect to begin at least one such additional trial in the second half of 2007. We also expect to begin a Phase II trial of VSF-173 for excessive sleepiness in mid 2007. Assuming successful outcomes of our clinical trials and approval by the FDA, we expect to commercialize iloperidone and VSF-173 with our own sales force in the U.S., and expect to commercialize VEC-162 through a partnership with a global pharmaceutical company, although we have not yet identified such a global partner.

Our three product candidates target large prescription markets with significant unmet medical needs. Sales of antipsychotic drugs were approximately \$16 billion in 2005, according to *World Review Analyst* by IMS, a leading pharmaceutical market research company. These sales were achieved despite the safety concerns, moderate efficacy and poor patient compliance that are associated with these drugs. We believe that iloperidone may address some of these shortcomings, based on its observed safety profile and based on further improvements to iloperidone that we plan to develop. According to IMS, in 2005 the insomnia market generated approximately \$4.5 billion in worldwide sales and the depression market accounted for worldwide sales in excess of \$19 billion. However, the approved drugs in both the sleep and mood disorders markets have sub-optimal safety and efficacy profiles. We believe VEC-162 may represent a breakthrough in each of these markets, based on the product s efficacy, safety and novel mechanism of action. The excessive sleepiness market generated approximately \$500 million in worldwide sales in 2005. Few drugs exist to treat this condition, and each of the available drugs has limitations. We believe that VSF-173 may represent a safe and effective alternative treatment in this growing market.

Our team includes experienced pharmaceutical industry executives, and our scientific team possesses deep expertise in clinical development and in pharmacogenetics and

pharmacogenomics, the scientific disciplines that examine both genetic variations among people that influence response to a particular drug and the multiple pathways through which drugs affect people. Our founder and Chief Executive Officer, Mihael H. Polymeropoulos, M.D., started our operations in early 2003 after establishing and leading the Pharmacogenetics Department at Novartis.

We believe that the combination of our clinical development expertise and our pharmacogenetics and pharmacogenomics expertise will enable us to shorten our drug development timeline relative to traditional approaches of drug discovery and development, and to provide additional differentiation for our product candidates. We also believe that our expertise will provide us with preferential access to compounds discovered by other pharmaceutical companies. This expertise allowed us to acquire the exclusive worldwide commercial rights to iloperidone and VSF-173 from Novartis and also allowed us to obtain exclusive worldwide commercial rights to VEC-162, which had originally been developed by Bristol-Myers Squibb Company (BMS).

## **Our strategy**

Our goal is to create a leading biopharmaceutical company focused on developing and commercializing products that address critical unmet medical needs through the application of our drug development expertise and our pharmacogenetics and pharmacogenemics expertise. The key elements of our strategy to accomplish this goal are to:

Pursue the clinical development and regulatory approval of our current product candidates. We have announced positive top-line results for our recently completed Phase III iloperidone trial in schizophrenia, and we believe that this trial will be the last trial required before filing an NDA for iloperidone. We recently completed a Phase III trial for VEC-162 in transient insomnia, for which we announced positive top-line results in November 2006. We believe that we will need to conduct additional Phase III trials of VEC-162 in chronic sleep disorders prior to filing an NDA for this compound. We intend to initiate a Phase II trial for VSF-173 in mid-2007. We have committed, and will continue to commit, substantial resources towards completing the development of, and obtaining regulatory approvals for, our product candidates.

Develop a focused commercialization capability in the United States. Because we believe that the number of physicians accounting for the majority of prescriptions in the United States for schizophrenia and excessive sleepiness is relatively small, we believe that we can cost-effectively develop our own sales force to market and sell iloperidone and VSF-173.

Enter into partnerships to extend our commercial reach. Given the large number of physicians treating sleep and mood disorders, we intend to enter into a global partnership with a large pharmaceutical company to market, distribute and sell VEC-162. Additionally, we intend to seek commercial partners for iloperidone and VSF-173 outside of the United States.

Apply our pharmacogenetics and pharmacogenomics expertise to differentiate our products. We believe that our pharmacogenetics and pharmacogenomics expertise will yield new insights into our product candidates. These insights may enable us to target our products to certain patient populations and to identify unexpected conditions for our product candidates to treat. We believe this expertise will enable us to differentiate and extend the lifecycle of each of our product candidates. Our expertise may allow us to develop companion diagnostic tests to help physicians identify patient populations that will realize greater benefits from our compounds.

Expand our product portfolio through the identification and acquisition of additional compounds. We intend to continue to draw upon our clinical development expertise and

pharmacogenetics and pharmacogenomics expertise to identify and pursue additional clinical-stage compounds.

## **Development programs**

We have the following product candidates in clinical trials:

Product candidate	Target indications	Clinical status
Iloperidone (Oral)	Schizophrenia	Phase III trial completed; NDA expected to be filed by the end of 2007
Iloperidone (Depot)	Bipolar Disorder Schizophrenia	Ready for Phase III trial Ready for Phase II trial
VEC-162	Insomnia	Phase III trial completed; additional Phase III trials to be conducted
VCE 172	Depression  Function Standings	Ready for Phase II trial
VSF-173	Excessive Sleepiness	Ready for Phase II trial

## **Iloperidone**

We are developing iloperidone, a compound for the treatment of schizophrenia and bipolar disorder. In December 2006 we announced positive top-line results from our Phase III trial of iloperidone in schizophrenia, which completed its enrollment in August 2006. The drug appeared to be safe and well-tolerated in the trial, and demonstrated statistically significant improvement in efficacy versus placebo on the Positive and Negative Symptoms Scale (PANSS), the trial s primary endpoint, as well as statistically significant improvements in other measures of efficacy. Based on our End of Phase IIb meeting with the FDA in September 2005, we believe we will be able to file an NDA for iloperidone for schizophrenia by the end of 2007. We expect to meet with the FDA in the first quarter of 2007 regarding this filing. If iloperidone obtains regulatory approval, we believe it will represent a unique new therapy for schizophrenia with distinct advantages over currently available therapies.

# Therapeutic opportunity

Schizophrenia is a chronic, debilitating mental disorder characterized by hallucinations, delusions, racing thoughts and other psychotic symptoms (collectively referred to as positive symptoms), as well as moodiness, anhedonia (inability to feel pleasure), loss of interest, eating disturbances and withdrawal (collectively referred to as negative symptoms), and additionally attention and memory deficits (collectively referred to as cognitive symptoms). Schizophrenia develops in late adolescence or early adulthood in approximately 1% of the world s population. Genetic and environmental factors are believed to be responsible for the disease. Most schizophrenia patients today are treated with drugs known as atypical antipsychotics, which were first approved in the U.S. in the late 1980s. These antipsychotics have been named atypical for their ability to treat a broader range of negative symptoms than the

first-generation typical antipsychotics, which were introduced in the 1950s and are now generic. Atypical antipsychotics are generally regarded as having improved side effect profiles and efficacy relative to typical antipsychotics and currently comprise 90% of schizophrenia prescriptions. The global market for atypical antipsychotics was in excess of \$12 billion in 2005. Currently

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approved atypical antipsychotics include olanzapine (Zyprexa<sup>®</sup>, Eli Lilly and Company), risperidone (Risperdal<sup>®</sup>, Johnson & Johnson), quetiapine (Seroquel<sup>®</sup>, AstraZeneca), aripiprazole (Abilify<sup>®</sup>, BMS), ziprasidone (Geodon<sup>®</sup>, Pfizer), paliperidone (Invega<sup>®</sup>, Johnson & Johnson) and generic clozapine.

## Limitations of current treatments

The treatment of schizophrenia remains challenging because currently approved antipsychotics, even atypical antipsychotics, often induce serious side effects and offer only modest and occasional efficacy. Side effects include weight gain, diabetes, extrapyramidal symptoms (involuntary bodily movements), hyperprolactinemia (an elevated secretion of the hormone prolactin which can lead to sexual dysfunction and breast development and milk secretion in women and men), increased somnolence (sleepiness) and cognition difficulties. The side effect profile and modest efficacy of currently available antipsychotics result in poor patient compliance to their prescribed drug regimen. Consequently, there remains a high degree of dissatisfaction with atypical antipsychotics among physicians and patients. Research by LEK Consulting LLC, a leading consulting firm, supports this, showing that physicians employ a trial-and-error approach of prescribing a series of different atypical antipsychotics as they attempt to balance side effects and symptom management in each patient. In addition, the recent CATIE (Clinical Antipsychotic Trials of Interventional Effectiveness) study, conducted by the National Institute of Mental Health and reported in *The New England Journal of Medicine*, found that 74% of patients taking antipsychotics discontinued treatment within 18 months. The average time to discontinuation for these patients in the CATIE study was approximately 6 months.

### Potential advantages of iloperidone

In addition to the efficacy observed in clinical trials to date, our experience with iloperidone thus far suggests that the compound may provide benefits to patients beyond those provided by currently available drugs:

Safety. Our Phase III trial and other short- and long-term safety trials have shown that patients who used iloperidone had reduced side effects relative to currently available antipsychotics, including low weight gain, no induction of diabetes, low extrapyramidal symptoms, including no akathisia (inability to sit still), no hyperprolactinemia, low incidence of sleepiness and low negative effects on cognition relative to placebo. Like other atypical antipsychotics, iloperidone is associated with a prolongation of the heart s QTc interval, but in no instance did any patient taking iloperidone in the controlled portion of a clinical trial have an interval exceeding a 500-millisecond threshold that the FDA has identified as being of particular concern. Two patients experienced a prolongation of 500 milliseconds or more during the open-label extension of one trial. We believe that the safety profile of iloperidone may result in improved patient compliance with their treatment regimen.

Extended-release injectable formulation. We are developing an extended-release injectable formulation for iloperidone, which only needs to be administered once every four weeks and which we believe will be a compelling complement to our oral formulation for both physicians and patients. Novartis conducted a two-month Phase I/IIa safety trial of this formulation in schizophrenia patients, in which it demonstrated the benefit of consistent release over a four-week time period with no greater side effects relative to oral dosing. The commercial potential for our extended-release injectable formulation has been demonstrated by the success of the injectable formulation for risperidone, Risperdal<sup>®</sup> Consta<sup>®</sup>, which achieved worldwide sales of in excess of \$550 million in 2005. We believe that our four-week formulation for iloperidone will be an attractive alternative to Risperdal Consta, which is

injected once every two weeks. Additionally, and unlike Risperdal<sup>®</sup> Consta<sup>®</sup>, we do not believe that the injectable formulation of iloperidone will require oral titration, which would result in simplified dosing.

Additionally, we plan to continue to apply our pharmacogenetics and pharmacogenomics expertise to develop tools that may allow physicians to avoid the trial-and-error approach to prescribing antipsychotic medications for their patients:

Pharmacogenetic evaluation of iloperidone s efficacy. Based on the results of our recently completed Phase III trial, as well as analyses of prior clinical data for iloperidone, we have determined that certain patients may be more likely to respond to iloperidone and to enjoy better treatment results relative to the general schizophrenia patient population. These patients have a common mutation of a gene, linked to central nervous system function, that is estimated to occur in approximately 70% of schizophrenia patients. We developed a genetic test which we used in our recently completed Phase III trial and confirmed this correlation. According to market research conducted by LEK Consulting, physicians treating schizophrenia patients would enthusiastically welcome a genetic test that would enable them to identify likely responders to iloperidone, given the unpredictable efficacy and serious side effects currently associated with atypical antipsychotics, and be more likely to prescribe iloperidone as a result.

Pharmacogenetic evaluation of iloperidone s safety. Based on the results of our recently completed Phase III trial, and other pharmacogenetic analysis, we have discovered that patients with an uncommon mutation of a well understood gene affecting drug metabolism experience higher levels of iloperidone in their blood and may experience longer QTc intervals while taking iloperidone. We estimate that this genetic attribute is found in approximately 25-30% of schizophrenia patients, comprised of poor metabolizers (approximately 5-10% of schizophrenia patients) and intermediate metabolizers (approximately 20% of schizophrenia patients). We believe that certain physicians may choose to test patients for this mutation if they have a concern about QTc interval prolongation with respect to a particular patient.

We intend to make one simple blood test for both markers available through national reference laboratories.

### Overview of our Phase III trial

In November 2005, we initiated our Phase III trial to evaluate iloperidone for the treatment of patients with schizophrenia. We completed enrollment for the trial in August 2006. The trial was a randomized, double-blind, placebo- and active-controlled Phase III trial of 604 patients with schizophrenia. Patients received four weeks of inpatient treatment in the trial. The iloperidone formulation being used in the study is an oral, twice-daily dose of 12 mg, or 24 mg per day. The trial was conducted in the United States and India by Quintiles Transnational, a contract research organization.

In December 2006, we reported positive top-line results for multiple endpoints of the trial using Mixed Method Repeated Measures (MMRM) statistical analysis. Specifically, iloperidone achieved statistically significant efficacy versus placebo in:

PANSS over the entire patient population (p = 0.006)

the positive symptoms subscale of PANSS (p = 0.0009)

the negative symptoms subscale of PANSS (p = 0.027)

an additional rating system of psychiatric symptoms called the Brief Psychiatric Rating Scale (p = 0.0128)

Iloperidone also achieved statistically significant efficacy in PANSS in the trial under a last observation carried forward (LOCF) statistical analysis. Our results confirm the conclusions we reached with respect to our retrospective analysis of three earlier Phase III clinical trials of iloperidone conducted by Novartis, in which iloperidone achieved statistical significance versus placebo for at least one dose in each Phase III trial. For regulatory purposes, only one of these three Phase III trials achieved success by demonstrating statistical significance at the dose that was the primary endpoint of the trial. The data for the three Phase III trials conducted by Novartis (ILP 3000, ILP 3004 and ILP 3005) and our Phase III trial (ILP 3101) are summarized in the following table:

			Positive and negative	
Trial number	Number of patients	Doses(1)	symptom scale improvement(2)	Significance vs. placebo(3)
ILP 3000	621	placebo 4 mg/day 8 mg/day(4) 12 mg/day(4)	-4.6 -9.0 -7.8 -9.9	n/a Not significant Not significant $p = 0.047$
ILP 3004	616	placebo 4-8 mg/day <b>10-16 mg/day</b>	-3.5 -9.5 <b>-11.1</b>	p = 0.017 $p = 0.002$
ILP 3005	710	placebo <b>12-16 mg/day</b> 20-24 mg/day	-7.6 <b>-11.0</b> -14.0	n/a Not significant $p = 0.005$
ILP 3101	604	placebo <b>24 mg/day</b>	-7.1 -12.0	p = 0.006

- (1) Declared dose (the dose for which a drug must show statistically significant improvement vs. placebo) is italicized and bolded.
- (2) As patients improve, their Positive and Negative Symptom Scale score decreases.
- (3) This is represented by p value, which measures likelihood that a difference between drug and placebo is due to random chance. A p<0.05 means the chance that the difference is due to random chance is less than 5%, and is a commonly accepted threshold for denoting a meaningful difference between drug and placebo.
- (4) Declared dose in this trial was a composite of 8 and 12 mg/day.

Vanda also evaluated iloperidone s efficacy and safety in patients with the common genetic mutation linked to central nervous system function described above, using its PG expertise. We developed a genetic test which we used in our

recently completed Phase III trial and confirmed that patients with this common genetic mutation, observed in approximately 70% of schizophrenia patients, were significantly more likely to respond to iloperidone than those in the general schizophrenia population. Patients with this common mutation achieved an improvement versus placebo in PANSS of 6.37 (p=0.002), compared to -0.09 improvement (p=0.981) in patients without the mutation and 4.93 (p=0.006) in all iloperidone patients.

In addition to our efficacy findings, iloperidone also appeared to be safe and well-tolerated in the trial. Vanda measured the effect of iloperidone on the QTc intervals of participating patients. The mean QTc prolongation at 14 days across participating patients (11.4 milliseconds) was consistent with previous trials of iloperidone. No patient experienced a QTc interval of over 500 milliseconds in the trial. The difference in mean QTc prolongation between patients with the uncommon genetic mutation affecting iloperidone metabolism described above (15.0 milliseconds) and patients without the mutation (10.4 milliseconds) was statistically significant at 14 days (p=0.008). The magnitude of QTc prolongation also diminished over time. At 28 days, the mean prolongation for patients without the common mutation described above was 5.0 milliseconds, while for patients with the mutation the prolongation fell to 12.9 milliseconds. The

difference in mean QTc prolongation between patients with the mutation and patients without the mutation was also statistically significant at 28 days (p = 0.002). The mean QTc prolongation for all participating patients at 28 days was 7.0 milliseconds.

We believe that, as of the conclusion of our Phase III trial, our data and documentation on iloperidone will be adequate to support an FDA filing of oral iloperidone. We conducted an End of Phase IIb meeting with the FDA in September 2005, during which the agency agreed that our trial s design is adequate to measure short-term efficacy in schizophrenia. The FDA also agreed that with success in this trial, the iloperidone package would be sufficient for filing an NDA. We expect to file an NDA for iloperidone by the end of 2007. We expect to meet with the FDA in the first quarter of 2007 regarding this NDA filing.

## Potential indication for bipolar disorder

In addition to schizophrenia, we believe iloperidone may be effective in treating bipolar disorder. All of the approved atypical antipsychotics have received approval for bipolar disorder subsequent to commercializing for the treatment of schizophrenia. Approximately 20% of antipsychotic prescriptions are for the treatment of bipolar disorder, according to LEK Consulting. Iloperidone is ready for an initial Phase III trial in bipolar disorder.

#### **Commercialization**

We expect to build our own sales force to market iloperidone directly to psychiatrists and other target physicians in the U.S. Because the U.S. psychiatric community is relatively small, we believe that we can cost-effectively develop our own sales force to market and sell iloperidone. Outside of the United States, we expect to find commercial partners for iloperidone.

## Intellectual property

Iloperidone and its metabolites, formulations, and uses are covered by a total of nine patent and patent application families worldwide. The primary new chemical entity patent covering iloperidone expires normally in 2011 in the United States and 2010 in most of the major markets in Europe. In the United States, the Hatch-Waxman Act of 1984 provides for an extension of new chemical entity patents for a period of up to five years following the expiration of the patent covering that compound to compensate for time spent in development. We believe that iloperidone will qualify for the full five-year patent term extension. In Europe, similar legislative enactments provide for five-year extensions of new chemical entity patents through the granting of Supplementary Protection Certificates, and we believe that iloperidone will qualify for this extension as well. Consequently, assuming that we are granted all available extensions by the FDA and European regulatory authorities and that we receive regulatory approval, we expect that our rights to commercialize iloperidone will be exclusive until 2016 in the United States and until 2015 in Europe. Additionally, the patent application covering the depot formulation of iloperidone, if it is granted, will expire normally in 2022. Several other patent applications covering uses, formulations and derivatives relating to iloperidone extend beyond 2020. Pursuant to a recent European Union directive, we may also acquire the exclusive right in most European Union countries to market iloperidone for a period of 10 years from the date of its regulatory approval in Europe (with the possibility for a further one-year extension), even though the European patents covering iloperidone will likely expire prior to the end of such 10-year period. No generic versions of iloperidone would be permitted to be marketed or sold during this 10-year period in most European countries. See Patents and Intellectual Property below for a more complete description of our intellectual property rights.

We acquired worldwide, exclusive rights to the new chemical entity patent covering iloperidone and certain related intellectual property from Novartis under a sublicense agreement we entered into in 2004. Please see License agreements below for a more complete description of the rights we acquired from Novartis with respect to iloperidone.

#### **VEC-162**

VEC-162 is an oral compound in development for sleep and mood disorders. The compound selectively binds the brain s melatonin receptors, which are thought to govern the body s natural sleep/wake cycle. Compounds that selectively bind to these receptors are thought to be able to help treat sleep disorders, and additionally are believed to offer potential benefits in mood disorders. We announced positive top-line results from our Phase III trial of VEC-162 in transient insomnia in November 2006. VEC-162 is also ready to commence a Phase II trial for the treatment of depression.

# Therapeutic opportunity

Industry sources estimate that of the 73 million U.S. adults who suffer from some form of insomnia, only approximately 11 million currently receive treatment. Sleep disorders are segmented into three major categories: primary insomnia, secondary insomnia and circadian rhythm sleep disorders. Insomnia is a symptom complex that comprises difficulty falling asleep or staying asleep, or non-refreshing sleep, in combination with daytime dysfunction or distress. The symptom complex can be an independent disorder (primary insomnia) or be a result of another condition such as depression or anxiety (secondary insomnia). Circadian rhythm sleep disorders result from a misalignment of the sleep/wake cycle and an individual s daily activities or lifestyle. The circadian rhythm is the rhythmic output of the human biological clock and is governed primarily by the hormone melatonin. Both the timing of behavioral events (activity, sleep, and social interactions) and the environmental light-dark cycle result in a sleep/wake cycle that follows the circadian rhythm. Examples of circadian rhythm sleep disorders include transient disorders such as jet lag and chronic disorders such as shift work sleep disorder. Market research we have conducted with LEK Consulting indicates that circadian rhythm sleep disorders represent a significant portion of the market for sleep disorders. In 2005, the sleep disorder drug market generated approximately \$4.5 billion in worldwide sales, according to IMS.

There are a number of drugs approved and prescribed for patients with sleep disorders. The most commonly prescribed drugs are hypnotics, such as zolpidem (Ambien®, Sanofi-Aventis), eszopiclone (Lunesta®, Sepracor) and zaleplon (Sonata®, King Pharmaceuticals). These drugs work by acting upon a set of brain receptors known as GABA receptors. Several drugs in development, including indiplon (Neurocrine Biosciences) and gaboxadol (Merck/Lundbeck), also utilize a similar mechanism of action. Members of the benzodiazapine class of sedatives are also approved for insomnia, but their usage has declined due to an inferior safety profile compared to hypnotics. Anecdotal evidence also suggests that sedative antidepressants, such as trazodone and doxepin, are prescribed off-label for insomnia. Recently, the FDA approved ramelteon (Rozerem Takeda), a compound with a mechanism of action similar to VEC-162, for the treatment of insomnia.

### Limitations of current treatments

We believe that each of the drugs used to treat insomnia has inherent limitations that leave patients underserved. The key limitations include the potential for abuse, significant side effects, and a failure to address the underlying causes of sleeplessness:

Many of the products prescribed commonly for sleep disorders, including Ambien, Lunesta, and Sonata, are classified as Schedule IV controlled substances by the DEA due to their potential for abuse, tolerance and withdrawal symptoms. Drugs that are classified as Schedule IV controlled substances are subject to restrictions on how such drugs are prescribed and dispensed.

Many drugs approved for and used in sleep disorders also induce a number of nuisance side effects beyond the more serious abuse and addiction effects associated with most approved products. These side effects include next-day grogginess, memory loss, unpleasant taste, dry mouth and hormonal changes.

We believe that none of the drugs used and approved for sleep, other than Rozerem, work through the body s natural sleep/wake cycle, which is governed by melatonin. We believe that, for patients whose sleep disruption is due to a misalignment of this sleep/wake cycle and the patients need to sleep (as is the case in circadian rhythm sleep disorders), a drug that naturally modulates the sleep/wake cycle would be an attractive new alternative because it would address the underlying cause of the sleeplessness, rather than merely addressing its symptoms.

### Potential advantages of VEC-162

We believe that VEC-162 may offer efficacy similar to the most efficacious of the approved sleep drugs, and that it may provide significant benefits to patients beyond those offered by the approved drugs. We believe that VEC-162 is unlikely to be scheduled as a controlled substance by the DEA, because Rozerem, which has a similar mechanism of action to VEC-162, was shown not to have potential for abuse and was not classified as a Schedule IV controlled substance by the DEA. However, despite the fact that the drugs have a similar mechanism of action, our Phase III results demonstrate that VEC-162 may offer superior sleep maintenance to Rozerem. VEC-162 also appears to be safe and well-tolerated, with no significant side effects or effects on next-day performance. For patients with circadian rhythm sleep disorders, VEC-162 may be able to align the patient s sleep/wake cycle with their lifestyle, something we believe no approved sleep therapy has demonstrated. For example, in our Phase II trial of VEC-162 in transient insomnia with 37 healthy participants, VEC-162 induced a statistically significant (p<0.025) shift in circadian rhythm of up to five hours on the first night.

### Overview of Phase III clinical trial

We recently completed a randomized, double-blind, multi-center, placebo-controlled Phase III trial that enrolled 412 adults in a sleep laboratory setting using a phase-advance, first-night assessment model of induced transient insomnia. The trial examined VEC-162 dosed 30 minutes before bedtime at 20, 50 and 100 milligrams versus placebo.

VEC-162 achieved significant results in multiple endpoints captured using polysomnography (PSG) including:

Reduced duration of wake after sleep onset. Wake after sleep onset is defined as the number of minutes awake from the time the participant falls asleep to the end of the evaluation period. There was a significant reduction in wake after sleep onset compared with

placebo of 24.2 (p = 0.017), 33.7 (p = 0.001), and 17.5 (p = 0.081) minutes at 20, 50, and 100 mg respectively.

Latency to Persistent Sleep. Patients experienced a reduction in the time it took to achieve persistent sleep (otherwise known as latency). Specifically, there was an improvement in latency to persistent sleep compared with placebo of 21.5 (p < 0.001), 26.3 (p < 0.001), and 22.8 (p < 0.001) minutes at 20, 50, and 100 mg respectively.

Latency to non-awake. Patients experienced a reduction in the time it took to fall into the initial stage of sleep, or latency to non-awake. Specifically, there was improvement in latency to non-awake compared to placebo of 11.1 (p = 0.006), 14.3 (p < 0.001) and 12.3 (p = 0.002) minutes at 20, 50, and 100 mg, respectively.

Total Sleep Time. Patients had improved total sleep times compared with placebo of 33.7 (p = 0.002), 47.9 (p < 0.001) and 29.6 (p = 0.005) minutes at 20, 50, and 100 mg respectively.

The Phase III trial also demonstrated that VEC-162 was safe and well-tolerated, with no significant side effects versus placebo and no impairment of next-day performance or mood. We believe that we will need to conduct additional Phase III trials in chronic sleep disorders to receive FDA approval of VEC-162 for the treatment of insomnia. We expect to begin at least one of these additional trials in the second half of 2007.

## Potential indication for depression

We believe that VEC-162 may also be effective in treating depression. Agomelatine, another drug that acts on the brain s melatonin receptors, has shown efficacy and safety that compared favorably to an approved antidepressant, Paxil® (paroxetine, GSK), in a Phase III trial. While the precise mechanism for the effect of drugs like VEC-162, agomelatine and Rozerem, which act on the brain s melatonin receptors, is currently unknown, it is possible that by improving sleep, these drugs could improve mood because depressed patients are likely to have sleep disorders. It is also possible that mood disorders such as depression have an association with circadian rhythm misalignments.

Approximately 29 million adults in the United States suffer from some form of depression, over 11 million of whom are currently treated with a prescription antidepressant medication. Sales of antidepressants exceeded \$19 billion globally in 2005.

We believe that VEC-162 will be differentiated from approved antidepressants in several ways. In the Phase III trial of agomelatine described above, agomelatine showed significantly improved mood in two weeks, vs. four weeks for Paxil<sup>®</sup>. Consequently, VEC-162 may, with its similar properties to agomelatine, offer a more rapid onset of action than approved antidepressants. We believe that VEC-162 should also have an improved side effect profile when compared to approved products because it should not have the sexual side effects, weight gain, and sleep disruption associated with these products.

VEC-162 is ready for Phase II trials in depression. It has demonstrated an antidepressant effect in animal models and has completed several Phase I trials, including one with four weeks of exposure, showing none of the serious side effects associated with the approved antidepressants.

#### **Commercialization**

Given the size of the prescribing physician base for sleep and mood disorders, we plan to partner with a global pharmaceutical company for the development and commercialization of VEC-162 worldwide, although we have not yet identified such a partner.

### Intellectual property

VEC-162 and its formulations and uses are covered by a total of five patent and patent application families worldwide. The primary new chemical entity patent covering VEC-162 expires normally in 2017 in the United States and in most European markets. We believe that, like iloperidone, VEC-162 will meet the various criteria of the Hatch-Waxman Act and will receive five additional years of patent protection for VEC-162 in the United States, which would extend its patent protection in the United States until 2022. In Europe, similar legislative enactments provide for five-year extensions of European new chemical entity patents through the granting of Supplementary Protection Certificates, and we believe that VEC-162 will qualify for such an extension, which would extend European patent protection for VEC-162 until 2022. Several other patent applications covering uses of VEC-162 will, if granted, provide exclusive rights for these uses until 2026.

Our rights to the new chemical entity patent covering VEC-162 and related intellectual property have been acquired through a license with BMS. Please see License agreements below for a discussion of this license.

### **VSF-173**

VSF-173 is an oral compound that has demonstrated effects on animal sleep/wake patterns and gene expression patterns suggestive of a stimulant effect. The compound also demonstrated a stimulant effect in humans during clinical trials conducted by Novartis for Alzheimer s Disease. As a result of these observations, we are currently planning to begin the clinical evaluation of VSF-173 in excessive sleepiness. We intend to initiate a Phase II trial for VSF-173 in mid-2007. We believe the market opportunity for VSF-173 is significant. Sales of drugs to treat excessive sleepiness were approximately \$500 million in 2005.

## Pharmacogenetics and pharmacogenomics expertise

Our expertise in pharmacogenetics and pharmacogenomics enables us to acquire high quality, patent-protected clinical compounds that have been discovered and developed by other pharmaceutical firms. We can capitalize on the discovery and early development efforts of other firms by acquiring compounds with clinical safety and possibly efficacy data that we believe can benefit from our extensive pharmacogenetics and pharmacogenomics expertise.

Pharmacogenetics and pharmacogenomics start from the premise that a given drug will not just affect the target/receptor for which it was initially developed, but will in fact interact with many systems within the body. Proof of this comes from two different sources. We know, for instance, that most drugs have side effects. These typically result from a drug s interaction not just with its intended receptor in its intended organ system, but also with either that receptor outside the intended organ system or with other receptors entirely. There are many examples of drugs that were developed initially for one indication but were then shown to be effective for another. One example of this is Viagra® (sildenafil, Pfizer), which was developed initially for hypertension (high blood pressure) but proved more effective for erectile dysfunction. Being compound-focused enables us to forego the costly discovery work and start with compounds already known to be drugs, in that they are safe and interact with at least one biological system.

Starting with safe compounds ones that have completed at least Phase I safety trials we use our pharmacogenetics and pharmacogenomics expertise to understand the disease or diseases for which the drug has the optimal biological (and clinical) effect. We have used this expertise to identify potential points of differentiation for iloperidone and VSF-173. Beyond these two, we have already identified a number of unexpected signaling pathways attributable to known compounds using these techniques, and we have filed a number of patent applications based

on these findings. For each compound, we may choose to confirm our findings in animal studies. Compounds clearing this hurdle will be ready for Phase II trials.

Compounds that we would most likely consider attractive candidates for applying our expertise would meet the following criteria:

were initially developed by an established biopharmaceutical company

have already completed Phase I trials

are free of significant formulation issues

have potential for strong patent protection through composition of matter patents, new doses or new formulations

# License agreements

Our rights to develop and commercialize our clinical-stage product candidates are subject to the terms and conditions of licenses granted to us by other pharmaceutical companies.

## **Iloperidone**

We acquired exclusive worldwide rights to patents for iloperidone through a sublicense agreement with Novartis. A predecessor company of Sanofi-Aventis, Hoechst Marion Roussel, Inc. (HMRI), discovered iloperidone and completed early clinical work on the compound. In 1996, following a review of its product portfolio, HMRI licensed its rights to the iloperidone patents to Titan Pharmaceuticals, Inc. on an exclusive basis. In 1997, soon after it had acquired its rights, Titan sublicensed its rights to iloperidone on an exclusive basis to Novartis. In June 2004, we acquired exclusive worldwide rights to these patents to develop and commercialize iloperidone through a sublicense agreement with Novartis. In partial consideration for this sublicense, we paid Novartis an initial license fee of \$500,000 and are obligated to make future milestone payments to Novartis of less than \$100 million in the aggregate (the majority of which are tied to sales milestones), as well as royalty payments to Novartis at a rate which, as a percentage of net sales, is in the mid-twenties. Our rights with respect to the patents to develop and commercialize iloperidone may terminate, in whole or in part, if we fail to meet certain regulatory or commercialization milestones relating to the time it takes for us to launch iloperidone commercially following regulatory approval, and the time it takes for us to receive regulatory approval following our submission of an NDA or equivalent foreign filing. Additionally our rights may terminate in whole or in part if we do not meet certain other obligations under our sublicense agreement to make royalty and milestone payments, if we fail to comply with requirements in our sublicense agreement regarding our financial condition, or if we do not abide by certain restrictions in our sublicense agreement regarding our other development activities. Additionally, if we do not cure any breaches by Novartis or Titan of their respective obligations under their agreements with Titan and Sanofi-Aventis, respectively, our rights to develop and commercialize iloperidone may revert back to Novartis.

### **VEC-162**

In February 2004, we entered into a license agreement with BMS under which we received an exclusive worldwide license under certain patents and patent applications, and other licenses to intellectual property, to develop and commercialize VEC-162. In partial consideration for the license, we paid BMS an initial license fee of \$500,000. We made a milestone payment to BMS of \$1,000,000 under this license earlier in 2006. We are also obligated to make future milestone payments to BMS of less than \$40 million in the aggregate (the majority of which are tied to sales milestones) as well as royalty payments based on the net sales of VEC-162 at a rate which,

as a percentage of net sales, is in the low teens. We are also obligated under this agreement to pay BMS a percentage of any sublicense fees, upfront payments and milestone and other payments (excluding royalties) that we receive from a third party in connection with any sublicensing arrangement, at a rate which is in the mid-twenties. We have agreed with BMS in our license agreement for VEC-162 to use our commercially reasonable efforts to develop and commercialize VEC-162 and to meet certain milestones in initiating and completing certain clinical work.

BMS holds certain rights with respect to VEC-162 in our license agreement. For example, if we have not agreed to one or more partnering arrangements to develop and commercialize VEC-162 in certain significant markets with one or more third parties after the completion of our Phase III program, which may consist of several Phase III trials, BMS has the option to exclusively develop and commercialize VEC-162 on its own on pre-determined financial terms, including milestone and royalty payments.

Either party may terminate the VEC-162 license agreement under certain circumstances, including a material breach of the agreement by the other. In the event that BMS has not exercised its option to reacquire the rights to VEC-162 and we terminate our license, or if BMS terminates our license due to our breach, all rights licensed and developed by us under this agreement will revert or otherwise be licensed back to BMS on an exclusive basis.

#### **VSF-173**

In June 2004, we entered into a license agreement with Novartis under which we received an exclusive worldwide license to develop and commercialize VSF-173. In consideration for the license, we paid Novartis an initial license fee of \$500,000. We are also obligated to make future milestone payments to Novartis of less than \$50 million in the aggregate (the majority of which are tied to sales milestones) and royalty payments at rates which, as a percentage of net sales, range from the low-to-mid teens. Novartis has the right to co-develop and exclusively commercialize VSF-173 on its own after Phase II and Phase III in exchange for certain milestones and royalty payments. In the event that Novartis chooses not to exercise either of these options and we decide to enter into a partnering arrangement to commercialize VSF-173, Novartis has a right of first refusal to negotiate such an agreement with us, as well as a right to submit a last matching counteroffer regarding such an agreement. In addition, our rights with respect to VSF-173 may terminate, in whole or in part, if we fail to meet certain development and commercialization milestones described in our license agreement relating to the time it takes us to complete our development work on VSF-173. These rights may also terminate in whole or in part if we fail to make royalty or milestone payments or if we do not comply with requirements in our license agreement regarding our financial condition. In the event of an early termination of our license agreement, all rights licensed and developed by us under this agreement may revert back to Novartis.

## **Government regulation**

Government authorities in the United States, at the federal, state and local level, as well as foreign countries and local foreign governments, regulate the research, development, testing, manufacture, labeling, promotion, advertising, distribution, sampling, marketing, import and export of our product candidates. All of our products will require regulatory approval by government agencies prior to commercialization. In particular, human pharmaceutical products are subject to rigorous pre-clinical and clinical trials and other approval procedures of the FDA and similar regulatory authorities in foreign countries. The process of obtaining these approvals

and the subsequent compliance with appropriate domestic and foreign laws, rules and regulations require the expenditure of significant time and human and financial resources.

#### **United States government regulation**

In the United States, the FDA regulates drugs under the Federal Food, Drug and Cosmetic Act and implementing regulations. If we fail to comply with the applicable requirements at any time during the product development process, approval process, or after approval, we may become subject to administrative or judicial sanctions. These sanctions could include the FDA s refusal to approve pending applications, withdrawals of approvals, clinical holds, warning letters, product recalls, product seizures, total or partial suspension of our operations, injunctions, fines, civil penalties or criminal prosecution. Any such sanction could have a material adverse effect on our business.

The steps required before a drug may be marketed in the United States include:

pre-clinical laboratory tests, animal studies and formulation studies under cGLP

submission to the FDA of an investigational new drug application, or IND, which must become effective before human clinical trials may begin

execution of adequate and well-controlled clinical trials to establish the safety and efficacy of the product for each indication for which approval is sought

submission to the FDA of an NDA

satisfactory completion of an FDA inspection of the manufacturing facility or facilities at which the product is produced to assess compliance with cGMP

FDA review and approval of the NDA

Pre-clinical studies generally are conducted in laboratory animals to evaluate the potential safety and activity of a product. Violation of the FDA s good laboratory practices regulations can, in some cases, lead to invalidation of the studies, requiring these studies to be replicated. In the United States, drug developers submit the results of pre-clinical trials, together with manufacturing information and analytical and stability data, to the FDA as part of the IND, which must become effective before clinical trials can begin in the United States. An IND becomes effective 30 days after receipt by the FDA unless before that time the FDA raises concerns or questions about issues such as the proposed clinical trials outlined in the IND. In that case, the IND sponsor and the FDA must resolve any outstanding FDA concerns or questions before clinical trials can proceed. If these concerns or questions are unresolved, the FDA may not allow the clinical trials to commence.

Pilot studies generally are conducted in a limited patient population, approximately three to 25 subjects, to determine whether the product candidate warrants further clinical trials based on preliminary indications of efficacy. These pilot studies may be performed in the United States after an IND has become effective or outside of the United States prior to the filing of an IND in the United States in accordance with government regulations and institutional procedures.

Clinical trials involve the administration of the investigational product candidate to human subjects under the supervision of qualified investigators. Clinical trials are conducted under protocols detailing, among other things, the objectives of the study, the parameters to be used in assessing the safety and the effectiveness of the drug. Each protocol must be submitted to the FDA as part of the IND prior to beginning the trial.

Typically, clinical evaluation involves a time-consuming and costly three-Phase sequential process, but the phases may overlap. Each trial must be reviewed, approved and conducted under the

auspices of an independent Institutional Review Board, and each trial must include the patient s informed consent.

Phase I: refers typically to closely-monitored clinical trials and includes the initial introduction of an investigational new drug into human patients or health volunteer subjects. Phase I trials are designed to determine the safety, metabolism and pharmacologic actions of a drug in humans, the potential side effects associated with increasing drug doses and, if possible, to gain early evidence of the product candidate s effectiveness. Phase I trials also include the study of structure-activity relationships and mechanism of action in humans, as well as studies in which investigational drugs are used as research tools to explore biological phenomena or disease processes. During Phase I trials, sufficient information about a drug s pharmacokinetics and pharmacological effects should be obtained to permit the design of well-controlled, scientifically valid Phase II studies. The total number of subjects and patients included in Phase I trials varies, but is generally in the range of 20 to 80 people.

Phase II: refers to controlled clinical trials conducted to evaluate appropriate dosage and the effectiveness of a drug for a particular indication or indications in patients with a disease or condition under study and to determine the common short-term side effects and risks associated with the drug. These trials are typically well controlled, closely monitored and conducted in a relatively small number of patients, usually involving no more than several hundred subjects.

Phase III: refers to expanded controlled and uncontrolled clinical trials. These trials are performed after preliminary evidence suggesting effectiveness of a drug has been obtained. Phase III trials are intended to gather additional information about the effectiveness and safety that is needed to evaluate the overall benefit-risk relationship of the drug and to provide an adequate basis for physician labeling. Phase III trials usually include from several hundred to several thousand subjects.

Phase I, II and III testing may not be completed successfully within any specified time period, if at all. The FDA closely monitors the progress of each of the three phases of clinical trials that are conducted in the United States and may, at its discretion, reevaluate, alter, suspend or terminate the testing based upon the data accumulated to that point and the FDA is assessment of the risk/benefit ratio to the patient. A clinical program is designed after assessing the causes of the disease, the mechanism of action of the active pharmaceutical ingredient of the product candidate and all clinical and pre-clinical data of previous trials performed. Typically, the trial design protocols and efficacy endpoints are established in consultation with the FDA. Upon request through a special protocol assessment, the FDA can also provide specific guidance on the acceptability of protocol design for clinical trials. The FDA or we may suspend or terminate clinical trials at any time for various reasons, including a finding that the subjects or patients are being exposed to an unacceptable health risk. The FDA can also request additional clinical trials be conducted as a condition to product approval. During all clinical trials, physicians monitor the patients to determine effectiveness and to observe and report any reactions or other safety risks that may result from use of the drug candidate.

Assuming successful completion of the required clinical trials, drug developers submit the results of pre-clinical studies and clinical trials, together with other detailed information including information on the manufacture and composition of the product, to the FDA, in the form of an NDA, requesting approval to market the product for one or more indications. In most cases, the NDA must be accompanied by a substantial user fee. The FDA reviews an NDA to determine, among other things, whether a product is safe and effective for its intended use.

Before approving an application, the FDA will inspect the facility or facilities where the product is manufactured. The FDA will not approve the application unless cGMP compliance is satisfactory. The FDA will issue an approval letter if it determines that the application, manufacturing process and manufacturing facilities are acceptable. If the FDA determines that the application, manufacturing process or manufacturing facilities are not acceptable, it will outline the deficiencies in the submission and will often request additional testing or information. Notwithstanding the submission of any requested additional information, the FDA may ultimately decide that the application does not satisfy the regulatory criteria for approval and refuse to approve the application by issuing a not approvable letter.

The testing and approval process requires substantial time, effort and financial resources, and each may take several years to complete. The FDA may not grant approval on a timely basis, or at all. We may encounter difficulties or unanticipated costs in our efforts to secure necessary governmental approvals, which could delay or preclude us from marketing our products. Furthermore, the FDA may prevent a drug developer from marketing a product under a label for its desired indications or place other conditions on distribution as a condition of any approvals, which may impair commercialization of the product. After approval, some types of changes to the approved product, such as adding new indications, manufacturing changes and additional labeling claims, are subject to further FDA review and approval. Similar regulatory procedures must also be complied with in countries outside the United States.

If the FDA approves the new drug application, the drug becomes available for physicians to prescribe in the United States. After approval, the drug developer must comply with a number of post-approval requirements, including delivering periodic reports to the FDA, submitting descriptions of any adverse reactions reported, and complying with drug sampling and distribution requirements. The holder of an approved NDA is required to provide updated safety and efficacy information and to comply with requirements concerning advertising and promotional labeling. Also, quality control and manufacturing procedures must continue to conform to cGMP after approval. Drug manufacturers and their subcontractors are required to register their facilities and are subject to periodic unannounced inspections by the FDA to assess compliance with cGMP which imposes certain procedural and documentation requirements relating to quality assurance and quality control. Accordingly, manufacturers must continue to expend time, money and effort in the area of production and quality control to maintain compliance with cGMP and other aspects of regulatory compliance. The FDA may require post market testing and surveillance to monitor the product safety or efficacy, including additional studies, known as Phase IV trials, to evaluate long-term effects.

In addition to studies requested by the FDA after approval, a drug developer may conduct other trials and studies to explore use of the approved compound for treatment of new indications, which require FDA approval. The purpose of these trials and studies is to broaden the application and use of the drug and its acceptance in the medical community.

We use, and will continue to use, third-party manufacturers to produce our products in clinical and commercial quantities. Future FDA inspections may identify compliance issues at our facilities or at the facilities of our contract manufacturers that may disrupt production or distribution, or require substantial resources to correct. In addition, discovery of problems with a product or the failure to comply with requirements may result in restrictions on a product, manufacturer or holder of an approved NDA, including withdrawal or recall of the product from the market or other voluntary or FDA-initiated action that could delay further marketing. 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. Also, new

government requirements may be established that could delay or prevent regulatory approval of our products under development.

#### Foreign regulation

Whether or not we obtain FDA approval for a product, we must obtain approval of a product by the comparable regulatory authorities of foreign countries before we can commence clinical trials or marketing of the product in those countries. The approval process varies from country to country, and the time may be longer or shorter than that required for FDA approval. The requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement also vary greatly from country to country. Although governed by the applicable country, clinical trials conducted outside of the United States typically are administered with the three-Phase sequential process that is discussed above under United States government regulation. However, the foreign equivalent of an IND is not a prerequisite to performing pilot studies or Phase I clinical trials.

Under European Union regulatory systems, we may submit marketing authorization applications either under a centralized or decentralized procedure. The centralized procedure which is available for products produced by biotechnology or which are highly innovative, provides for the grant of a single marketing authorization that is valid for all European Union member states. This authorization is a marketing authorization approval. The decentralized procedure provides for mutual recognition of national approval decisions. Under this procedure, the holder of a national marketing authorization may submit an application to the remaining member states. Within 90 days of receiving the applications and assessment report, each member state must decide whether to recognize approval. This procedure is referred to as the mutual recognition procedure.

In addition, regulatory approval of prices is required in most countries other than the United States. We face the risk that the resulting prices would be insufficient to generate an acceptable return to us or our collaborators.

#### Third-party reimbursement and pricing controls

In the United States and elsewhere, sales of pharmaceutical products depend in significant part on the availability of reimbursement to the consumer from third-party payors, such as government and private insurance plans. Third-party payors are increasingly challenging the prices charged for medical products and services. It will be time consuming and expensive for us to go through the process of seeking reimbursement from Medicare and private payors. Our products may not be considered cost-effective, and coverage and reimbursement may not be available or sufficient to allow us to sell our products on a competitive and profitable basis. The passage of the Medicare Prescription Drug and Modernization Act of 2003 imposes new requirements for the distribution and pricing of prescription drugs which may affect the marketing of our products.

In many foreign markets, including the countries in the European Union, pricing of pharmaceutical products is subject to governmental control. In the United States, there have been, and we expect that there will continue to be, a number of federal and state proposals to implement similar governmental pricing control. While we cannot predict whether such legislative or regulatory proposals will be adopted, the adoption of such proposals could have a material adverse effect on our business, financial condition and profitability.

#### Marketing and sales

We currently have no sales, marketing or distribution capabilities. However, we plan to develop these capabilities internally to the extent that it is practical to do so, and enter into partnering arrangements to the extent that we believe large sales and marketing forces will be necessary. More specifically, in the United States, we expect to build our own sales force to market iloperidone and VSF-173 directly to psychiatrists and other target physicians. Because we believe that the number of physicians that would generate the majority of prescriptions for iloperidone and VSF-173 is relatively small, we believe that we can cost-effectively develop our own sales force to market and sell iloperidone and VSF-173. Outside of the U.S., we intend to find commercial partners for iloperidone and VSF-173. We will seek a global commercial partner for VEC-162.

## Patents and proprietary rights; Hatch-Waxman protection

We will be able to protect our products from unauthorized use by third parties only to the extent that our products are covered by valid and enforceable patents either licensed in from third parties or generated internally that give us sufficient proprietary rights. Accordingly, patents and other proprietary rights are essential elements of our business.

Our three current compounds in clinical development are covered by new chemical entity and other patents. These new chemical entity patents cover the active portions of our compounds and provide patent protection for all other compounds and formulations containing these active portions. The new chemical entity patent for iloperidone is owned by Sanofi-Aventis, and other patents and patent applications relating to iloperidone are owned by Sanofi-Aventis and Novartis. Novartis also owns the new chemical entity patent for VSF-173 and Bristol-Myers Squibb owns the new chemical entity patent for VEC-162. For all three compounds we have obtained exclusive worldwide rights to develop and commercialize the compounds covered by these patents through license and sublicense arrangements. For more on these license and sublicense arrangements, please see License agreements above. In addition, we have generated intellectual property, and filed patent applications covering this intellectual property, for each of the three compounds.

The new chemical entity patent covering iloperidone expires normally in 2011 in the United States and in 2010 in most European markets. The new chemical entity patent covering VEC-162 expires in 2017 in the United States and most European markets. The new chemical entity patent covering VSF-173 expires in 2014 in the United States and in 2012 in most European markets. Additionally, for each of our late-stage compounds, an additional period of exclusivity in the United States of up to five years following the expiration of the patent covering that compound may be obtained pursuant to the United States Drug Price Competition and Patent Term Restoration Act of 1984, more commonly known as the Hatch-Waxman Act. Assuming we gain such a five-year extension and that we continue to have our intellectual property rights under our sublicense and license agreements, we would have exclusive new chemical entity patent rights in the U.S. for iloperidone until 2016, for VEC-162 until 2022 and for VSF-173 until 2019. In Europe, similar legislative enactments may allow us to obtain five-year extensions of the European new chemical entity patents covering our product candidates through the granting of Supplementary Protection Certificates, which would allow us to have exclusive European new chemical entity patent rights for iloperidone until 2015, for VEC-162 until 2022 and for VSF-173 until 2017. Additionally, a recent directive in the European Union allows companies who receive European regulatory approval for a new compound to have a 10-year period of market exclusivity in most European countries for that compound (with the possibility of a further one-year extension), beginning on the date of such European regulatory approval, regardless of

when the European new chemical entity patent covering such compound expires. No generic version of an approved drug may be marketed or sold in most European countries during this 10-year period. This directive may be of particular importance with respect to iloperidone, since the European new chemical entity patent for iloperidone will likely expire prior to the end of this 10-year period of market exclusivity.

Aside from the new chemical entity patents covering our current late-stage compounds, as of December 31, 2006 we had 15 pending provisional patent applications in the United States and three pending Patent Cooperation Treaty applications. The claims in these various patents and patent applications are directed to compositions of matter, including claims covering other product candidates, pharmaceutical compositions, and methods of use.

For proprietary know-how that is not appropriate for patent protection, processes for which patents are difficult to enforce and any other elements of our discovery process that involve proprietary know-how and technology that is not covered by patent applications, we rely on trade secret protection and confidentiality agreements to protect our interests. We require all of our employees, consultants and advisors to enter into confidentiality agreements. Where it is necessary to share our proprietary information or data with outside parties, our policy is to make available only that information and data required to accomplish the desired purpose and only pursuant to a duty of confidentiality on the part of those parties.

## Manufacturing

We currently depend and expect to continue to depend on a small number of third-party manufacturers to produce sufficient quantities of our product candidates for use in our clinical studies. We are not obligated to obtain our product candidates from any particular third-party manufacturer and we believe that we would be able to obtain our product candidates from a number of third-party manufacturers at comparable cost.

If any of our product candidates are approved for commercial use, we plan to rely on third-party contract manufacturers to produce sufficient quantities for large-scale commercialization. If we do enter into commercial manufacturing arrangements with third parties, these third-party manufacturers will be subject to extensive governmental regulation. Specifically, regulatory authorities in the markets which we intend to serve will require that drugs be manufactured, packaged and labeled in conformity with cGMP or equivalent foreign standards. We intend to engage only those contract manufacturers who have the capability to manufacture drug products in compliance with cGMP and other applicable standards in bulk quantities for commercial use.

## Competition

The pharmaceutical industry and the central nervous system segment of that industry in particular, is highly competitive and includes a number of established large and mid-sized companies with greater financial, technical and personnel resources than we have and significantly greater commercial infrastructures than we have. Our market segment also includes several smaller emerging companies whose activities are directly focused on our target markets and areas of expertise. If approved, our product candidates will compete with numerous therapeutic treatments offered by these competitors. While we believe that our product candidates will have certain favorable features, existing and new treatments may also possess advantages. Additionally, the development of other drug technologies and methods of disease prevention are occurring at a rapid pace. These developments may render our product candidates or technologies obsolete or noncompetitive.

We believe the primary competitors for each of our product candidates are as follows:

For iloperidone in the treatment of schizophrenia, the atypical antipsychotics Risperdal® (risperidone) by Johnson & Johnson (including the depot formulation Risperdal® Consta®), Zyprexa® (olanzapine) by Eli Lilly, Seroquel® (quetiapine) by AstraZeneca, Abilify® (aripiprazole) by BMS/Otsuka, Geodon® (ziprasidone) by Pfizer, Invega® (paliperidone) by Johnson & Johnson, and generic clozapine, as well as the typical antipsychotics haloperidol, chlorpromazine, thioridazine, and sulpiride (all of which are generic). In addition to the approved products, compounds in Phase III trials (or for which an NDA has recently been filed) for the treatment of schizophrenia include bifeprunox (Wyeth/Solvay/Lundbeck A/S), and asenapine (Organon International).

For VEC-162 in the treatment of insomnia, Rozerem<sup>tm</sup> (ramelteon) by Takeda, hypnotics such as Ambien<sup>®</sup> (zolpidem) by Sanofi-Aventis (including Ambien CR<sup>®</sup>), Lunesta<sup>®</sup> (eszopiclone) by Sepracor and Sonata<sup>®</sup> (zaleplon) by King Pharmaceuticals, generic compounds such as trazodone and doxepin, and over-the-counter remedies such as Benadryl<sup>®</sup> and Tylenol PM<sup>®</sup>. In addition to the approved products, compounds in Phase III trials for insomnia include indiplon (Neurocrine Biosciences), gaboxadol (Merck/Lundbeck A/S), and low-dose doxepin (Silenor<sup>tm</sup>, Somaxon).

For VEC-162 in the treatment of depression, antidepressant drugs such as Paxil® (paroxetine) by GSK, Zoloft® (sertraline) by Pfizer, Prozac® (fluoxetine) by Eli Lilly, and Lexapro (escitalopram) by Lundbeck A/S /Forest Pharmaceuticals Inc., Effexor® (venlafaxine) by Wyeth as well as other compounds such as Wellbutrin® (buproprion) by GSK and Cymbalta® (duloxetine) by Eli Lilly. In addition to the approved products, compounds in Phase III trials for depression include agomelatine (Novartis and Les Laboratoires Servier).

For VSF-173 in the treatment of excessive sleepiness, Provigil® (modafinil) and NuVigil® (armodafinil) by Cephalon, and Xyrem® (sodium oxybate) by Jazz Pharmaceuticals, Inc.

Our ability to compete successfully will depend in part on our ability to utilize our pharmacogenetics and pharmacogenomics and drug development expertise to identify, develop, secure rights to and obtain regulatory approvals for promising pharmaceutical compounds before others are able to develop competitive products. Our ability to compete successfully will also depend on our ability to attract and retain skilled and experienced personnel. Additionally, our ability to compete may be affected because insurers and other third-party payors in some cases seek to encourage the use of cheaper, generic products, which could make our products less attractive.

## **Employees**

As of December 31, 2006 we had 44 full-time employees, 32 of whom were primarily engaged in research and development activities. 40 of our full-time employees work at our facility in Rockville, Maryland, and 4 of our full-time employees work at our Singapore research facility. None of our employees are represented by a labor union. We have not experienced any work stoppages and consider our employee relations to be good.

## **Facilities**

Our current headquarters are located in Rockville, Maryland, consisting of approximately 17,000 square feet of office and laboratory space. Our annual rent under our lease for this facility is approximately \$433,000, with an annual increase of 3% per year, until the expiration of the lease in 2016.

In January, 2006, we vacated our previous headquarters in Rockville, Maryland, and intend to exercise our sublease rights under the lease governing this facility. Pending such a sublease, we

remain obligated to make rent payments under this lease. Our annual rent under this lease for 2006 is approximately \$233,000, with an annual increase of 3% per year. The lease expires in 2008.

We have also entered into a lease for a research facility in Singapore. Our annual rent for this facility for 2006 is approximately \$76,000; the lease for the facility expired in December 2006. We intend to renew the Singapore in January 2007.

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## Management

#### **Executive officers and directors**

The following are our executive officers and directors as of December 31, 2006.

Name	Age	Position
Mihael H. Polymeropoulos, M.D.	46	President and Chief Executive Officer, Director
Paolo Baroldi, M.D., Ph.D.	55	Senior Vice President and Chief Medical Officer
William D. Chip Clark	38	Senior Vice President, Chief Business Officer and
		Secretary
Steven A. Shallcross	45	Senior Vice President, Chief Financial Officer and
		Treasurer
Thomas Copmann, Ph.D.	54	Vice President of Regulatory Affairs
Deepak Phadke, Ph.D.	56	Vice President of Manufacturing
Argeris N. Karabelas, Ph.D.(1),(3)	54	Director and Chairman of the Board of Directors
Richard W. Dugan(2)	64	Director
Brian K. Halak, Ph.D.(2),(3)	35	Director
H. Thomas Watkins(1),(3)	54	Director
David Ramsay(2)	43	Director
James B. Tananbaum, M.D.(1)	43	Director

- (1) Member of compensation committee.
- (2) Member of audit committee.
- (3) Member of nominating/governance Committee.

Mihael H. Polymeropoulos, M.D. has served as Chief Executive Officer and a Director of Vanda since May of 2003. Prior to joining Vanda, Dr. Polymeropoulos was Vice President and Head of the Pharmacogenetics Department at Novartis from 1998 to 2003. Prior to his tenure at Novartis, he served as Chief of the Gene Mapping Section, Laboratory of Genetic Disease Research, National Human Genome Research Institute, from 1992 to 1998. Dr. Polymeropoulos is the co-founder of the Integrated Molecular Analysis of Genome Expression (IMAGE) Consortium. Dr. Polymeropoulos holds a degree in Medicine from the University of Patras.

Paolo Baroldi, M.D., Ph.D. has served as a Senior Vice President and Chief Medical Officer at Vanda since July 2006. Prior to joining Vanda, Dr. Baroldi served as Vice President Corporate Drug Development at Chiesi Farmaceutici SpA, in Parma, Italy, from 2003 to 2006. Prior to his tenure at Chiesi, Dr. Baroldi was the Global Head of Clinical Pharmacology at Novartis AG from 1998 to 2002. Dr. Baroldi holds degrees in Medicine and Surgery and a Ph.D. in Clinical Pharmacology from the University of Milan, and an Executive Masters in Business Administration from Harvard University.

William D. Chip Clark has served as Senior Vice President and Chief Business Officer of Vanda since September of 2004 and served as a Director of Vanda from 2003 to 2004. Prior to joining Vanda, Mr. Clark was a Principal at Care Capital, LLC, a venture capital firm investing in biopharmaceutical companies, from 2000 to 2004. Prior to his tenure at Care Capital, he served in a variety of commercial roles at SmithKline Beecham (now part of GlaxoSmithKline), from 1990 to 2000. Mr. Clark holds a B.A. from Harvard University and an M.B.A. from The Wharton School at the University of Pennsylvania.

Steven A. Shallcross has served as Senior Vice President, Chief Financial Officer and Treasurer of Vanda since November of 2005. From October 2001 to November 2005, Mr. Shallcross was the Senior Vice President, Chief Financial Officer and Treasurer at Advancis Pharmaceutical Corporation, a specialty pharmaceutical company. Mr. Shallcross was the Vice President of Finance and Chief Financial Officer at Bering Truck Corporation, a truck manufacturer, from 1997 to 2001. From 1994 to 1997, Mr. Shallcross served as Vice President of Operations at Precision Scientific, Inc., a manufacturer of scientific laboratory equipment. He was the Controller of Precision Scientific from 1993 to 1994. Mr. Shallcross has over 20 years of senior financial and operations experience in emerging organizations, including acquisitions and restructurings. Mr. Shallcross received a bachelor s degree in accounting from the University of Illinois and an M.B.A. from the University of Chicago, Graduate School of Business. Mr. Shallcross is also a certified public accountant.

Thomas Copmann, Ph.D. has served as Vice President of Regulatory Affairs at Vanda since April of 2005. Prior to joining Vanda, Dr. Copmann served as Senior Director of Regulatory Affairs at Eli Lilly, from 2000 to 2005 and as a Director from 1995 to 2000. Prior to his tenure at Eli Lilly, Dr. Copmann was the Associate Vice President for Regulatory Affairs and Executive Director for the Commission on Drugs for Rare Diseases at the Pharmaceutical Manufacturers Association, from 1989 to 1995. Dr. Copmann holds an M.S. in Endocrinology and a Ph.D. in Physiology from Kent State University.

Deepak Phadke, Ph.D. has served as Vice President of Manufacturing at Vanda since August of 2005. Prior to joining Vanda, Dr. Phadke served as Executive Director of Pharmaceutical Sciences at Beckloff Associates, a pharmaceutical research and development consulting company located in the Kansas City area, from 1998 to 2005. Prior to his tenure at Beckloff Associates, Dr. Phadke served as a manager and research scientist in the formulation development departments at Hoechst Marion Roussel and its predecessor companies in Kansas City and Indianapolis, from 1986 to 1998. Dr. Phadke also worked as a senior pharmaceutical chemist at Rorer Group Inc. in Fort Washington, Pennsylvania from 1983 to 1986. Dr. Phadke holds a B.S. and an M.S. in Pharmacy and Pharmaceutics, respectively, from Nagpur University in India, and a Ph.D. in Pharmaceutics from Rutgers University.

Argeris N. Karabelas, Ph.D. has served as a Director and Chairman of the Board since 2003, when he co-founded Vanda with Dr. Polymeropoulos. Dr. Karabelas has served as a Partner of Care Capital, LLC since 2001. Prior to his tenure at Care Capital, Dr. Karabelas was the Founder and Chairman of the Novartis BioVenture Fund, from July 2000 to December 2001. From 1998 to 2000, he served as Head of Healthcare and CEO of Worldwide Pharmaceuticals for Novartis. Prior to joining Novartis, Dr. Karabelas was Executive Vice President of SmithKline Beecham responsible for U.S. operations, European operations, Regulatory, and Strategic Marketing, from 1981 to 1998. He is a member of the Scientific Advisory Council of the Massachusetts General Hospital, the Harvard-MIT Health Science and Technology Visiting Committee, Chairman of Human Genome Sciences, Inc., Chairman and interim Chief Executive Officer of NitroMed, Inc., Chairman of SkyePharma plc, Chairman of Inotek, Inc., a director of Renovo, plc and a Trustee of Fox Chase Cancer Center and the Philadelphia University of the Sciences. Dr. Karabelas holds a Ph.D. in Pharmacokinetics from the Massachusetts College of Pharmacy.

Richard W. Dugan has served as a Director of Vanda since December of 2005. From 1976 to September 2002, Mr. Dugan served as a Partner with Ernst & Young, LLP, where he served in a variety of managing and senior partner positions, including Mid-Atlantic Area Senior Partner from 2001 to 2002, Mid-Atlantic Area Managing Partner from 1989 to 2001 and Pittsburgh Office Managing Partner from 1979 to 1989. Mr. Dugan retired from Ernst & Young, LLP in September 2002. Mr. Dugan currently serves on the board of directors of two other publicly-

traded pharmaceutical companies, Advancis Pharmaceutical Corporation and Critical Therapeutics, Inc. and on the board of directors of a privately-owned pharmaceutical company, Xanthus Pharmaceuticals, Inc. Mr. Dugan holds a B.S.B.A. from Pennsylvania State University.

Brian K. Halak, Ph.D. has served as a Director of Vanda since 2004. Dr. Halak has served as a Principal at Domain Associates, a venture capital firm based in Princeton, New Jersey, since 2001 and became a Partner in January 2006. Prior to joining Domain Associates, he served as an Associate of the venture capital firm Advanced Technology Ventures, from 2000 to 2001. Dr. Halak serves on the Investment Advisory Council for Ben Franklin Technology Partners and BioAdvance, both seed stage investment groups in Philadelphia. Dr. Halak holds a B.S.E. from the University of Pennsylvania and a Ph.D. in Immunology from Thomas Jefferson University.

H. Thomas Watkins has served as a Director of Vanda since September 2006. Mr. Watkins has served as the President and Chief Executive Officer of Human Genome Sciences, Inc. and as a member of its board of directors since 2004. Prior to his tenure at Human Genome Sciences Inc., Mr. Watkins served as President of TAP Pharmaceutical Products, Inc. Mr. Watkins previously held a series of executive positions over the course of nearly twenty years with Abbott Laboratories. Mr. Watkins also serves on the Board of Trustees of the College of William and Mary Foundation, and is a member of the College of William and Mary Mason School of Business Foundation. He holds a bachelor s degree from the College of William and Mary, and a master s degree in business administration from the University of Chicago Graduate School of Business.

David Ramsay has served as a Director of Vanda since 2004. Mr. Ramsay has served as a Partner of Care Capital, LLC, which he co-founded in 2000. Prior to founding Care Capital, Mr. Ramsay served as a Managing Director of the Rhône Group, LLC, from 1997 to 2000 and co-founded Rhône Capital, LLC, a private equity investment fund. Mr. Ramsay previously worked at Morgan Stanley Capital Partners. Mr. Ramsay holds an A.B. in Mathematics from Princeton University and an M.B.A. from the Stanford University Graduate School of Business.

James B. Tananbaum, M.D. has served as a Director of Vanda since 2004. Dr. Tananbaum has served as a Managing Partner of Prospect Venture Partners II, a dedicated life science venture fund group which he co-founded in 2000. Prior to co-founding Prospect Venture Partners, he co-founded and served as Chief Executive Officer of Theravance, Inc. from 1997 to 2000. Dr. Tananbaum also served as a Partner at Sierra Ventures, from 1993 to 1997. Dr. Tananbaum co-founded GelTex Pharmaceuticals, Inc. in 1991. He is an officer of the Young Presidents Organization, Golden Gate Chapter and a member of the World Economic Forum and the Harvard-MIT Health Science and Technology Visiting Committee. Dr. Tananbaum serves as a director of numerous public and private healthcare companies, including Cogentus Pharmaceuticals, Inc., Jazz Pharmaceuticals, Inc., PathWorks, Inc. and Novavax, Inc. Dr. Tananbaum holds a bachelor s degree and a B.S.E.E. from Yale University and an M.D. and an M.B.A. from Harvard University.

## **Election of officers**

Our officers are elected by our board of directors on an annual basis and serve until their successors are duly elected and qualified. There are no family relationships among any of our officers or directors.

## **Board composition**

Our amended and restated certificate of incorporation provides for a classified board of directors consisting of three classes of directors, each serving a staggered three-year term. As a result, a portion of our board of directors are elected each year. Dr. Tananbaum and Messrs. Ramsay and Watkins have been designated Class I directors whose term will expire at

the 2007 annual meeting of stockholders. Dr. Halak and Mr. Dugan have been designated Class II directors whose term will expire at the 2008 annual meeting of stockholders. Drs. Polymeropoulos and Karabelas have been designated Class III directors whose term expires at the 2009 annual meeting of stockholders. Our amended and restated bylaws provide that the number of authorized directors may be changed only by resolution of a number of directors that is more than half of the number of directors then authorized (including any vacancies). Any additional directorships resulting from an increase in the number of authorized directors will be distributed among the three classes so that, as nearly as reasonably possible, each class will consist of one-third of the directors. The classification of the board of directors may have the effect of delaying or preventing changes in control of our company.

We believe that each of our board members other than Dr. Polymeropoulos is an independent director under Nasdaq Marketplace Rule 4200(a)(15).

#### **Committees of the board of directors**

Our board of directors has a compensation committee, an audit committee and a nominating/corporate governance committee.

*Compensation committee.* Three directors comprise the compensation committee of the board of directors: Argeris N. Karabelas, Ph.D., James B. Tananbaum, M.D. and H. Thomas Watkins. Dr. Karabelas chairs the compensation committee.

The compensation committee reviews and makes recommendations to the board of directors regarding the overall compensation strategy and policies for the Company. Specifically, the committee reviews and makes recommendations to the board of directors regarding corporate performance goals and objectives relevant to the compensation of our executive officers and other senior management; reviews and makes recommendations to the board of directors regarding the compensation and other terms of employment of our Chief Executive Officer and other executive officers; reviews and makes recommendations to the board of directors regarding the individual bonus programs in effect for the Chief Executive Officer, other executive officers and key employees for each fiscal year; recommends to the board of directors the compensation of the directors; recommends to the board of directors the adoption or amendment of equity and cash incentive plans; recommends to the board of directors amendments to such plans; grants (subject to the ratification of the full board of directors in the case of the Company s executives) stock options and other stock-related awards; and administers our Second Amended and Restated Management Plan and 2006 Equity Incentive Plan. A more detailed description of the committee s functions can be found in our compensation committee charter. The charter is published in the corporate governance section of our website at www.vandapharma.com.

The compensation committee met five times during the fiscal year. The Chief Executive Officer does not participate in the determination of his own compensation or the compensation of directors. However, he makes recommendations to the committee regarding the amount and form of the compensation of the other executive officers and key employees, and he often participates in the committee s deliberations about their compensation. No other executive officers participate in the determination of the amount or form of the compensation of executive officers or directors.

The compensation committee retained Towers Perrin as its independent compensation consultant for a compensation review in November 2006. The consultant served at the pleasure of the committee, and the consultant served approved by the committee. The consultant provided

the committee with a report regarding the compensation paid by the Company s competitors and other employers who compete with the Company for executives. The Towers Perrin report is described below in Executive compensation benchmarking of base compensation and equity holdings.

*Audit committee*. The members of our audit committee are Messrs. Dugan and Ramsay and Dr. Halak. Mr. Dugan chairs the audit committee. Mr. Dugan serves as our audit committee financial expert and is an independent director under applicable SEC and Nasdaq rules. Our audit committee, among other duties:

appoints a firm to serve as independent accountant to audit our financial statements

discusses the scope and results of the audit with the independent accountant, and reviews with management and the independent accountant our interim and year-end operating results

considers the adequacy of our internal accounting controls and audit procedures

approves (or, as permitted, pre-approves) all audit and non-audit services to be performed by the independent accountant

The audit committee has the sole and direct responsibility for appointing, evaluating and retaining our independent auditors and for overseeing their work. All audit services and all non-audit services, other than de minimis non-audit services, to be provided to us by our independent auditors must be approved in advance by our audit committee. A more detailed description of the audit committee s responsibilities can be found in our audit committee charter. The charter is published in the corporate governance section of our website at <a href="https://www.vandapharma.com">www.vandapharma.com</a>. We believe that the composition of our audit committee meets the requirements for independence under the current Nasdaq Global Market and SEC rules and regulations.

Nominating/corporate governance committee. The members of our nominating/corporate governance committee are Drs. Halak and Karabelas and Mr. Watkins. Dr. Halak chairs the nominating/corporate governance committee. Our nominating/corporate governance committee identifies, evaluates and recommends nominees to our board of directors and committees of our board of directors, conducts searches for appropriate directors, and evaluates the performance of our board of directors and of individual directors. The nominating/corporate governance committee is also responsible for reviewing developments in corporate governance practices, evaluating the adequacy of our corporate governance practices and reporting and making recommendations to the board of directors concerning corporate governance matters. A more detailed description of the nominating/corporate governance committee is responsibilities can be found in our nominating/corporate governance charter. The charter is published in the corporate governance section of our website, <a href="https://www.vandapharma.com">www.vandapharma.com</a>.

## Compensation committee interlocks and insider participation

None of the members of the compensation committee was at any time during the 2006 fiscal year an officer or employee of the Company. No executive officer of the Company serves as a member of the board of directors or compensation committee of any other entity that has one or more executive officers serving as a member of our board of directors or compensation committee.

## **Director compensation**

On December 19, 2005, our board of directors adopted a compensation program for outside directors. Pursuant to this program, each member of our board of directors who is not our

employee receives a \$25,000 annual fee as well as \$2,500 for each board meeting attended in person (\$1,250 for meetings attended by telephone conference). The chairman of the board of directors receives an additional annual fee of \$10,000, and the chairman of each committee of the board of directors receives an additional annual fee of \$2,000. Each director receives \$1,000 for each meeting of any committee of the board of directors attended in person or by telephone conference.

Under the director compensation program adopted on December 19, 2005, each member of our board of directors who is not our employee and who is elected after December 19, 2005 initially receives a nonstatutory option to purchase 35,000 shares of our common stock upon election, and each member of our board of directors who is not our employee will also receive, upon the conclusion of each annual meeting of our stockholders, an option to purchase 15,000 shares of our common stock. The stock option granted upon election vests and becomes exercisable in equal monthly installments over a period of four years from the date of the grant, except that in the event of a change of control the option will accelerate and become immediately exercisable. Each annual stock option vests and becomes exercisable in equal monthly installments over a period of one year from the date of grant, except that in the event of a change of control the option will accelerate and become immediately exercisable. All of these options have an exercise price equal to the fair market value of our common stock on the date of the grant. In cases where a director is serving as such on behalf of an entity, we may issue a warrant directly to such entity as consideration for the services provided in lieu of granting an option to the director himself.

The following table shows the compensation earned by each of our non-officer directors for the year ended December 31, 2006:

	Fee	es earned or		
Name and principal position	pai	id in cash(\$)	Option awards(\$)(4)	Total(\$)
Argeris N. Karabelas, Ph.D. (Chairman)(1)	\$	56,750		\$ 56,750
Richard W. Dugan	\$	52,500	\$ 37,622(5)	\$ 90,122
Brian K. Halak, Ph.D.(1)	\$	58,250		\$ 58,250
Wayne T. Hockmeyer, Ph.D.(1)(2)	\$	37,750		\$ 37,750
David A. Ramsay(1)	\$	51,500		\$ 51,500
James B. Tananbaum, M.D.(1)	\$	43,500		\$ 43,500
H. Thomas Watkins(3)	\$	27,417	\$ 34,166(6)	\$ 61,583

- (1) Fees earned by Dr. Karabelas, Dr. Halak, Dr. Hockmeyer, Mr. Ramsay and Dr. Tananbaum were paid to the management companies of the venture capital funds affiliated with these directors.
- (2) Dr. Hockmeyer resigned from our board of directors effective September 12, 2006.
- (3) Mr. Watkins was appointed to our board of directors effective September 12, 2006.
- (4) This column reflects the compensation cost for the year ended December 31, 2006 of each director s options, calculated in accordance with SFAS 123(R) and using a Black-Scholes valuation model. See note 4 of Notes to condensed consolidated financial statements for a discussion of assumptions made by the Company in determining the grant date fair value and compensation costs of our equity awards.

- (5) As of December 31, 2006, Mr. Dugan held options to purchase an aggregate of 10,571 shares of our common stock, 2,642 shares of which were vested as of December 31, 2006.
- (6) As of December 31, 2006, Mr. Watkins held options to purchase an aggregate of 35,000 shares of our common stock, 2,916 shares of which were vested as of December 31, 2006.

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#### **Executive compensation**

## Compensation discussion and analysis

This section discusses the principles underlying our executive compensation decisions and the most important factors relevant to an analysis of these decisions. It provides qualitative information regarding the manner and context in which compensation is awarded to and earned by our executive officers and places in perspective the data presented in the tables and other quantitative information that follows this section.

Our compensation of executives is designed to attract, as needed, individuals with the skills necessary for us to achieve our business plan, to reward those individuals fairly over time, and to retain those individuals who continue to perform at or above our expectations. Our executives compensation has three primary components salary, a yearly cash incentive bonus, and stock option awards. In addition, we provide our executives with benefits that are generally available to our salaried employees. We fix the base salary of each of our executives at a level we believe enables us to hire and retain individuals in a competitive environment and rewards satisfactory individual performance and a satisfactory level of contribution to our overall business goals. We also take into account the base salaries paid by similarly situated companies in the field of biotechnology and the base salaries of other private and public companies with which we believe we compete for talent. To this end, we subscribe to certain executive compensation surveys and other databases and review them periodically and when making a crucial executive hiring decision and annually when we review executive compensation. We designed the cash incentive bonuses for each of our executives to focus him on achieving key clinical, operational and/or financial objectives within a yearly time horizon, as described in more detail below. We use stock options to reward long-term performance; these options are intended to produce significant value for each executive if the Company s performance is outstanding and if the executive has an extended tenure.

We view the three components of our executive compensation as related but distinct. Although our compensation committee does review total compensation, we do not believe that significant compensation derived from one component of compensation should negate or reduce compensation from other components. We determine the appropriate level for each compensation component based in part, but not exclusively, on our view of internal equity and consistency, individual performance and other information we deem relevant, such as the survey data referred to above. We believe that, as is common in the biotechnology sector, stock option awards are the primary motivator in attracting and retaining executives, and that salary and cash incentive bonuses are secondary considerations. Except as described below, our compensation committee has not adopted any formal or informal policies or guidelines for allocating compensation between long-term and currently paid out compensation, between cash and non-cash compensation, or among different forms of compensation. This is due to the small size of our executive team and the need to tailor each executive s award to attract and retain that executive.

Our compensation committee s current intent is to perform annually a strategic review of our executive officers cash compensation and share and option holdings to determine whether they provide adequate incentives and motivation to our executive officers and whether they adequately compensate our executive officers relative to comparable officers in other companies. Our compensation committee s most recent review occurred in November 2006, and utilized a report from Towers Perrin, a well-known consulting firm specializing in executive compensation. This review is described in more detail below. Compensation committee meetings typically have included, for all or a portion of each meeting, not only the committee members

but also our President and Chief Executive Officer, our Chief Business Officer and our Chief Financial Officer. For compensation decisions, including decisions regarding the grant of equity compensation relating to executive officers (other than our President and Chief Executive Officer), the compensation committee typically considers the recommendations of our President and Chief Executive Officer.

We account for the equity compensation expense for our employees under the rules of SFAS 123R, which requires us to estimate and record an expense for each award of equity compensation over the service period of the award. Accounting rules also require us to record cash compensation as an expense at the time the obligation is accrued. Until we achieve sustained profitability, the availability to us of a tax deduction for compensation expense is not material to our financial position. We structure cash incentive bonus compensation so that it is taxable to our employees at the time it becomes available to them. It is not anticipated that any executive officer s annual cash compensation will exceed \$1 million, and the Company has accordingly not made any plans to qualify for any compensation deductions under Section 162(m) of the Internal Revenue Code.

# Benchmarking of base compensation and equity holdings

At its November 2006 meeting, our compensation committee determined that our respective executive officers salaries, cash incentive bonuses and equity holdings were at or near the median of executives with similar roles at comparable pre-public and recently public companies and that no material changes should be made to the compensation levels of our executive officers until our annual executive performance reviews are conducted early in the first quarter of 2007, other than the grant of additional options made in December 2006 to Drs. Baroldi, Copmann and Phadke reflected in the Grants of plan-based awards table below. This median was derived based on a report we obtained from Towers Perrin. The report compared our executive compensation with that of 22 comparable companies, including Myogen, Inc., New River Pharmaceuticals Inc., Keryx Biopharmaceuticals, Inc., Idenix Pharmaceuticals, Inc., AtheroGenics, Inc., CV Therapeutics, Inc., Xeno Port, Inc., Renovis, Inc., Santarus, Inc., Dendreon Corporation, Osiris Therapeutics, Inc., Coley Pharmaceutical Group, Inc., Somaxon Pharmaceuticals, Inc., Inspire Pharmaceuticals, Inc., Tercica, Inc., CoTherix, Inc., Barrier Therapeutics, Inc., Insmed Incorporated, Dynavax Technologies Corporation, ISTA Pharmaceuticals, Inc., Acorda Therapeutics, Inc. and DOV Pharmaceutical, Inc., analyzing various factors such as geography, employee headcount, research and development expenses, capitalization, product candidate pipeline, and therapeutic focus. Our compensation committee realizes that benchmarking the Company s compensation against the compensation earned at comparable companies may not always be appropriate, but believes that engaging in a comparative analysis of the Company s compensation practices is useful at this point in the life cycle of the Company. In instances where an executive officer is uniquely key to our success, the compensation committee may provide compensation above the median referred to above. The committee s choice not to recommend to the board of directors immediate material changes to the compensation levels following its review of the Towers Perrin report reflects our consideration of stockholders interests in paying what is necessary, but not significantly more than necessary, to achieve our corporate goals while conserving cash and equity as much as is practicable. We believe that, given the industry in which we operate and the corporate culture we have created, our compensation levels are generally sufficient to retain our existing executive officers and to hire new executive officers when and as required.

#### Equity compensation

All option grants made prior to our initial public offering on April 12, 2006 were made at what our board of directors determined to be the fair market value of our common stock on the respective grant dates. Beginning in the fourth quarter of 2005 our board of directors analyzed these grants for the years ended December 31, 2004 and December 31, 2005 retrospectively. As a result of this retrospective analysis, we determined that the fair value of our common stock on a fully-diluted basis steadily increased from \$3.21 per share at March 31, 2004 to \$17.18 per share at December 31, 2005, even though our options were granted between the range of \$0.33 to \$4.73 per share on those dates. For more information on this retrospective analysis, please see the section of this prospectus entitled Management s discussion and analysis of financial condition and results of operations Stock-based compensation. Since our initial public offering on April 12, 2006 we have made option grants based on the closing market value of our stock as reported on The Nasdaq Global Market on the date of grant. The value of the shares subject to our 2006 option grants to executive officers is reflected in the Summary compensation table and Grants of plan-based awards tables below.

We do not have any program, plan or obligation that requires us to grant equity compensation to any executive on specified dates. The authority to make equity grants to executive officers rests with our compensation committee (subject to ratification by the full board of directors), although, as noted above, the compensation committee does consider the recommendations of its President and Chief Executive Officer in setting the compensation of our other executives. All of our option grants made prior to our initial public offering, including all grants to executives, were made under our Second Amended and Restated Management Equity Plan. All of our option grants since our initial public offering have been made under our 2006 Equity Incentive Plan.

#### Cash incentive bonuses

Yearly cash incentive bonuses for our executives are established as part of their respective individual employment agreements. Each of these employment agreements provides that the executive will receive a cash incentive bonus determined in the discretion of our board of directors, with a target bonus amount specified for that executive based on individualized objective and subjective criteria. These criteria are established by the compensation committee and approved by the full board of directors on an annual basis, and include specific objectives relating to the achievement of clinical, regulatory, business and/or financial milestones. The target cash incentive bonus amount for each of our executives is as follows:

Mihael H. Polymeropoulos, M.D., President and Chief Executive Officer: 40% of base salary

Paolo Baroldi, M.D., Ph.D., Senior Vice President and Chief Medical Officer: 25% of base salary

William D. Chip Clark, Senior Vice President and Chief Business Officer: 25% of base salary

Steven A. Shallcross, Senior Vice president and Chief Financial Officer: 25% of base salary

Thomas Copmann, Ph.D., Vice President of Regulatory Affairs: 28% of base salary

Deepak Phadke, Vice President of Manufacturing: 15% of base salary

The compensation committee has not yet reviewed the extent to which our executive officers have met the individualized performance criteria established by the committee for the year ended December 31, 2006. Accordingly, the compensation committee has not determined the cash incentive bonus amounts that should be earned by our executive officers for the year ended December 31, 2006. These amounts will be determined and paid early in the first

quarter of 2007. Given that the Company has not yet generated revenues, the compensation committee

has not considered whether the Company would attempt to recover cash incentive bonuses to the extent that they were paid based on our financial performance and our quarterly revenue or net income is restated in a downward direction.

## Severance and change in control benefits

Each of our executives has a provision in his employment agreement providing for certain severance benefits in the event of termination without cause, as well as a provision providing for the acceleration of his then unvested options in the event of termination without cause following a change in control of the Company. These severance and acceleration provisions are described in the Employment Agreements section below, and certain estimates of these change of control benefits are provided in Estimated payments and benefits upon termination below.

## Other benefits

Our executives are eligible to participate in all of our employee benefit plans, such as medical, dental, vision, group life and disability insurance and our 401(k) plan, in each case on the same basis as our other employees. There were no special benefits or perquisites provided to any executive officer in 2006.

## **Summary compensation table**

The following table sets forth all of the compensation awarded to, earned by, or paid to the Company s principal executive officer, principal financial officer and the four other highest paid executive officers (our named executive officers) for the year ended December 31, 2006:

			Ontion	All other	
Name and principal position	Year	<b>Salary(\$)Bonus (\$)(1)</b>	Option awards <b>(\$)</b> mp	pensation (\$)	<b>Total</b> (\$)(1)
Mihael H. Polymeropoulos, M.D. President and Chief Executive Officer	2006	\$ 375,533	\$ 2,976,966(3)	\$ 6,582(5)	\$ 3,359,081
Paolo Baroldi, M.D., Ph.D. Senior Vice President and Chief Medical Officer(2)	2006	\$ 122,917 \$ 30,000	\$ 103,197(3)	\$ 63,619(6)	\$ 319,732
William D. Chip Clark Senior Vice President, Chief Business Officer and Secretary	2006	\$ 235,971	\$ 1,493,222(3)	\$ 3,039(5)	\$ 1,732,232
Steven A. Shallcross Senior Vice President, Chief Financial Officer and Treasurer	2006	\$ 250,000	\$ 585,619(3)	\$ 6,600(5)	\$ 842,219
Thomas Copmann, Ph.D. Vice President of Regulatory Affairs	2006	\$ 207,333	\$ 135,860(4)	\$ 6,600(5)	\$ 349,793
Deepak Phadke, Ph.D. Vice President of Manufacturing	2006	\$ 176,233 \$ 10,000	\$ 115,835(3)	\$ 4,800(5)	\$ 306,868

- (1) Bonus and total amounts do not include 2006 cash incentive bonuses accrued in 2006 pursuant to our named executive officers employment agreements, as described in Compensation discussion and analysis Cash incentive bonuses above. These amounts will be determined and paid early in the first quarter of 2007. The target amounts of these cash incentive bonuses are set forth as target estimated future payouts under Non-equity incentive plan awards in the Grants of plan based awards table below.
- (2) Dr. Baroldi joined the Company in July 2006.
- (3) Amount reflects the compensation cost for the year ended December 31, 2006 of the named executive officer s options, calculated in accordance with SFAS 123(R) and using a Black-Scholes valuation model. See note 4 of Notes to condensed consolidated financial statements for a discussion of assumptions made by the Company in determining the grant date fair value and compensation costs of our equity awards.
- (4) Amount reflects the aggregate compensation cost for the year ended December 31, 2006 of Dr. Copmann s options and also 5,665 shares of restricted stock which were issued upon the early exercise of an option owned by Dr. Copmann and which vested to him during the year ended December 31, 2006, in each case calculated in accordance with SFAS 123(R) and using a

Black-Scholes valuation model. See note 4 of Notes to condensed consolidated financial statements for a discussion of assumptions made by the Company in determining the grant date fair value and compensation costs of our equity awards.

- (5) Includes matching contributions made by the Company to executives respective 401(k) plan contributions.
- (6) Includes \$938 in matching contributions made by the Company to Dr. Baroldi s 401(k) plan contributions, \$49,934 in relocation expenses paid by the Company, and \$12,747 in tax costs paid by the Company relating to such relocation expenses.

#### **Employment agreements**

We have entered into offer letters or employment agreements with each of Mihael H. Polymeropoulos, M.D., our Chief Executive Officer, Paolo Baroldi, M.D., Ph.D., our Chief Medical Officer, William D. Chip Clark, our Chief Business Officer, Steven A. Shallcross, our Chief Financial Officer, Thomas Copmann, our Vice President of Regulatory Affairs, and Deepak Phadke, our Vice President of Manufacturing.

Mihael Polymeropoulos, M.D. We entered into an employment agreement in February 2005 with Dr. Polymeropoulos, our President and Chief Executive Officer, which provides for an annual base salary of not less than \$362,250 and the possibility of an annual target cash incentive bonus amount equal to 40% of his annual base salary upon achievement of certain performance goals. If Dr. Polymeropoulos employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 12 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount determined as follows: (i) if he is terminated prior to the first anniversary of this agreement, a pro-rata portion of the anticipated first-year target cash incentive bonus will be given to him; (ii) if he is terminated on or following the first anniversary and prior to the third, the amount will equal the greater of the most recent target cash incentive bonus or the average target cash incentive bonuses awarded for the prior years; or (iii) if he is terminated on or following the third anniversary, the amount will be equal to the greater of the most recent target cash incentive bonus or the average target cash incentive bonus awarded for the prior three years. In addition, if, following a change in control, Dr. Polymeropoulos is terminated without cause, or he terminates his employment for good reason, he will become vested in 100% of his then unvested shares and options.

Paolo Baroldi, M.D., Ph.D. We entered into an employment agreement in July 2006 with Dr. Baroldi, our Senior Vice President and Chief Medical Officer, which provides for an annual base salary of not less than \$250,000 and the possibility of an annual target cash incentive bonus equal to 25% of his annual base salary, prorated for fiscal year 2006 based on the number of days worked in that year. Additionally, Dr. Baroldi received a one-time bonus of \$30,000 and the Company paid \$62,681 for relocation expenses and tax costs relating to such relocation expenses. If Dr. Baroldi s employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 12 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount determined as follows: (i) if he is terminated prior to the first anniversary of this agreement, a pro-rata portion of the anticipated first-year target cash incentive bonus will be given to him; (ii) if he is terminated on or following the first anniversary and prior to the third, the amount will equal the greater of the most recent target cash incentive bonus or the average target cash incentive bonuses awarded for the prior years; or (iii) if he is terminated on or following the third anniversary, the amount will be equal to the greater of the most recent target cash incentive bonus or the average target cash incentive bonus awarded for the prior three years. In addition, if, following a change in control,

Dr. Baroldi is terminated without cause, or he terminates his employment for good reason, he will become vested in 24 months worth of his then unvested shares and options.

William D. Chip Clark. We entered into an employment agreement in February 2005 with Mr. Clark, our Senior Vice President, Chief Business Officer and Secretary, which provides for an annual base salary of not less than \$227,625 and the possibility of an annual target cash incentive bonus equal to 25% of his annual base salary upon achievement of certain performance criteria. If Mr. Clark s employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 12 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount determined as follows: (i) if he is terminated prior to the first anniversary of this agreement, a pro-rata portion of the anticipated first-year target cash incentive bonus will be given to him; (ii) if he is terminated on or following the first anniversary and prior to the third, the amount will equal the greater of the most recent target cash incentive bonus or the average target cash incentive bonuses awarded for the prior years; or (iii) if he is terminated on or following the third anniversary, the amount will be equal to the greater of the most recent target cash incentive bonus or the average target bonus awarded for the prior three years. In addition, if, following a change in control, Mr. Clark is terminated without cause, or he terminates his employment for good reason, he will become vested in 24 months worth of his then unvested shares and options.

Steven A. Shallcross. We entered into an employment agreement in October 2005 with Mr. Shallcross, our Senior Vice President, Chief Financial Officer and Treasurer, which provides for an annual base salary of not less than \$250,000 and the possibility of an annual target cash incentive bonus equal to 25% of his annual base salary upon achievement of certain performance criteria. If Mr. Shallcross employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 12 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount determined as follows: (i) if he is terminated prior to the first anniversary of this agreement, a pro-rata portion of the anticipated first-year target cash incentive bonus will be given to him; (ii) if he is terminated on or following the first anniversary and prior to the third, the amount will equal the greater of the most recent target cash incentive bonus or the average target cash incentive bonuses awarded for the prior years; or (iii) if he is terminated on or following the third anniversary, the amount will be equal to the greater of the most recent target cash incentive bonus or the average target cash incentive bonus awarded for the prior three years. In addition, if, following a change in control, Mr. Shallcross is terminated without cause, or he terminates his employment for good reason, he will become vested in 24 months worth of his then unvested shares and options.

Thomas Copmann, Ph.D. We entered into an employment agreement in May 2005 with Dr. Copmann, our Vice President of Regulatory Affairs, which provides for an annual base salary of not less than \$200,000 and the possibility of an annual target cash incentive bonus equal to 28% of his annual base salary upon achievement of certain performance criteria. If Dr. Copmann s employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 6 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount equal to a pro-rata portion of the annual target cash incentive bonus for the year of his termination. In addition, if, following a change in control, Dr. Copmann is terminated without

cause, or he terminates his employment for good reason, he will become vested in 12 months worth of his then unvested shares and options.

Deepak Phadke, Ph.D. We entered into an offer letter in July 2005 with Dr. Phadke, our Vice President of Manufacturing, which provides for a sign-on bonus of \$20,000, \$10,000 of which was awarded in his first pay period and the remainder of which was awarded on the one year anniversary of his start date. We also entered into an employment agreement in August 2005 with Dr. Phadke, which provides for an annual base salary of not less than \$170,000 and the possibility of an annual target bonus equal to 15% of his annual base salary upon achievement of certain performance criteria. If Dr. Phadke s employment is terminated without cause, he becomes permanently disabled, or he terminates his employment for good reason, he will receive the following severance benefits following his employment termination: (a) a cash payment of his monthly base salary for 6 months; (b) payment of his monthly COBRA health insurance premiums; and (c) a bonus in an amount equal to a pro-rata portion of the annual target cash incentive bonus for the year of his termination. In addition, if, following a change in control, Dr. Phadke is terminated without cause, or he terminates his employment for good reason, he will become vested in 12 months worth of his then unvested shares and options.

#### Grants of plan-based awards

The following table sets forth each equity award granted to the Company s named executive officers during the year ended December 31, 2006.

	Grant	Estimated future  payouts under non-equity incentive plan awards	All other option awards: Number of securities underlying	Exercise or base price of option awards	Grant date fair value of stock option
Name	date	<b>Target</b> (\$)(1)	options	(\$/Sh)	awards(\$)(2)
Mihael H. Polymeropoulos, M.D. Paolo Baroldi, M.D., Ph.D.		\$150,696 30,735			
	07/06/06		60,427	\$ 8.30	\$ 348,368
	12/13/06		35,000	24.71	595,175
William D. Chip Clark		59,183			
Steven A. Shallcross		62,500			
Thomas Copmann, Ph.D.		58,240			
Deepak Phadke, Ph.D.		26,520			

- (1) This column sets forth the target amount of each executive s cash incentive bonus for 2006, as set forth in such executive s employment agreement and described in Compensation discussion & analysis and Employment agreements above.
- (2) Represents the fair value of each stock option as of the date it was granted, in accordance with SFAS 123(R) and using a Black-Scholes valuation model.

## Description of certain awards granted in 2006

On July 6, 2006, we granted an option to Dr. Paolo Baroldi to purchase a total of 60,427 shares of our common stock. The option vests with respect to 15,106 shares on July 6, 2007, provided Dr. Baroldi has remained employed with us through that date. The option then vests with respect to an additional 1,258 shares for each month after July 2007 that Dr. Baroldi remains employed with us.

On December 13, 2006, we granted an option to Dr. Baroldi to purchase a total of 35,000 shares of our common stock. The option vests with respect to 8,750 shares on December 13, 2007, provided Dr. Baroldi has remained employed with us through that date. The option then vests with respect to an additional 729 shares for each month after December 2007 that Dr. Baroldi remains employed with us.

# Outstanding equity awards at fiscal year-end

The following table sets forth information regarding each unexercised option held by each of our named executive officers as of December 31, 2006.

		Option awards				
	Number of securities underlying unexercised options(#)	Number of securities underlying unexercised options		Option exercise	Option expiration	
Name	exercisable	(#)unexercisable		price(\$)	date	
Mihael H. Polymeropoulos, M.D.		15,530(2)	\$	0.33	05/05/13	
• •	25,070	69,555(3)	\$	0.33	02/10/15	
	129,255	284,364(4)	\$	0.33	09/28/15	
	47,593	142,780(5)	\$	4.73	12/29/15	
Paolo Baroldi, M.D., Ph.D.		60,327(6)	\$	8.30	07/06/16	
		35,000(7)	\$	24.71	12/13/16	
William D. Chip Clark	17,562	40,106(8)	\$	0.33	09/01/04	
	22,156	26,185(9)	\$	0.33	02/10/15	
	64,231	141,310(10)	\$	0.33	09/28/15	
	9,976	29,931(11)	\$	4.73	12/29/15	
Steven A. Shallcross	12,502	60,585(12)	\$	0.83	11/14/15	
	16,995	50,985(13)	\$	4.73	12/29/15	
Thomas Copmann, Ph.D.(1)	3,172	9,517(14)	\$	4.73	12/29/15	
		5,000(15)	\$	25.50	12/18/16	
Deepak Phadke, Ph.D.	315	10,071(16)	\$	0.33	08/15/15	
	3,275	9,825(17)	\$	4.73	12/29/15	
		20,000(18)	\$	25.50	12/18/16	

<sup>(1)</sup> Does not include a total of 22,660 shares issued to Dr. Copmann upon the exercise of an option held by him. Of these 22,660 shares, 9,913 shares were vested to Dr. Copmann on December 31, 2006, and the remaining 12,747 shares were not yet vested to Dr. Copmann on December 31, 2006. 472 of these remaining shares vest each month after December, 2006, provided that Dr. Copmann remains employed with us.

<sup>(2)</sup> The option vests with respect to 3,105 additional shares each month after December, 2006, provided that Dr. Polymeropoulos remains employed with us.

<sup>(3)</sup> The option vests with respect to 2,675 additional shares each month after December, 2006, provided that Dr. Polymeropoulos remains employed with us.

- (4) The option vests with respect to 8,617 additional shares each month after December, 2006, provided that Dr. Polymeropoulos remains employed with us.
- (5) The option vests with respect to 3,966 additional shares each month after December, 2006, provided that Dr. Polymeropoulos remains employed with us.
- (6) The option vests with respect to 15,106 shares on July 6, 2007 and with respect to 1,256 additional shares for each month after July 2007, provided that Dr. Baroldi remains employed with us.
- (7) The option vests with respect to 8,750 shares on December 13, 2007 and with respect to 729 additional shares for each month after December 2007, provided that Dr. Baroldi remains employed with us.
- (8) The option vests with respect to 1,909 additional shares each month after December, 2006, provided that Mr. Clark remains employed with us.
- (9) The option vests with respect to 1,007 additional shares each month after December, 2006, provided that Mr. Clark remains employed with us.
- (10) The option vests with respect to 4,282 additional shares each month after December, 2006, provided that Mr. Clark remains employed with us.
- (11) The option vests with respect to 831 additional shares each month after December, 2006, provided that Mr. Clark remains employed with us.
- (12) The option vests with respect to 1,730 additional shares each month after December, 2006, provided that Mr. Shallcross remains employed with us.
- (13) The option vests with respect to 1,416 additional shares each month after December, 2006, provided that Mr. Shallcross remains employed with us.

- (14) The option vests with respect to 264 additional shares each month after December, 2006, provided that Dr. Copmann remains employed with us.
- (15) The option vests with respect to 1,250 shares on December 18, 2007 and with respect to 104 additional shares for each month after December 2007, provided that Dr. Copmann remains employed with us.
- (16) The option vests with respect to 314 additional shares each month after December, 2006, provided that Dr. Phadke remains employed with us.
- (17) The option vests with respect to 272 additional shares each month after December, 2006, provided that Dr. Phadke remains employed with us.
- (18) The option vests with respect to 5,000 shares on December 18, 2007 and with respect to 416 additional shares for each month after December 2007, provided that Dr. Phadke remains employed with us.

#### Option exercises and stock vested

The following table shows the number of shares acquired upon exercise of options by each named executive officer during the year ended December 31, 2006 and the number of shares of restricted stock held by each named executive officer that vested during the year ended December 31, 2006.

	Option awards Number of				
Name	shares acquired on exercise(#)	Value realized on exercise(\$)			
Mihael H. Polymeropoulos, M.D. Paolo Baroldi, M.D., Ph.D.	167,327	4,007,783			
William D. Chip Clark	34,000	598,602			
Steven A. Shallcross Thomas Copmann, Ph.D(1).	10,000	256,200			
Deepak Phadke, Ph.D.	4,720	116,772			

(1) Does not include 5,665 shares of restricted stock which were issued to Dr. Copmann upon the early exercise of an option owned by him and which vested during the year ended December 31, 2006.

#### **Option repricings**

The Company has not engaged in any option repricings or other modifications to any of its outstanding equity awards during fiscal year 2006.

## Severance and change in control arrangements

See Employment agreements above for a description of the severance and change in control arrangements for Drs. Polymeropoulos, Baroldi, Copmann and Phadke and Messrs. Clark and Shallcross. Drs. Polymeropoulos, Baroldi, Copmann and Phadke and Messrs. Clark and Shallcross will only be eligible to receive severance payments if each officer signs a general release of claims.

The compensation committee of our board of directors, as plan administrator of our Second Amended and Restated Management Equity Plan and our 2006 Equity Incentive Plan, has the authority to provide for accelerated vesting of the shares of common stock subject to outstanding options held by our named executive officers and any other person in connection with certain changes in control of Vanda.

In each employment agreement, a change in control is defined as (i) the consummation of a merger or consolidation of the Company with or into another entity, if persons who were not stockholders of the Company immediately prior to such merger or consolidation own immediately after such merger or consolidation 50% or more of the voting power of the outstanding securities of each of (A) the continuing or surviving entity and (B) any direct or indirect parent

corporation of such continuing or surviving entity; or (ii) the sale, transfer or other disposition of all or substantially all of the Company s assets. A transaction shall not constitute a change in control if its sole purpose is to change the state of the Company s incorporation or to create a holding company that will be owned in substantially the same proportions by the persons who held the Company s securities immediately before such transaction.

# Estimated payments and benefits upon termination

The amount of compensation and benefits payable to each named executive officer in various termination situations has been estimated in the tables below.

#### Mihael H. Polymeropoulos, M.D.

The following table describes the potential payments and benefits upon employment termination for Dr. Polymeropoulos, the Company s President and Chief Executive Officer, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

	Voluntary resignation		Voluntary	Tei	rminationTe	rmination		Involuntary termination in connection with or
Executive benefits and payments upon termination	not for good reason		resignation for good reason	by	company not for cause	by company for cause		following change in control
Compensation:		¢	276 740(1)	¢	276 740(1)		¢	276 740(1)
Base salary Highest target cash incentive bond Stock options	ıs	\$ \$	376,740(1) 181,000(2)		376,740(1) 181,000(2)		\$ \$	376,740(1) 181,000(2)
Unvested and accelerated							\$	11,829,202(3)
Benefits and perquisites:								
Health care		\$	23,398(4)	\$	23,398(4)		\$	23,398(4)
Accrued vacation pay		\$	5,161(5)	\$	5,161(5)		\$	5,161(5)
Total		\$	586,299	\$	586,299		\$	12,415,501

<sup>(1)</sup> Last monthly base salary prior to the termination for a period of 12 months following the date of the termination.

<sup>(2)</sup> Greater of the most recent target cash incentive bonus or the average of target cash incentive bonuses awarded for the prior years.

- (3) All options held by Dr. Polymeropoulos will become fully vested in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Dr. Polymeropoulos begins employment with another company that offers comparable benefits.
- (5) Based on 5 vacation days available to Dr. Polymeropoulos at December 31, 2006.

### Paolo Baroldi, M.D., Ph.D.

The following table describes the potential payments and benefits upon employment termination for Dr. Baroldi, the Company s Senior Vice President and Chief Medical Officer, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

Executive benefits and payments upon termination	Voluntary resignation not for good reason	Voluntary resignation for good reason	b	erminatioffe by company ot for cause	rmination by company for cause	Involuntary termination in connection with or owing change in control
Compensation:						
Base salary		\$ 250,000(1)	\$	250,000(1)		\$ 250,000(1)
Highest target cash incentive bonus Stock options	S	\$ 62,500(2)	\$	62,500(2)		\$ 62,500(2)
Unvested and accelerated						\$ 986,346(3)
Benefits and perquisites:						
Health care		\$ 23,398(4)	\$	23,398(4)		\$ 23,398(4)
Accrued vacation pay		\$ 2,397(5)		2,397(5)		\$ 2,397(5)
Total		\$ 338,295	\$	338,295		\$ 1,324,642

- (1) Last monthly base salary prior to the termination for a period of 12 months following the date of the termination.
- (2) Current target cash incentive bonus.
- (3) Acceleration of 24 months worth of Dr. Baroldi s then unvested options will occur in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Dr. Baroldi begins employment with another company that offers comparable benefits.
- (5) Based on 3.5 vacation days available to Dr. Baroldi at December 31, 2006.

# William D. Chip Clark

The following table describes the potential payments and benefits upon employment termination for Mr. Clark, the Company s Senior Vice President, Chief Business Officer and Secretary, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

Voluntary		Voluntary	T	erminatio <b>T</b> e			Involuntary termination in connection with or
resignation not for good reason		resignation for good reason			by company for cause	following change in control	
	\$	236,730(1)	\$	236,730(1)		\$	236,730(1)
us	\$	62,600(2)	\$	62,600(2)		\$	62,600(2)
						\$	4,460,057(3)
	\$	23,398(4)	\$	23,398(4)		\$	23,398(4)
	\$	1,946(5)	\$	1,946(5)		\$	1,946(5)
	\$	324.674	\$	324.674		\$	4,784,731
	resignation not for good	resignation not for good reason  \$ us \$	resignation not for good reason reason resignation for good reason \$ 236,730(1) \$ 62,600(2) \$ \$ 23,398(4) \$ 1,946(5)	resignation not for good for good reason	resignation not for good reason reason reason by company not for good reason not for cause  \$ 236,730(1) \$ 236,730(1) \$ 62,600(2) \$ 62,600(2)  \$ 23,398(4) \$ 23,398(4) \$ 1,946(5) \$ 1,946(5)	resignation not for good reason reason by company company for not for cause \$ 236,730(1) \$ 236,730(1) \$ 62,600(2) \$ 62,600(2) \$ 1,946(5) \$ 1,946(5)	resignation not for good for good reason reason not for cause cause  \$ 236,730(1) \$ 236,730(1) \$ 30,000(2) \$ 30,00

- (1) Last monthly base salary prior to the termination for a period of 12 months following the date of the termination.
- (2) Greater of the most recent target cash incentive bonus or the average of target cash incentive bonuses awarded for the prior years.
- (3) Acceleration of 24 months worth of Mr. Clark s then unvested options will occur in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Mr. Clark begins employment with another company that offers comparable benefits.
- (5) Based on 3 vacation days available to Mr. Clark at December 31, 2006.

#### Steven A. Shallcross

The following table describes the potential payments and benefits upon employment termination for Mr. Shallcross, the Company s Senior Vice President, Chief Financial Officer and Treasurer, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

Voluntary		Voluntary	Te	rminationTe	rmination		Involuntary termination in connection with or
not for good reason	]	resignation for good reason	by	company not for cause	by company for cause	change	following change in control
	\$	250,000(1)	\$	250,000(1)		\$	250,000(1)
ıs	\$		\$			\$	62,500(2)
						\$	2,458,756(3)
	\$	23,398(4)	\$	23,398(4)		\$	23,398(4)
	\$	3,425(5)	\$	3,425(5)		\$	3,425(5)
	\$	339,323	\$	339,323		\$	2,789,079
	resignation not for good	resignation not for good reason  \$	resignation not for resignation good for good reason	resignation not for resignation by good for good reason so	resignation not for resignation by company good for good reason reason cause  \$ 250,000(1) \$ 250,000(1) \$ 62,500(2) \$ 62,500(2) \$ 62,500(2) \$ 3,425(5) \$ 3,425(5)	resignation not for resignation by company company good for good reason reason solution solution (a) \$ 250,000(1) \$ 250,000(1) \$ 62,500(2) \$ 62,500(2) \$ 3,425(5) \$ 3,425(5)	Voluntary resignation         Voluntary not for resignation good reason         TerminationTermination           \$ 250,000(1)         \$ 250,000(1)         \$ 250,000(1)         \$ 3,425(5) </td

- (1) Last monthly base salary prior to the termination for a period of 12 months following the date of the termination.
- (2) Greater of the most recent target cash incentive bonus or the average of target cash incentive bonuses awarded for the prior years.
- (3) Acceleration of 24 months worth of Mr. Shallcross then unvested options will occur in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Mr. Shallcross begins employment with another company that offers comparable benefits.
- (5) Based on 5 vacation days available to Mr. Shallcross at December 31, 2006.

# Thomas Copmann, Ph.D.

The following table describes the potential payments and benefits upon employment termination for Dr. Copmann, the Company s Vice President of Regulatory Affairs, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

	Voluntary		•				Involuntary termination n connection with or
Executive benefits and payments upon termination	resignation not for good reason		resignation for good reason	by	company not for cause	by company for cause	following change in control
Compensation:							
Base salary		\$	104,000(1)	\$	104,000(1)		\$ 104,000(1)
Target cash incentive bonus Stock options		\$	58,240(2)	\$	58,240(2)		\$ 58,240(2)
Unvested and accelerated							\$ 200,964(3)
Benefits and perquisites:							
Health care		\$	23,398(4)	\$	23,398(4)		\$ 23,398(4)
Accrued vacation pay		\$	2,849(5)	\$	2,849(5)		\$ 2,849(5)
Total		\$	188,487	\$	188,487		\$ 389,451

- (1) Last monthly base salary prior to the termination for a period of 6 months following the date of the termination.
- (2) 2006 target bonus.
- (3) Acceleration of 12 months worth of Dr. Copmann s then unvested options will occur in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Dr. Copmann begins employment with another company that offers comparable benefits.
- (5) Based on 5 vacation days available to Dr. Copmann at December 31, 2006.

# Deepak Phadke, Ph.D.

The following table describes the potential payments and benefits upon employment termination for Dr. Phadke, the Company s Vice President of Manufacturing, as if his employment terminated as of December 29, 2006, the last business day of our last fiscal year.

	Voluntary		· ·				Involuntary termination n connection with or
Executive benefits and payments upon termination	resignation not for good reason		resignation for good reason	by	company not for cause	by company for cause	following change in control
Compensation:							
Base salary		\$	88,400(1)	\$	88,400(1)		\$ 88,400(1)
Target cash incentive bonus Stock options		\$	26,520(2)	\$	26,520(2)		\$ 26,520(2)
Unvested and accelerated							\$ 157,082(3)
Benefits and perquisites:							
Health care		\$	23,398(4)	\$	23,398(4)		\$ 23,398(4)
Accrued vacation pay		\$	2,422(5)	\$	2,422(5)		\$ 2,422(5)
Total		\$	140,740	\$	140,740		\$ 297,822

- (1) Last monthly base salary prior to the termination for a period of 6 months following the date of the termination.
- (2) 2006 target bonus.
- (3) Acceleration of 12 months worth of Dr. Phadke s then unvested options will occur in the event of an involuntary termination following a change of control.
- (4) Payment of the COBRA health insurance premiums up to 18 months or until Dr. Phadke begins employment with another company that offers comparable benefits.
- (5) Based on 5 vacation days available to Dr. Phadke at December 31, 2006.

The value of the option vesting acceleration was calculated for each of the above tables based on the assumption that the change in control and the executive s employment termination occurred on December 29, 2006. The closing price of the Company s stock as of December 29, 2006, was \$24.65, which was used as the value of the Company s stock in the change in control. The value of the vesting acceleration was calculated by multiplying the number of unvested option shares subject to vesting acceleration as of December 29, 2006 by the difference between the closing price of the Company s stock as of December 29, 2006 and the exercise price for such unvested option shares.

### **Equity benefit plans**

#### Second Amended and Restated Management Equity Plan

Share reserve. Our Second Amended and Restated Management Equity Plan was adopted by us in December 2004. As of December 31, 2006, options for a total of 318,158 shares were exercised under the plan, and options for a total of 1,347,205 shares remained outstanding. No further option grants were made under this plan after the effective date of our initial public offering. The options that were outstanding under the plan after the effective date of our initial public offering continue to be governed by their terms. After the effective date of our initial public offering, any shares that remained available for grants under the plan and any shares subject to options or share awards under the plan that are canceled, forfeited or repurchased will not be available for future grants or awards. The plan is administered by our compensation committee.

*Eligibility*. Employees, non-employee members of our board of directors and consultants are eligible to participate in our Second Amended and Restated Management Equity Plan.

Types of awards. Our Second Amended and Restated Management Equity Plan provides for the purchase of shares of our common stock, and incentive and nonstatutory stock options to purchase shares of our common stock. The exercise price for incentive stock options and nonstatutory stock options granted under the plan may not be less than 100% and 30%, respectively, of the fair market value of our common stock on the option grant date. Optionees may pay the purchase price or the exercise price by using cash, shares of common stock that the optionee already owns, a full-recourse promissory note, by rendering services to us, by an immediate sale of the option shares through a broker designated by us, or with a loan from a broker designated by us and secured by the option shares. In most cases, our options vest over a four-year period following the date of grant and generally expire 10 years after they are granted, unless the optionee separates from service with us.

*Change in control.* If we merge or consolidate with another company, an option granted under the Second Amended and Restated Management Equity Plan will be subject to the terms of the merger or consolidation agreement, which may provide that the option continues, is assumed or substituted, becomes vested and exercisable in full, or is canceled and the optionees receive a payment.

Amendments or termination. Our board of directors may amend or terminate the Second Amended and Restated Management Equity Plan at any time. If our board amends the plan, it does not need to seek stockholder approval of the amendment unless the number of shares reserved under the plan increases or the class of person eligible for the grant of incentive stock options materially changes. The plan will automatically terminate 10 years after its adoption by our board of directors.

### 2006 Equity Incentive Plan

Our 2006 Equity Incentive Plan was adopted in March 2006 and took effect as of the closing of the initial public offering with the following material terms:

Share reserve. We have reserved 1,500,000 shares of our common stock for issuance under the 2006 Equity Incentive Plan. As of December 31, 2006, awards for a total of 359,527 shares were outstanding under the plan. On January 1 of each year, starting with the year 2007, the number of shares in the reserve will automatically increase by 4% of the total number of shares of common stock that are outstanding at that time or, if less, by 1,500,000 shares (or such lesser number as may be approved by the board of directors). If options or shares awarded under the 2006 Equity Incentive Plan are forfeited, then those options or shares will again become available for awards under this plan.

Administration. The compensation committee of our board of directors administers the 2006 Equity Incentive Plan. The committee has complete discretion to make all decisions relating to the interpretation and operation of the plan, including the discretion to determine who will receive an award, what type of award it will be, how many shares will be covered by the award, what the vesting requirements will be, if any, and what the other features and conditions of each award will be. The compensation committee has the ability to reprice outstanding options and modify outstanding awards in other ways.

Eligibility. The following groups of individuals are eligible to participate in the 2006 Equity Incentive Plan:

employees

members of our board of directors who are not employees

consultants

Types of awards. The 2006 Equity Incentive Plan provides for the following types of awards:

options to purchase shares of our common stock

stock appreciation rights

restricted shares of our common stock

stock units (sometimes called phantom shares)

Options and stock appreciation rights. Both incentive stock options and nonstatutory stock options are available for grant under the 2006 Equity Incentive Plan. An optionee who exercises an incentive stock option may qualify for favorable tax treatment under section 422 of the Internal Revenue Code of 1986. On the other hand, nonstatutory stock options do not qualify for such favorable tax treatment. The exercise price of options and stock appreciation rights granted under the 2006 Equity Incentive Plan may not be less than 100% of the fair market value of our common stock on the grant date. Optionees may pay the exercise price by using:

cash

shares of common stock that the optionee already owns

an immediate sale of the option shares through a broker designated by us

a full-recourse promissory note

Options and stock appreciation rights vest at the time or times determined by the compensation committee. In most cases, our options vest over the four-year period following the date of grant. Options and stock appreciation rights generally expire 10 years after they are granted, except that they generally expire earlier if the optionees service terminates earlier. The 2006 Equity Incentive Plan provides that no participant may receive options covering more than 500,000 shares and stock appreciation rights covering more than 500,000 shares in the same year, except that a newly hired employee may receive options covering up to 1,000,000 shares and stock appreciation rights covering up to 1,000,000 shares in the first year of employment.

The 2006 Equity Incentive Plan also provides for automatic annual option grants to members of our board of directors who are not our employees. See Director compensation.

Restricted shares and stock units. Restricted shares may be awarded under the 2006 Equity Incentive Plan in return for:

cash

a full-recourse promissory note; or

services

Restricted shares and stock units vest at the time or times determined by the compensation committee.

Change in control. If a change in control of Vanda occurs, an award under the 2006 Equity Incentive Plan will vest on an accelerated basis to the extent determined by the compensation committee. The compensation committee may determine that outstanding grants will vest in full or in part at the time of the change in control. It may also determine that the grants will

vest on an accelerated basis only if the participant is actually or constructively discharged within a specified period of time after the change in control. Finally, the committee has the discretion to determine that the grants will remain outstanding without acceleration of vesting, except that if the surviving corporation fails to assume an outstanding award or replace it with a comparable award or cash payment, then the award will always become fully vested as a result of the change in control. A change in control includes the following events for purposes of the 2006 Equity Incentive Plan:

- a merger of Vanda after which our own stockholders own 50% or less of the surviving corporation
- a sale of all or substantially all of our assets
- a proxy contest that results in the replacement of 50% or more of our directors over a 24-month period

an acquisition of 50% or more of our outstanding stock by any person or group, other than a person related to Vanda (such as a holding company owned by our stockholders)

Amendments or termination. Our board of directors may amend or terminate the 2006 Equity Incentive Plan at any time. If our board were to amend the plan, it would not need to ask for stockholder approval of the amendment unless applicable law requires it. The 2006 Equity Incentive Plan will automatically terminate on April 12, 2016.

#### 401(k) plan

We have established a 401(k) plan to allow our employees to save on a tax-favorable basis for their retirements. We match up to 50% of the first 6% of contributions made by employees pursuant to the plan.

# Limitation of liability and indemnification of officers and directors

Our amended and restated certificate of incorporation and amended and restated bylaws provide that we will indemnify our directors and officers to the fullest extent permitted by Delaware law, as it now exists or may in the future be amended, against all expenses and liabilities reasonably incurred in connection with their service for or on behalf of us. In addition, our amended and restated certificate of incorporation provides that our directors will not be personally liable for monetary damages to us for breaches of their fiduciary duty as directors, unless they violated their duty of loyalty to us or our stockholders, acted in bad faith, knowingly or intentionally violated the law, authorized illegal dividends or redemptions or derived an improper personal benefit from their actions as directors. We maintain liability insurance which insures our directors and officers against certain losses and which insures us against our obligations to indemnify our directors and officers.

In addition, we have entered into indemnification agreements with each of our directors and officers. These agreements, among other things, require us to indemnify each director and officer to the fullest extent permitted by Delaware law, including indemnification of expenses such as attorneys fees, judgments, fines and settlement amounts incurred by the director or officer in any action or proceeding, including any action or proceeding by or in right of us, arising out of the person s services as a director or officer. At present, we are not aware of any pending or threatened litigation or proceeding involving any of our directors, officers, employees or agents in which indemnification would be required or permitted. We believe provisions in our amended and restated certificate of incorporation and indemnification agreements are necessary to attract and retain qualified persons as directors and officers.

## Certain relationships and related party transactions

# 2004 Securityholder Agreement

We have entered into a 2004 Securityholder Agreement with certain holders of our common stock, including significant holders that are affiliates of certain of our directors. Under the Securityholders Agreement, these holders have the right to demand the registration of our common stock and to participate in other public offerings of our common stock (for more information regarding the registration rights granted pursuant to the 2004 Securityholder Agreement, see Description of capital stock Registration rights )

### **Indemnification agreements**

We have entered into indemnification agreements with each of our directors and officers. These agreements, among other things, require us to indemnify each director and officer to the fullest extent permitted by Delaware law, including indemnification of expenses such as attorneys fees, judgments, fines and settlement amounts incurred by the director or officer in any action or proceeding, including any action or proceeding by or in right of us, arising out of the person s services as a director or officer.

# Relationship with Care Capital, LLC

From time to time, we reimbursed Care Capital, LLC ( Care Capital ), an affiliate of our stockholders, Care Capital Investments II, LP and Care Capital Offshore Investments II, LP, for certain expenses incurred by Care Capital on our behalf. We reimbursed Care Capital for approximately \$54,000 and approximately \$299,000 for the year ended December 31, 2004 and for the period from March 13, 2003 (inception) to December 31, 2003, respectively.

We also used the services of a Care Capital employee and reimbursed Care Capital for such personnel services related to occupancy and salary expenses incurred on our behalf. Reimbursements related to such expenses were approximately \$49,000 and \$34,000 for the year ended December 31, 2004 and the period from March 13, 2003 (inception) to December 31, 2003, respectively.

# **Principal stockholders**

The following table sets forth certain information known to us regarding beneficial ownership of our common stock as of December 31, 2006 and as adjusted to reflect the sale of the shares of common stock in this offering by:

each person known by us to be the beneficial owner of more than 5% of our common stock our named executive officers each of our directors all executive officers and directors as a group

Unless otherwise indicated in the footnotes, to our knowledge, each stockholder possesses sole voting and investment power over the shares listed, except for shares owned jointly with that person s spouse.

Beneficial ownership is determined in accordance with the rules of the Securities and Exchange Commission and generally includes voting or investment power with respect to securities. Options and warrants to purchase shares of our common stock that are exercisable within 60 days of December 31, 2006, are deemed to be beneficially owned by the persons holding these options for the purpose of computing percentage ownership of that person, but are not treated as outstanding for the purpose of computing any other person s ownership percentage.

Percentage of shares beneficially owned before the offering is based on 22,128,534 shares of common stock outstanding as of December 31, 2006. Percentage of shares beneficially owned after the offering is based on 25,628,534 shares of common stock as of December 31, 2006, after giving effect to the sale of 3,500,000 shares in this offering.

		Percentage of share beneficially owne			
Name and address of beneficial owner(1)	Number of shares beneficially owned	Before offering	After offering		
5% Stockholders					
Domain Partners VI, L.P.(2)	1,995,976	9.02%	7.79%		
One Palmer Square, Suite 515					
Princeton, NJ 08542					
Care Capital Investments II, LP(3)	1,893,730	8.56%	7.39%		
47 Hulfish St., Ste 310					
Princeton, NJ 08542					
Versant Capital Management LLC(4)	1,500,000	6.78%	5.85%		
45 Rockefeller Plaza					
Suite 2074					
New York, NY 10111					
Steven A. Cohen(5)	1,448,332	6.55%	5.65%		
72 Cummings Point Road					
Stanford, CT 06902					
Davidson Kempner Partners(6)	1,401,000	6.33%	5.46%		
65 East 55th Street, 19th Floor					
New York, NY 10022					
Executive Officers and Directors					
Mihael H. Polymeropoulos, M.D.(7)	238,467	1.07%	*		
Paolo Baroldi, M.D., Ph.D.(8)					
William D. Chip Clark(9)	131,896	*	*		
Steven A. Shallcross(10)	35,791	*	*		
Argeris N. Karabelas, Ph.D.(11)	1,893,730	8.56%	7.39%		
Richard W. Dugan(12)	3,083	*	*		
Brian K. Halak, Ph.D.					
H. Thomas Watkins(13)	3,645	*	*		
David Ramsay(14)	1,893,730	8.56%	7.39%		
James B. Tananbaum, M.D.(15)	736,077	3.33%	2.87%		
Deepak Phadke, Ph.D.(16)	4,764	*	*		
Thomas Copmann, Ph.D.(17)	26,360	*	*		
All executive officers and directors as a group	3,073,993	13.62%	11.79%		

<sup>\*</sup> Represents beneficial ownership of less than one percent of our outstanding common stock.

- (1) Unless otherwise indicated, the address for each beneficial owner is c/o Vanda Pharmaceuticals Inc., 9605 Medical Center Drive, Suite 300, Rockville, Maryland 20850.
- (2) Includes 1,954,450 shares held of record by Domain Partners VI, L.P., 21,068 shares held of record by DP VI Associates, L.P. and 20,458 shares held of record by One Palmer Square Associates: VI, L.L.C. Voting and/or dispositive decisions with respect to the shares held by Domain Partners VI, L.P. and DP VI Associates, L.P. are made by the managing members of their general partner, One Palmer Square Associates VI, L.L.C.: James C. Blair, Ph.D., Brian H. Dovey, Robert J. More, Kathleen K. Schoemaker, Jesse I. Treu, Ph.D. and Nicole Vitullo,

- each of whom disclaims beneficial ownership of such shares except to the extent of his or her pecuniary interest therein, the amount of which cannot currently be determined.
- (3) Includes 1,772,184 shares held of record by Care Capital Investments II, LP and 121,546 shares held of record by Care Capital Offshore Investments II, LP. Voting and/or dispositive decisions with respect to the shares held by Care Capital Investments II, LP and Care Capital Offshore Investments II, LP are made by the managing members of their general partner, Care Capital II, LLC: Jan Leschly, Argeris N. Karabelas, Ph.D. and David R. Ramsay, each of whom disclaims beneficial ownership of such shares except to the extent of his pecuniary interest therein, the amount of which cannot currently be determined.
- (4) Based on Schedule 13G filed by Versant Capital Management LLC and Herriot Tabuteau on November 17, 2006. Harriot Tabuteau is the managing member of Versant Capital Management LLC and so may be deemed to beneficially own such shares of Common Stock. Mr. Tabuteau disclaims such beneficial ownership.

- (5) Based on Schedule 13G filed jointly on December 21, 2006 by (i) S.A.C. Capital Advisors, LLC, (SAC Capital Advisors ), beneficially owned by S.A.C. Capital Associates, LLC ( SAC Capital Associates ) and S.A.C. MultiQuant Fund, LLC ( SAC MultiQuant ), (ii) S.A.C. Capital Management, LLC, ( SAC Capital Management ) with respect to shares beneficially owned by SAC Capital Associates and SAC MultiQuant, (iii) SAC Capital Associates with respect to shares beneficially owned by it, (iv) Sigma Capital Management, LLC ( Sigma Management ) with respect to shares beneficially owned by Sigma Capital Associates, LLC ( Sigma Capital Associates ), and (v) Steven A. Cohen with respect to shares beneficially owned by SAC Capital Advisors, SAC Capital Management, SAC Capital Associates, SAC MultiQuant, Sigma Management and Sigma Capital Associates. SAC Capital Advisors, SAC Capital Management, Sigma Management and Mr. Cohen do not directly own any shares. Pursuant to investment agreements, each of SAC Capital Advisors and SAC Capital Management share all investment and voting power with respect to the securities held by SAC Capital Associates and SAC MultiQuant. Pursuant to an investment management agreement, Sigma Management maintains investment and voting power with respect to the securities held by Sigma Capital Associates. Mr. Cohen controls each of SAC Capital Advisors, SAC Capital Management and Sigma Management. By reason of the provisions of Rule 13d-3 of the Securities Exchange Act of 1934, as amended, each of (i) SAC Capital Advisors, SAC Capital Management and Mr. Cohen may be deemed to own beneficially 1,348,332 shares; and (ii) Sigma Management and Mr. Cohen may be deemed to own beneficially an additional 100,000 shares. Each of SAC Capital Advisors, SAC Capital Management, Sigma Management and Mr. Cohen disclaim beneficial ownership of any shares.
- (6) Based on Schedule 13G filed jointly on December 11, 2006 by Davidson Kempner Partners, a New York limited partnership ( DKP ), Davidson Kempner Institutional Partners, L.P., a Delaware limited partnership ( DKIP ), M. H. Davidson & Co., a New York limited partnership (CO), Davidson Kempner International, Ltd., a British Virgin Islands corporation ( DKIL ), Serena Limited, a Cayman Islands corporation ( Serena ), Davidson Kempner Healthcare Fund LP, a Delaware limited partnership (DKHF), Davidson Kempner Healthcare International Ltd., a Cayman Islands corporation ( DKHI ), MHD Management Co., a New York limited partnership and the general partner of DKP (MHD), Davidson Kempner Advisers Inc., a New York corporation and the general partner of DKIP (DKAI), Davidson Kempner International Advisors, L.L.C., a Delaware limited liability company and the manager of DKIL and Serena (DKIA), DK Group LLC, a Delaware limited liability company and the general partner of DKHF ( DKG ), DK Management Partners LP, a Delaware limited partnership and the investment manager of DKHI ( DKMP ), DK Stillwater GP LLC, a Delaware limited liability company and the general partner of DKMP ( DKS ), and Messrs. Thomas L. Kempner, Jr., Marvin H. Davidson, Stephen M. Dowicz, Scott E. Davidson, Michael J. Leffell, Timothy I. Levart, Robert J. Brivio, Jr., Anthony A. Yoseloff, Eric P. Epstein and Avram Z. Friedman (collectively, the Principals ), who are the general partners of CO and MHD, the sole managing members of DKIA and DKG and the sole stockholders of DKAI. Messrs. Thomas L. Kempner, Jr. and Timothy I. Levart are Executive Managing Member and Deputy Executive Managing Member, respectively, of DKS. Each of Messrs. Kempner and Levart, together with Messrs. Marvin H. Davidson, Stephen M. Dowicz, Scott E. Davidson, Michael J. Leffell, Robert J. Brivio, Jr., Anthony A. Yoseloff, Eric P. Epstein and Avram Z. Friedman are limited partners of DKMP. The Principals may be deemed to beneficially own an aggregate of 1,401,100 shares as a result of their voting and dispositive power over the 1,401,100 shares beneficially owned by DKP, DKIP, DKIL, Serena, CO, DKHF and DKHI. DKIA may be deemed to beneficially own the 226,978 shares beneficially owned by DKIL and the 5,604 shares beneficially owned by Serena as a result of its voting and dispositive power over those shares. DKAI may be deemed to beneficially own the 141,513 shares beneficially owned by DKIP as a result of its voting and dispositive power over those shares. MHD may be deemed to beneficially own the 86,868 shares beneficially owned by DKP as a result of its voting and dispositive power over those shares. DKG may be deemed to beneficially own the 479,175 shares beneficially owned by DKHF as a result of its voting and dispositive power over those shares. DKMP and DKS may be deemed to beneficially own the 451,154 shares beneficially owned by DKHI as a result of their voting and dispositive

power over those shares.

- (7) Excludes 475,501 shares unexercisable within 60 days of December 31, 2006.
- (8) Excludes 95,427 shares unexercisable within 60 days of December 31, 2006
- (9) Excludes 219,561 shares unexercisable within 60 days of December 31, 2006.
- (10) Excludes 105,276 shares unexercisable within 60 days of December 31, 2006.
- (11) Includes 1,772,184 shares held of record by Care Capital Investments II, LP and 121,546 shares of record held by Care Capital Offshore Investments II, LP. Dr. Karabelas is a managing member of Care Capital II, LLC. Care Capital II, LLC is the general partner of Care Capital Investments II, LP and Care Capital Offshore Investments II, LP. Dr. Karabelas disclaims beneficial ownership of the shares held by Care Capital Investments II, LP and Care Capital Offshore Investments II, LP except to the extent of his pecuniary interest therein, the amount of which cannot currently be determined.
- (12) Excludes 7,488 shares unexercisable within 60 days of December 31, 2006.
- (13) Excludes 31,355 shares unexercisable within 60 days of December 31, 2006.
- (14) Includes 1,772,184 shares held of record by Care Capital Investments II, LP and 121,546 shares held of record held by Care Capital Offshore Investments II, LP. Mr. Ramsay is a Partner of Care Capital, LLC. Care Capital, LLC is the general partner of Care Capital Investments II, LP and Care Capital Offshore Investments II, LP. Mr. Ramsay disclaims beneficial ownership of the shares held by Care Capital Investments II, LP and Care Capital Offshore Investments II, LP except to the extent of his pecuniary interest therein, the amount of which cannot currently be determined.
- (15) Includes 725,036 shares held of record by Prospect Venture Partners II, L.P. and 11,041 shares held of record by Prospect Associates II, L.P. Dr. Tananbaum serves as a managing member of Prospect Management Co. II, L.L.C., the general partner of Prospect Venture Partners II, L.P. and Prospect Associates II, L.P. He disclaims beneficial ownership of the shares held of record by Prospect Venture Partners II, L.P. and Prospect Associates II, L.P. except to the extent of his pecuniary interest therein the amount of which cannot currently be determined. Also includes 2,303 shares held directly by Dr. Tananbaum.
- (16) Excludes 38,722 shares unexercisable within 60 days of December 31, 2006.
- (17) Excludes 13,989 shares unexercisable within 60 days of December 31, 2006. Includes 12,747 restricted shares which are subject to vesting restrictions as of December 31, 2006.

## **Description of capital stock**

#### General

The following is a summary of the rights of our common stock and related provisions of our restated certificate of incorporation and bylaws as in effect upon the closing of this offering. For more detailed information, please see our Amended and Restated Certificate of Incorporation, Amended and Restated Bylaws, and our 2004 Securityholder Agreement, which are filed as exhibits to the Registration Statement on Form S-1 for our initial public offering and which are each incorporated by reference in this prospectus.

Our authorized capital stock consists of 150,000,000 shares of common stock and 20,000,000 shares of undesignated preferred stock, each with a par value of \$0.001 per share.

As of December 31, 2006, we had outstanding 22,128,534 shares of common stock. In addition, as of December 31, 2006, a total of 1,706,732 shares of our common stock were subject to outstanding options. At December 31, 2006, a total of 91,533 shares of our outstanding common stock were held by our employees and consultants (not including shares underlying outstanding options). 28,610 of these shares are subject to a lapsing right of repurchase in our favor, under which we may repurchase these shares upon the termination of the holder s employment or consulting relationship.

#### Common stock

### **Voting rights**

Unless otherwise provided for in our amended and restated certificate of incorporation or required by applicable law, on all matters submitted to our stockholders for vote, our common stockholders are entitled to one vote per share, voting together as a single class.

### **Dividends**

Our amended and restated certificate of incorporation provides that the holders of common stock shall be entitled to share equally in any dividends that our board of directors may determine to issue from time to time. We do not currently expect to pay dividends.

# Liquidation

Our amended and restated certificate of incorporation provides that upon our liquidation, dissolution or winding-up, the holders of common stock are entitled to share equally in all assets remaining after the payment of any liabilities and the liquidation preferences on any outstanding preferred stock.

### Anti-takeover effects of our amended and restated certificate of incorporation, bylaws and Delaware law

Some provisions of Delaware law and our amended and restated certificate of incorporation and bylaws could make the following transactions more difficult:

our acquisition by means of a tender offer

our acquisition by means of a proxy contest or otherwise

removal of our incumbent officers and directors

These provisions, summarized below, are expected to discourage and prevent coercive takeover practices and inadequate takeover bids. These provisions are designed to encourage persons seeking to acquire control of us to first negotiate with our board of directors, and also are

intended to provide management with flexibility to enhance the likelihood of continuity and stability in our composition if our board of directors determines that a takeover is not in our best interests or the best interests of our stockholders. These provisions, however, could have the effect of discouraging attempts to acquire us, which could deprive our stockholders of opportunities to sell their shares of common stock at prices higher than prevailing market prices. We believe that the benefits of these provisions, including increased protection of our potential ability to negotiate with the proponent of an unfriendly or unsolicited proposal to acquire or restructure us, outweigh the disadvantages of discouraging takeover proposals because negotiation of takeover proposals could result in an improvement of their terms.

Election and removal of directors. Our board of directors is divided into three classes serving staggered three-year terms. This system of electing directors may tend to discourage a third party from making a tender offer or otherwise attempting to obtain control of us because generally at least two stockholders meetings will be required for stockholders to effect a change in control of the board of directors. Our amended and restated certificate of incorporation and our bylaws contain provisions that establish specific procedures for appointing and removing members of the board of directors. Under our amended and restated certificate of incorporation, vacancies and newly created directorships on the board of directors may be filled only by a majority of the directors then serving on the board, and under our bylaws, directors may be removed by the stockholders only for cause.

*Stockholder meetings*. Under our amended and restated bylaws, only the board of directors, the Chairman of the board or our Chief Executive Officer may call special meetings of stockholders.

Requirements for advance notification of stockholder nominations and proposals. Our amended and restated bylaws contain advance notice procedures with respect to stockholder proposals and the nomination of candidates for election as directors, other than nominations made by or at the direction of the board of directors or a committee of the board of directors.

Delaware anti-takeover law. We are subject to Section 203 of the Delaware General Corporation Law, an anti-takeover law. In general, Section 203 prohibits a publicly held Delaware corporation from engaging in a business combination with an interested stockholder for a period of three years following the date the person became an interested stockholder, unless the business combination or the transaction in which the person became an interested stockholder is approved in a prescribed manner. Generally, a business combination includes a merger, asset or stock sale, or another transaction resulting in a financial benefit to the interested stockholder. Generally, an interested stockholder is a person who, together with affiliates and associates, owns, or within three years prior to the date of determination of interested stockholder status did own, 15% or more of the corporation s voting stock. The existence of this provision may have an anti-takeover effect with respect to transactions that are not approved in advance by our board of directors, including discouraging attempts that might result in a premium over the market price for the shares of common stock held by stockholders.

*No stockholder action by written consent.* Our amended and restated certificate of incorporation prohibits stockholders from taking action without a meeting.

*No cumulative voting.* Our amended and restated certificate of incorporation and bylaws do not provide for cumulative voting in the election of directors. Cumulative voting allows a minority stockholder to vote a portion or all of its shares for one or more candidates for seats on the board of directors. Without cumulative voting, a minority stockholder will not be able to gain as many seats on our board of directors based on the number of shares of our stock the stockholder holds as the stockholder would be able to gain if cumulative voting were permitted.

The absence of cumulative voting makes it more difficult for a minority stockholder to gain a seat on our board of directors to influence our board s decision regarding a takeover.

*Undesignated preferred stock*. The authorization of undesignated preferred stock in our amended and restated certificate of incorporation makes it possible for our board of directors to issue preferred stock with voting or other rights or preferences that could impede the success of any attempt to acquire us.

These and other provisions may have the effect of deferring hostile takeovers or delaying changes in control or management.

#### Limitation of liability of directors

To the fullest extent permitted by the Delaware General Corporation Law as it now exists or hereafter may be amended, our directors will not be personally liable to us or our stockholders for monetary damages for breach of fiduciary duty as a director.

If the Delaware General Corporation Law is later amended to authorize the further elimination or limitation of the liability of directors, then the liability of our directors, in addition to the limitation on personal liability provided in our certificate of incorporation, will be limited to the fullest extent permitted by the amended Delaware General Corporation Law. Any repeal or modification of the provisions in our certificate of incorporation by our stockholders relating to the limitation of the liability of our directors will be prospective only and will not adversely affect any limitation on the personal liability of our directors existing at the time of the repeal or modification.

#### **Registration rights**

As of December 31, 2006, the holders of 6,103,579 shares of our common stock are entitled to rights with respect to the registration of their shares under the Securities Act. These registration rights are contained in our 2004 Securityholder Agreement and are described below. These registration rights will expire on April 18, 2011.

# **Demand registration rights**

Stockholders holding at least 25% of the total shares of common stock held by our venture capital investors as of our initial public offering may demand the registration of such shares under our 2004 Securityholder Agreement. We are only obligated to effect two registrations in response to these demand registration rights, and we are not obligated to effect any demand registration for shares having an aggregate market value of less than \$5,000,000 as of the date notice is given to us to effect such a registration. We may also postpone the filing of a registration statement for up to 90 days once in any 12-month period if our board of directors determines in good faith that the filing would be significantly disadvantageous to us and our affiliates, taken as a whole. We must pay all expenses incurred in connection with demand registration rights.

# **Incidental registration rights**

Stockholders with incidental registration rights under our 2004 Securityholder Agreement have the right to include their shares in any registration of shares to be issued by the Company, subject to specified exceptions. The underwriters of any underwritten offering have the right to limit the number of shares registered by these stockholders due to marketing reasons. We must pay all expenses incurred in connection with these incidental registration rights. These incidental registration rights have been waived with respect to this offering.

### S-3 registration rights

If we are eligible to file a registration statement on Form S-3, the stockholders with S-3 registration rights under the 2004 Securityholder Agreement can request that we register their shares, provided that the total price of the shares of common stock offered to the public is at least \$1,000,000 (before deduction of underwriting discounts and commissions). The holders of S-3 registration rights may only require us to file one Form S-3 registration statement in any 12-month period. We may postpone the filing of a Form S-3 registration statement for up to 90 days once in any 12-month period if our board of directors determines in good faith that the filing would be seriously detrimental to us.

# Transfer agent and registrar

The transfer agent and registrar for our common stock and the rights is American Stock Transfer and Trust Company.

## Material United States federal tax consequences

The following is a general discussion of the material U.S. federal income and estate tax consequences of the ownership and disposition of common stock by a beneficial owner that is a non-U.S. holder , other than a non-U.S. holder that owns, or has owned, actually or constructively, more than 5% of the company s common stock. A non-U.S. holder is a person or entity that, for U.S. federal income tax purposes, is a:

non-resident alien individual, other than certain former citizens and residents of the United States subject to tax as expatriates

foreign corporation

foreign estate or trust

A non-U.S. holder does not include an individual who is present in the United States for 183 days or more in the taxable year of disposition and is not otherwise a resident of the United States for U.S. federal income tax purposes. Such an individual is urged to consult his or her own tax advisor regarding the U.S. federal income tax consequences of the sale, exchange or other disposition of common stock.

This discussion is based on the Internal Revenue Code of 1986, as amended (the Code ), and administrative pronouncements, judicial decisions and final, temporary and proposed Treasury Regulations, changes to any which subsequent to the date of this prospectus may affect the tax consequences described herein. This discussion does not address all aspects of U.S. federal income and estate taxation that may be relevant to non-U.S. holders in light of their particular circumstances such as non-U.S. holders subject to special tax treatment under U.S. federal tax laws (including partnerships or other pass-through entities, controlled foreign corporations, passive foreign investment companies, banks and insurance companies, dealers in securities, holders of securities held as part of a straddle, hedge, conversion transaction or other risk-reduction transaction, non-U.S. holders that do not hold our common stock as a capital asset and persons who hold or receive common stock as compensation). In addition, this discussion does not address any tax consequences arising under the laws of any state, local or foreign jurisdiction.

We have not requested a ruling from the Internal Revenue Service ( IRS ) in connection with the tax consequences described herein. Accordingly, the discussion below neither binds the IRS nor precludes it from adopting a contrary position.

IN VIEW OF THE FOREGOING AND BECAUSE THE FOLLOWING DISCUSSION IS INTENDED AS A GENERAL SUMMARY ONLY, YOU ARE URGED TO CONSULT YOUR OWN TAX ADVISORS AS TO THE SPECIFIC TAX CONSEQUENCES OF THE OWNERSHIP OR DISPOSITION OF OUR STOCK, INCLUDING THE APPLICABLE FEDERAL, STATE, LOCAL AND FOREIGN TAX CONSEQUENCES, IN LIGHT OF YOUR OWN PARTICULAR TAX SITUATION.

# **Dividends**

As discussed under Dividend policy above, the company does not currently expect to pay dividends. In the event that the company does pay dividends, dividends paid to a non-U.S. holder of common stock generally will be subject to withholding tax at 30% rate or a reduced rate specified by an applicable income tax treaty. In order to obtain a reduced rate of withholding, a non-U.S. holder will be required to provide an IRS Form W-8BEN certifying its entitlement to benefits under a treaty.

The withholding tax does not apply to dividends paid to a non-U.S. holder who provides a Form W-8ECI, certifying that the dividends are effectively connected with the non-U.S. holder  $\,$  s

conduct of a trade or business within the United States. Instead, the effectively connected dividends will be subject to regular U.S. income tax as if the non-U.S. holder were a U.S. resident, subject to an applicable income tax treaty providing otherwise. A non-U.S. corporation receiving effectively connected dividends may also be subject to an additional branch profits tax imposed at a rate of 30% (or a lower treaty rate).

# Gain on disposition of common stock

A non-U.S. holder generally will not be subject to U.S. federal income tax on gain realized on a sale or other disposition of common stock unless:

the gain is effectively connected with a trade or business of the non-U.S. holder in the United States, subject to an applicable treaty providing otherwise

the company is or has been a U.S. real property holding corporation, as defined below, at any time within the five-year period preceding the disposition or the non-U.S. holder s holding period, whichever period is shorter, and its common stock has ceased to be traded on an established securities market prior to the beginning of the calendar year in which the sale or disposition occurs

In general, we would be a U.S. real property holding corporation if interests in U.S. real estate comprised the majority of our assets. The company believes that it is not, and does not anticipate becoming, a U.S. real property holding corporation.

## Information reporting requirements and backup withholding

Information returns will be filed with the IRS in connection with payments of dividends and the proceeds from a sale or other disposition of common stock. A non-U.S. holder may have to comply with certification procedures to establish that it is not a United States person in order to avoid information reporting and backup withholding tax requirements. The certification procedures required to claim a reduced rate of withholding under a treaty will satisfy the certification requirements necessary to avoid the backup withholding tax as well. The amount of any backup withholding from a payment to a non-U.S. holder will be allowed as a credit against such holder s United States federal income tax liability and may entitle such holder to a refund, provided that the required information is furnished to the IRS.

# Federal estate tax

Individual non-U.S. holders and entities the property of which is potentially includible in such an individual s gross estate for U.S. federal estate tax purposes (for example, a trust funded by such an individual and with respect to which the individual has retained certain interests or powers), should note that, absent an applicable treaty benefit, the common stock will be treated as U.S. situs property subject to U.S. federal estate tax.

#### **Underwriters**

Under the terms and subject to the conditions contained in an underwriting agreement dated the date of this prospectus, the underwriters named below, for whom J.P. Morgan Securities Inc. and Morgan Stanley & Co., Incorporated are acting as representatives, have severally agreed to purchase, and we have agreed to sell to them, severally, the number of shares indicated below:

Underwriters Number of shares

J.P. Morgan Securities Inc. Morgan Stanley & Co., Incorporated Banc of America Securities LLC Natexis Bleichroeder Inc.

Total 3,500,000

The underwriters are offering the shares of common stock subject to their acceptance of the shares from us and subject to prior sale. The underwriting agreement provides that the obligations of the several underwriters to pay for and accept delivery of the shares of common stock offered by this prospectus are subject to the approval of specified legal matters by their counsel and to other conditions. The underwriters are obligated to take and pay for all of the shares of common stock offered by this prospectus if any such shares are taken. However, the underwriters are not required to take or pay for the shares covered by the underwriters—over-allotment option described below.

The underwriters initially propose to offer the shares of common stock directly to the public at the public offering price listed on the cover page of this prospectus and to certain dealers at a price that represents a concession not in excess of \$ per share under the public offering price. Any underwriter may allow, and such dealers may re-allow, a concession not in excess of \$ per share to other underwriters or to certain dealers. After the public offering of the shares of common stock, the offering price and other selling terms may from time to time be varied by the representatives.

We have granted to the underwriters an option, exercisable for 30 days from the date of this prospectus, to purchase up to an aggregate of 525,000 additional shares of common stock at the public offering price listed on the cover page of this prospectus, less underwriting discounts and commissions. The underwriters may exercise this option solely for the purpose of covering over-allotments, if any, made in connection with the offering of the shares of common stock offered by this prospectus. To the extent the option is exercised, each underwriter will become obligated, subject to certain conditions, to purchase about the same percentage of the additional shares of common stock as the number listed next to the underwriter s name in the preceding table bears to the total number of shares of common stock listed next to the names of all underwriters in the preceding table. If the underwriters option is exercised in full, the total price to the public would be \$ , the total underwriters discounts and commissions would be \$ and the total proceeds to us would be \$

The underwriters have informed us that they do not intend sales to discretionary accounts to exceed five percent of the total number of shares of common stock offered by them.

We, each of our directors and officers and certain venture capital funds affiliated with certain of our directors that held an aggregate of 5,543,183 shares of our common stock as of

December 31, 2006 have agreed that, without the prior written consent of J.P. Morgan Securities Inc. and Morgan Stanley & Co., Incorporated on behalf of the underwriters, we and they will not, during the period ending 30 days after the date of this prospectus:

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 of or otherwise transfer or dispose of, directly or indirectly, any shares of common stock or any securities convertible into or exercisable or exchangeable for common stock

enter into any swap or other agreement that transfers to another, in whole or in part, any of the economic consequences of ownership of the common stock

whether any transaction described above is to be settled by delivery of common stock or such other securities, in cash or otherwise. These restrictions do not apply to:

in the case of a corporation, the transfer of shares of our common stock or any shares convertible into common stock to any wholly-owned subsidiary of such corporation, provided that in such case, the transferee will execute an agreement stating that the transferee is subject to the restrictions described above

transactions relating to shares of common stock or other securities acquired in open market transactions after the completion of the offering of the shares, provided that no filing or other public announcement by any party under the Exchange Act shall be required or made in connection with subsequent sales of common stock or other securities acquired in such open market transactions (other than a filing on a Form 5 made after the expiration of the 30-day period referred to above)

transfers of any shares of common stock or other securities convertible into common stock made as a gift, to a trust, to limited partners, limited liability company members or stockholders of our executive officers, directors, or holders of substantially all of our stock, or to immediate family members, provided that the transferee agrees to be bound by the restrictions described above and if the donor or transferor is a reporting person subject to Section 16(a) of the Exchange Act, any gifts or transfers made in accordance with this paragraph will not require such person to and such person will not voluntarily, file a report of such transaction on Form 4 under the Exchange Act

sales or other transfers of up to 100,000 shares of common stock in the aggregate by non-executive employees of the Company

sales or other transfer by certain executive officers of the Company pursuant to trading plans established, pursuant to Rule 1065-1 under the Securities Exchange Act of 1934, prior to the date of this prospectus

Notwithstanding the foregoing, if (1) during the last 17 days of the 30-day restricted period, we issue an earnings release or material news or a material event relating to us occurs; or (2) prior to the expiration of the 30-day restricted period, we announce that we will release earnings results during the 16-day period beginning on the last day of the 30-day period, the restrictions described above shall continue to apply until the expiration of the 18-day period beginning on the issuance of the earnings release or the occurrence of the material news or material event.

The following table shows the underwriting discounts and commissions that we are to pay to the underwriters in connection with this offering. These amounts are shown assuming both no

exercise and full exercise of the underwriters option to purchase additional shares of our common stock.

Paid by Vanda	No exercise	Full exercise
Per Share	\$	\$
Total	\$	\$

We estimate that the total expenses of this offering payable by us, not including the underwriting discounts and commissions, will be approximately \$1,250,000.

In order to facilitate the offering of the common stock, the underwriters may engage in transactions that stabilize, maintain or otherwise affect the price of the common stock. Specifically, the underwriters may sell more shares than they are obligated to purchase under the underwriting agreement, creating a short position. A short sale is covered if the short position is no greater than the number of shares available for purchase by the underwriters under the over-allotment option. The underwriters can close out a covered short sale by exercising the over-allotment option or purchasing shares in the open market. In determining the source of shares to close out a covered short sale, the underwriters will consider, among other things, the open market price of shares compared to the price available under the over-allotment option. The underwriters may also sell shares in excess of the over-allotment option, creating a naked short position. The underwriters must close out any naked short position by purchasing shares in the open market. 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 after pricing that could adversely affect investors who purchase in this offering. In addition, to stabilize the price of the common stock, the underwriters may bid for, and purchase, shares of common stock in the open market. Finally, the underwriting syndicate may reclaim selling concessions allowed to an underwriter or a dealer for distributing the common stock in this offering, if the syndicate repurchases previously distributed common stock in transactions to cover syndicate short positions or to stabilize the price of the common stock. Any of these activities may stabilize or maintain the market price of the common stock above independent market levels. The underwriters are not required to engage in these activities, and may end any of these activities at any time.

We trade our common stock on The Nasdaq Global Market under the trading symbol VNDA.

We and the underwriters have agreed to indemnify each other against certain liabilities, including liabilities under the Securities Act of 1933, as amended.

Certain of the underwriters or their affiliates have provided from time to time, and may provide in the future, investment and commercial banking and financial advisory services to Vanda and its affiliates in the ordinary course of business, for which they have received and may continue to receive customary fees and commissions.

A prospectus in electronic format will be made available on the websites maintained by one or more of the lead managers of this offering and may also be made available on websites maintained by other underwriters. The underwriters may agree to allocate a number of shares to underwriters for sale to their online brokerage account holders. Internet distributions will be allocated by the lead managers to underwriters that may make Internet distributions on the same basis as other allocations.

Each underwriter acknowledges and agrees that it has only communicated or caused to be communicated and will only communicate or cause to be communicated any invitation or inducement to engage in investment activity (within the meaning of Section 21 of the Financial

Services and Markets Act 2000, also known as the FSMA) received by it in connection with the issue or sale of any common stock in circumstances in which Section 21(1) of the FSMA does not apply to us. Each underwriter further acknowledges and agrees that it has complied and will comply with all applicable provisions of the FSMA with respect to anything done by it in relation to the shares in, from or otherwise involving the United Kingdom.

Regarding each Member State of the European Economic Area which has implemented European Union Directive 2003/71/EC, also known as the EU Prospectus Directive, each underwriter has acknowledged and agreed that, from and including the date on which the EU Prospectus Directive is implemented in such Member State, such underwriter has not made and will not make an offer of our securities to the public in that Member State prior to the publication of a prospectus in relation to such securities which has been approved by the competent authority in that Member State or, where appropriate, approved in another applicable Member State and notified to the competent authority in that Member State, all in accordance with the EU Prospectus Directive, except that it may, with effect from and including the relevant implementation date of the EU Prospectus Directive, make an offer of shares to the public in that Member State at any time

to legal entities which are authorized or regulated to operate in the financial markets or, if not so authorized or regulated, whose corporate purpose is solely to invest in securities

to any legal entity which has two or more of (1) an average of at least 250 employees during the last financial year; (2) a total balance sheet of more than 43,000,000 and (3) an annual net turnover of more than 50,000,000, as shown in its last annual or consolidated accounts

to fewer than 100 natural or legal persons (other than qualified investors as defined in the EU Prospectus Directive) subject to obtaining the prior consent of the book-running manger(s) for any such offer

in any other circumstances which do not require the publication by the Issuer of a prospectus pursuant to Article 3 of the EU Prospectus Directive

For the purposes of this provision, the expression an offer of shares to the public in relation to any shares in any Member State that has implemented the EU Prospectus Directive 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 that Member State by any measure implementing the EU Prospectus Directive in that Member State.

No prospectus (including any amendment, supplement or replacement thereto) has been prepared in connection with this offering that has been approved by the Autorité des marchés financiers in France or by the competent authority of another applicable Member State that has provided notice to the Autorité des marchés financiers. No securities to be registered pursuant to this offering have been offered or sold or will be offered or sold, directly or indirectly, to the public in France except to permitted investors, such permitted investors consisting of persons licensed to provide the investment service of portfolio management for the account of third parties, qualified investors (investisseurs qualifiés) acting for their own account and/or investors belonging to a limited circle of investors (cercle restreint d investisseurs) acting for their own account, with qualified investors and limited circle of investors having the meaning ascribed to them in Articles L. 411-2, D. 411-1, D. 411-2, D. 734-1, D. 744-1, D. 754-1 and D. 764-1 of the French Code Monétaire et Financier and applicable regulations thereunder. None of this prospectus or any other materials related to the offering or information contained therein relating to the securities to be registered pursuant to this offering has been released, issued or distributed to the public in France except to the permitted investors described above, and the

direct or indirect resale to the public in France of any Securities acquired by any such permitted investors may be made only as provided by Articles L. 411-1, L. 411-2, L. 412-1 and L. 621-8 to L. 621-8-3 of the French Code Monétaire et Financier and applicable regulations thereunder.

This offering has not been cleared by the Italian Securities Exchange Commission (Company s Commissione Nazionale per le Società e la Borsa, also known as CONSOB) pursuant to Italian securities legislation and, accordingly, each underwriter has acknowledged and agreed that the Company s common stock may not and will not be offered, sold or delivered, nor may or will copies of the prospectus or any other documents relating to such common stock be distributed in Italy, except (i) to professional investors (operatori qualificati), as defined in Article 31, second paragraph, of CONSOB Regulation No. 11522 of July 1, 1998, as amended, or (ii) in other circumstances which are exempted from the rules on solicitation of investments pursuant to Article 100 of Legislative Decree No. 58 of February 24, 1998 (also known as the Financial Services Act) and Article 33, first paragraph, of CONSOB Regulation No. 11971 of May 14, 1999, as amended.

Any offer, sale or delivery of the common stock or distribution of copies of the prospectus or any other document relating to the common stock in Italy may and will be effected in accordance with all Italian securities, tax, exchange control and other applicable laws and regulations, and, in particular, will be made by an investment firm, bank or financial intermediary permitted to conduct such activities in Italy in accordance with the Financial Services Act, Legislative Decree No. 385 of September 1, 1993, as amended, the above-mentioned CONSOB Regulation No. 11522, and any other applicable laws and regulations. Such offer, sale or delivery will further be made in compliance with Article 129 of the above-mentioned Legislative Decree No. 385 and the implementing guidelines of the Bank of Italy, and will be made in compliance with any other applicable notification requirement or limitation which may be imposed by CONSOB or the Bank of Italy.

Any investor purchasing the common stock in the offering is solely responsible for ensuring that any offer or resale of the common stock it purchased in the offering occurs in compliance with applicable laws and regulations.

The prospectus and the information contained therein are intended only for the use of its recipient and, unless in circumstances which are exempted from the rules on solicitation of investments pursuant to Article 100 of the Financial Services Act and Article 33, first paragraph, of CONSOB Regulation No. 11971 of May 14, 1999, as amended, is not to be distributed, for any reason, to any third party resident or located in Italy. No person resident or located in Italy other than the original recipients of this document may rely on it or its content.

Italy has only partially implemented the EU Prospectus Directive. The provisions above regarding the EU Prospectus Directive shall apply with respect to Italy only to the extent that the relevant provisions of the EU Prospectus Directive have already been implemented in Italy.

Insofar as the requirements above are based on laws which are superseded at any time pursuant to the implementation of the EU Prospectus Directive, such requirements shall be replaced by the applicable requirements under the EU Prospectus Directive.

#### Legal matters

Gunderson Dettmer Stough Villeneuve Franklin & Hachigian, LLP will pass upon the validity of the common stock offered by this prospectus. Hoffman, Warnick & D Alessandro LLC, will pass upon certain intellectual property matters. Davis Polk & Wardwell will pass upon certain legal matters for the underwriters.

#### **Experts**

The consolidated financial statements for the period from March 13, 2003 (date of inception) to December 31, 2003, and for the years ended December 31, 2004 and 2005 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 Securities and Exchange Commission (SEC), Washington, D.C. 20549, a registration statement on Form S-1 under the Securities Act of 1933, with respect to our common stock offered hereby. This prospectus, which forms part of the registration statement, does not contain all of the information set forth in the registration statement and the exhibits and schedules to the registration statement. Some items are omitted in accordance with the rules and regulations of the SEC. For further information about us and our common stock, we refer you to the registration statement and the exhibits and schedules to the registration statement filed as part of the registration statement, as well as the reports and other information that we have filed with the SEC. Statements contained in this prospectus as to the contents of any contract or other document filed as an exhibit are qualified in all respects by reference to the actual text of the exhibit. You may read and copy the registration statement, including the exhibits and schedules to the registration statement, as well as the reports and other information we have filed with the SEC, at the SEC s Public Reference Room at 100 F Street N.E., Washington, D.C. 20549. You can obtain information on the operation of the Public Reference Room by calling the SEC at 1-800-SEC-0330. In addition, the SEC maintains an Internet site at www.sec.gov, from which you can electronically access the registration statement, including the exhibits and schedules to the registration statement.

#### **Index to consolidated financial statements**

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#### Report of Independent Registered Public Accounting Firm

To the Board of Directors and Stockholders of Vanda Pharmaceuticals Inc. (A development stage company):

In our opinion, the accompanying consolidated balance sheets and the related consolidated statements of operations, changes in stockholders—equity and cash flows present fairly, in all material respects, the financial position of Vanda Pharmaceuticals Inc. and its subsidiary (a development stage company) at December 31, 2004 and 2005, and the results of operations and cash flows for the period from March 13, 2003 (date of inception) to December 31, 2004 and 2005 and for the period from March 13, 2003 (date of inception) to December 31, 2005 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 McLean, Virginia February 15, 2006, except for Note 8 as to which the date is April 12, 2006

#### Vanda Pharmaceuticals Inc. (A development stage company) Consolidated Balance Sheets

	December 31 2004	2005
Assets		
Current assets		
Cash and cash equivalents	\$ 16,259,770	\$ 21,012,815
Short-term investments		10,141,189
Prepaid expenses and other current assets	190,604	2,217,960
Total current assets	16,450,374	33,371,964
Property and equipment, net	1,251,867	1,110,576
Deposits	50,000	840,000
Restricted cash	23,233	430,230
Total assets	\$ 17,752,241	\$ 35,752,770
Liabilities and stockholders equity Current liabilities	\$ 718,606	\$ 2,254,897
Accounts payable Accrued liabilities	689,428	2,528,091
Deferred rent and credit on lease concession, current	3,549	8,131
Current portion of long-term debt	173,929	142,461
Current portion of capital lease	37,241	142,401
Deferred grant revenue	37,271	129,950
Total current liabilities	1,622,753	5,063,530
Deferred rent and credit on lease concession, less current portion	30,371	24,433
Long-term debt, less current portion	142,487	21,133
Capital lease, less current portion	13,043	
Total liabilities	1,808,654	5,087,963
Commitments Stockholders equity		

Series A Preferred Stock, \$0.001 par value; 10,000,000 shares authorized, issued and outstanding at December 31, 2004 and 2005,		
respectively; liquidation preference of \$10,000,000	9,963,541	9,963,541
Series B Preferred Stock, \$0.001 par value; 42,276,437 shares		
authorized; 15,040,654 and 42,276,437 shares issued and outstanding at		
December 31, 2004 and 2005, respectively; liquidation preference of		
\$52,000,018	18,345,023	51,831,646
Common stock, \$0.001 par value; 50,000,000 and 70,000,000 shares		
authorized and 3,020 and 98,945 shares issued and outstanding at		
December 31, 2004 and 2005, respectively	3	99
Additional paid-in capital	340,637	23,982,981
Deferred stock-based compensation	(257,934)	(18,766,443)
Accumulated other comprehensive loss	(2,576)	(17,609)
Deficit accumulated during the development stage	(12,445,107)	(36,329,408)
Total stockholders equity	\$ 15,943,587	\$ 30,664,807
Total liabilities and stockholders equity	\$ 17,752,241	\$ 35,752,770

The accompanying notes are an integral part of these consolidated financial statements.

# Vanda Pharmaceuticals Inc. (A development stage company) Consolidated Statements of Operations

	Period from March 13, 2003 (inception) to December 31, 2003		Year ended December 31, 2004 2005			•	Period from March 13, 2003 (inception) to December 31, 2005		
Revenues from services	\$	47,565	\$	33,980	\$		\$	81,545	
Operating expenses:	·	,	·	,				,	
Research and development	2	,010,532	,	7,442,983	]	16,890,615		26,344,130	
General and administrative	1	,052,659	2	2,119,394		7,396,038		10,568,091	
Total operating expenses	3	,063,191	ý	9,562,377	2	24,286,653		36,912,221	
Loss from operations Other income (expense):	(3	,015,626)	(9	9,528,397)	(2	24,286,653)	(	(36,830,676)	
Interest income		52,595		100,785		435,537		588,917	
Interest expense		(8,090)		(41,934)		(25,629)		(75,653)	
Other income		300		209		93		602	
Total other income		44,805		59,060		410,001		513,866	
Loss before tax provision Tax provision	(2	,970,821)	(9	9,469,337) 4,949	(2	23,876,652) 7,649	(	(36,316,810) 12,598	
Net loss Beneficial conversion feature deemed	(2	,970,821)	(9	9,474,286)	(2	23,884,301)	(	(36,329,408)	
dividend to preferred stockholders					(3	33,486,623)	(	(33,486,623)	
Net loss attributable to common stockholders	\$ (2	,970,821)	\$ (9	9,474,286)	\$ (5	57,370,924)	\$ (	(69,816,031)	
Basic and diluted net loss per share attributable to common stockholders	\$	(983.72)	\$	(3,137.18)	\$	(3,374.33)			

Shares used in calculation of basic and diluted net loss per share attributable to common stockholders

3,020 3,020 17,002

The accompanying notes are an integral part of these consolidated financial statements.

#### Vanda Pharmaceuticals Inc. (A development stage company) Statements of Changes in Stockholders Equity

Series A preferred stock		Series B preferred stock		Common stock		Additional	Deferred	other	Deficit d accumulated during the	Comp
preferre	u Stuck	preferre	EU SIUCK	Stoc	k Par	paiu-iii	Stock-Dasett	ompi enensi	veaeveiopinent	Comp
Shares	Par value	Shares	Par value	Shares	value	capital	compensation	n loss	stage	Ì
	\$		\$		\$	\$	\$	\$	\$	\$
0,000,000	9,963,541									
				3,020	3	3,997				
						12,628				
									(2,970,821)	(2,
								(2,315)		
										(2,9
0,000,000	9,963,541			3,020	3	16,625		(2,315)	(2,970,821)	
		15,040,654	18,345,023							
						27,945				
						281,130	(281,130) 23,196			

14,937

(9,474,286) (9,

(261)

(9,

# Vanda Pharmaceuticals Inc. (A development stage company) Statements of Changes in Stockholders Equity (continued)

erie errec	s A d stock	Serio preferre		Common		Additional paid-in	Deferred stock-based	Accumulated other comprehensive	Deficit accumulated during the development
	Par value	Shares	Par value	Shares	Par value	capital	compensation	loss	stage
000	9,963,541	15,040,654	18,345,023	3,020	3	340,637	(257,934)	(2,576)	(12,445,107)
		27,235,783	33,486,623						
				95,925	96	31,658			
						18,788,385	(18,788,385)		
						1,702,625	(1,702,625)		
						3,119,676			
							1,982,501		
						33,486,623			
						(33,486,623)			

(23,884,301)

(17,711)

2,678

00 \$ 9,963,541 42,276,437 \$ 51,831,646 98,945 \$ 99 \$ 23,982,981 \$ (18,766,443) \$ (17,609) \$ (36,329,408)

# Vanda Pharmaceuticals Inc. (A development stage company) Consolidated Statements of Cash Flows

	Period from March 13, 2003 (inception) to December 31, 2003	2004	Year ended December 31, 2005	Period from March 13, 2003 (inception) to December 31, 2005
Cash flows from operating activities				
Net loss	\$ (2,970,821)	\$ (9,474,286)	\$ (23,884,301)	\$ (36,329,408)
Adjustments to reconcile net loss to net cash used in operating activities				
Depreciation and amortization	79,891	376,709	423,828	880,428
Stock-based compensation		66,078	5,102,177	5,168,255
Accretion of discount on investments Changes in assets and liabilities:			(42,335)	(42,335)
Accounts receivable	(28,489)	28,489		
Prepaid expenses and other current	(==, :==)			
assets	(97,044)	(93,024)	(2,027,544)	(2,217,960)
Deposits	450,600	(50,000)	(790,000)	(840,000)
Accounts payable Accrued expenses	458,608 432,474	415,506 99,335	1,514,868 1,860,539	2,254,897 2,528,091
Deferred grant revenue	432,474	77,333	129,950	129,950
Deferred rent and credit on lease			,	,,
concession	17,661	16,259	(1,356)	32,564
Net cash used in operating activities	(2,107,720)	(8,614,934)	(17,714,174)	(28,435,518)
Cash flows from investing activities				
Purchases of property and equipment	(1,161,921)	(414,531)	(291,978)	(1,868,430)
Purchases of short-term investments			(11,846,176)	(11,846,176)
Maturities of short-term investments Investments in restricted cash			1,750,000 (430,230)	1,750,000 (430,230)
investments in restricted cash			(430,230)	(+30,230)
Net cash used in investing activities	(1,161,921)	(414,531)	(10,818,384)	(12,394,836)
Cash flows from financing activities				
Proceeds from borrowings on note payable	515,147			515,147
payaore	313,147			313,147

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Principal payments on obligations under capital lease Principal payments on note payable Proceeds from the issuance of	(45,010)	(42,887) (156,446)	(51,569) (172,617)	(94,456) (374,073)
preferred stock, net of issuance costs Proceeds from exercise of stock options	9,963,541	18,345,023	33,486,623 31,754	61,795,187 31,754
Proceeds from issuance of common stock	4,000			4,000
Net cash provided by financing activities	10,437,678	18,145,690	33,294,191	61,877,559
Effect of foreign currency translation	(2,315)	(22,177)	(8,588)	(34,390)
Net increase in cash and cash equivalents	7,165,722	9,094,048	4,753,045	21,012,815
Cash and cash equivalents Beginning of period		7,165,722	16,259,770	
End of period	\$ 7,165,722	\$ 16,259,770	\$ 21,012,815	\$ 21,012,815
Supplemental disclosure Cash payments for interest Supplemental disclosure of noncash financing activities Equipment acquired through	\$ 4,221	\$ 41,354	\$ 25,043	\$ 70,618
obligation under capital lease	\$	\$ 95,305	\$	\$ 95,305

The accompanying notes are an integral part of these consolidated financial statements.

Vanda Pharmaceuticals Inc.
(A development stage company)
Notes to consolidated financial statements (continued)

#### 1. Business organization and presentation

#### **Business organization**

Vanda Pharmaceuticals Inc. ( Vanda or the Company ) was founded in November 2002 and commenced its operations on March 13, 2003. Vanda is a biopharmaceutical company focused on the development and commercialization of small molecule therapeutics, with exclusive worldwide commercial rights to three product candidates in clinical development for various central nervous system disorders. The Company s lead product candidate, iloperidone, is a compound for the treatment of schizophrenia and bipolar disorder and is in a Phase III trial for schizophrenia. The Company s second product candidate, VEC-162, is a compound for the treatment of sleep and mood disorders and has recently completed a Phase III trial for transient insomnia. VEC-162 is also ready for Phase II trials for the treatment of depression. The Company s third product candidate, VSF-173, is a compound for the treatment of excessive sleepiness and is ready for a Phase II trial. Each of these product candidates benefits from new chemical entity (NCE) patent protection and may offer substantial advantages over approved therapies.

The Company expects to complete its Phase III trial for iloperidone in the first half of 2007. If this trial is successful, the Company will file a New Drug Application (NDA) for approval with the Food and Drug Administration (FDA) later that year. The Company recently completed an efficacy and safety Phase III trial of VEC-162 for insomnia and announced top-line results in a Phase III trial in November 2006. The Company also expects to begin a Phase II trial of VSF-173 for excessive sleepiness in the second half of 2006.

Vanda Pharmaceuticals Pte. Ltd. ( Vanda Singapore ) is a limited liability company domiciled and incorporated in Singapore on February 24, 2003 as a wholly-owned subsidiary of Vanda Pharmaceuticals Inc. Vanda Singapore s principal activity is drug research using genetic and genomic sciences.

#### Capital resources

Although the Company was incorporated in November 2002, the Company did not commence operations until March 13, 2003, the date on which the Company first issued capital stock and began incurring expenses. Prior to March 13, 2003, the Company did not have any assets or liabilities, directly incur any expenses, or indirectly incur any expenses by a related party. Since its inception, the Company has devoted substantially all of its efforts to business planning, research and development, recruiting management and technical staff, acquiring operating assets and raising capital. Accordingly, the Company is considered to be in the development stage as defined in Statement of Financial Accounting Standards (SFAS) No. 7, Accounting and Reporting by Development Stage Enterprises.

The Company s activities will necessitate significant uses of working capital throughout 2006 and beyond. Additionally, the Company s capital requirements will depend on many factors, including the success of the Company s research and development efforts, payments received under contractual agreements with other parties, if any, and the status of competitive products. The Company plans to continue financing operations with the cash received from the

private placement of Series B Preferred Stock (see Note 8) and the Company plans to seek additional sources of funding in 2006. The Company s failure to raise additional capital, as and when

needed, could have a negative impact on the financial condition and the ability of the Company to execute its business strategy. In the absence of our ability to raise additional private equity capital, we are also prepared and have the ability to curtail our existing clinical trial commitments and extend them in such a manner so that we have operating funds through the end of 2007.

#### **Basis of presentation**

The accompanying consolidated financial statements include the accounts of the Company and its wholly-owned subsidiary. All inter-company balances and transactions have been eliminated. The accompanying consolidated financial statements are prepared in accordance with accounting principles generally accepted in the United States of America.

#### Unaudited pro forma balance sheet

The unaudited pro forma balance sheet gives effect to the conversion of the Series A and B Preferred Stock in the event of an initial public offering ( IPO ), as if it occurred on December 31, 2005. The shares of the Company s Series A and B Preferred Stock shall be converted into common stock on a 3.309755-to-one basis automatically upon consummation of an IPO, after giving effect to the reverse split of the Company s common stock described in Note 8 below.

#### 2. Summary of significant accounting policies

#### Cash and cash equivalents

For purposes of the consolidated balance sheet and consolidated statement of cash flows, cash equivalents represent all highly-liquid investments with an original maturity date of three months or less. At December 31, 2005, the Company maintained all of its cash and cash equivalents in two financial institutions. Deposits held with these institutions may exceed the amount of insurance provided on such deposits. Generally, these deposits may be redeemed upon demand, and the Company believes there is minimal risk of losses on such cash balances.

#### **Short-term investments**

The Company maintained highly-liquid investments throughout the period ending December 31, 2005, which were classified as available-for-sale because they can be utilized for current operations. The Company s investment policy requires the selection of high-quality issuers, with bond ratings of AAA to A1+/P1. These available-for-sale securities are accounted for at their fair market value and unrealized gains and losses on these securities, if any, are included in accumulated other comprehensive loss in stockholders—equity. Interest and dividend income is recorded when earned and included in interest income. Premiums and discounts on short-term investments are amortized and accreted to maturity and included in interest income. The Company uses the specific identification method in computing realized gains and losses on the sale of investments, which would be included in the consolidated statements of operations when generated. For the period from March 13, 2003 (inception) to December 31, 2003 and for the years ended December 31, 2004 and 2005, the Company did not have any realized gains or losses.

The following is a summary of the Company s available-for-sale short-term investments as of December 31, 2005:

	Amortized cost		Gross unrealized gains		Gross unrealized losses	Fair market value	
U.S. government agencies U.S. corporate debt	\$	6,054,023 4,084,488	\$	847 1,831	\$	\$	6,054,870 4,086,319
	\$	10,138,511	\$	2,678	\$	\$	10,141,189

#### **Concentrations of credit risk**

Financial instruments which potentially subject the Company to significant concentrations of credit risk consist primarily of cash and cash equivalents and short-term investments. The Company places its cash and cash equivalents and short-term investments with highly-rated financial institutions.

#### Fair value of financial instruments

The carrying amounts of the Company s financial instruments, which include cash and cash equivalents, short-term investments, and accounts payable, approximate their fair values due to their short maturities. The fair value of the long-term debt approximates its carrying value based on the variable nature of interest rates and current market rates available to the Company.

#### **Property and equipment**

Property and equipment are stated at cost less accumulated depreciation and amortization. Depreciation of property and equipment is provided on a straight-line basis over the estimated useful lives of the assets, generally three to seven years. Amortization of leasehold improvements is provided on a straight-line basis over the shorter of their estimated useful life or the lease term. The costs of additions and betterments are capitalized, and repairs and maintenance costs are charged to operations in the period incurred.

Upon retirement or disposition of property and equipment, the cost and accumulated depreciation are removed from the accounts and any resulting gain or loss is reflected in general and administrative expenses for that period.

#### Impairment of long-lived assets

The Company assesses the recoverability of its long-lived assets by determining whether the carrying value of such assets can be recovered through undiscounted future operating cash flows. If impairment is indicated, the Company

measures the amount of such impairment by comparing the fair value to the carrying value. There have been no indicators of impairment through December 31, 2005.

#### Restricted cash

During 2005, in conjunction with the lease of the office and laboratory space building, the Company provided the landlord with a letter of credit, which was collateralized with a restricted cash deposit in the amount of \$430,230 (see Note 6). The deposit is recorded as non-current restricted cash at December 31, 2005.

#### **Deferred grant revenue**

Vanda Singapore entered into an agreement with the Economic Development Board of Singapore (EDB) to provide a grant for a Development Project. During 2005, the Company submitted its first asset-related claim with the EDB and received a cash payment of \$127,866. Given that the Company has not met the conditions attached to the grant, the payment has been recorded as deferred grant revenue on the balance sheet at December 31, 2005. Management expects that a resolution is likely to be reached with the EDB in the near future.

#### Translation of foreign currency

The functional currency of the Company s wholly-owned foreign subsidiary located in Singapore is the local currency. Assets and liabilities of the Company s foreign subsidiary are translated to United States dollars based on exchange rates at the end of the reporting period. Income and expense items are translated at weighted average exchange rates prevailing during the reporting period. Translation adjustments are accumulated in a separate component of stockholder s equity. Translation gains or losses are included in the determination of operating results.

#### Other comprehensive income (loss)

SFAS No. 130, *Reporting Comprehensive Income*, requires a full set of general-purpose financial statements to include the reporting of comprehensive income. Other comprehensive loss is composed of two components, net loss and other comprehensive income. At December 31, 2004 and 2005, other comprehensive loss of \$2,576 and \$17,609, respectively, consists of cumulative translation adjustments due to foreign currency and unrealized gains on short-term investments.

#### **Revenue recognition**

Revenue is recognized upon delivery of products to customers. Revenue earned under research and development contracts are recognized in accordance with the proportional performance method outlined in Staff Accounting Bulletin No. 104 whereby the extent of progress toward completion is measured on the cost-to-cost basis; however, revenue recognized at any point will not exceed the cash received. When the current estimates of total contract revenue and contract cost indicate a loss, a provision for the entire loss on the contract is made in the period which it becomes probable. All costs related to these agreements are expensed as incurred. Revenue is derived principally from consulting agreements the Company entered into during its start-up phase to defray research costs. Vanda completed its obligations under these agreements during the year ended December 31, 2004, and no longer seeks such arrangements.

The Company will use the substantive milestone payment method for its revenues recognition policy. Under this method, revenue is recognized when all milestones to be received under contractual arrangements are determined to be substantive, at-risk and the culmination of an

earnings process. Substantive milestones are payments that are conditioned upon an event requiring substantive effort, when the amount of the milestone is reasonable relative to the time, effort and risk involved in achieving the milestones and when the milestones are reasonable relative to each other and the amount of any up-front payment. If these criteria are not met, the timing of the recognition of revenue from the milestone payment may be deferred.

#### Research and development expenses

Research and development costs are expensed as incurred and include the cost of salaries, building costs, utilities, allocation of indirect costs, and expenses to third parties who conduct research and development, pursuant to development and consulting agreements, on behalf of the Company. Costs related to the acquisitions of intellectual property are expensed as incurred since the underlying technology associated with these acquisitions were made in connection with the Company s research and development efforts and have no alternative future use.

#### General and administrative expenses

General and administrative costs are expensed as incurred and consist primarily of salaries and other related costs for personnel serving executive, finance, accounting, information technology and human resource functions. Other costs include facility costs not otherwise included in research and development expense and professional fees for legal and accounting services.

#### Interest income and expense

Interest income consists of interest earned on the Company s cash and cash equivalents and short-term investments. Interest expense consists of interest incurred on equipment debt.

#### Accounting for stock-based compensation

As provided by SFAS No. 123, *Accounting for Stock-Based Compensation* (SFAS 123), the Company has elected to continue to account for its stock-based compensation programs according to the provisions of Accounting Principles Board Opinion No. 25, (APB 25) *Accounting for Stock Issued to Employees*. Accordingly, compensation expense has been recognized to the extent of employee or director services rendered based on the intrinsic value of compensatory options or shares granted under the plans. Under APB 25, compensation expense is recognized over the vesting period of the option to the extent that the fair value of the stock exceeds the exercise price of the stock at the date of grant.

Variable stock-based compensation awards are amortized and expensed in accordance with Financial Accounting Standards Board (FASB) Interpretation No. 28, *Accounting for Stock Appreciation Rights and Other Variable Stock Option Plan or Award Plans*, an accelerated vesting model. Under this model, all stock based employee compensation charges are amortized over the vesting periods of the individual stock awards.

Had the Company determined compensation cost based on the fair value at the grant date for its stock options under SFAS 123, the Company s net loss and basic and diluted net loss

### Vanda Pharmaceuticals Inc. (A development stage company)

#### Notes to consolidated financial statements (continued)

attributable to common stockholders per share would have been changed to the following pro forma amounts:

	Period from March 13, 2003 (inception) to		
	December 31, 2003	Year ended 2004	December 31, 2005
Net loss attributable to common stockholders Add: Stock based employee compensation	\$ (2,970,821)	\$ (9,474,286)	\$ (57,370,924)
expense included in net loss Less: Stock-based employee compensation		38,133	5,102,177
expense determined under SFAS 123	(33,160)	(57,954)	(5,167,246)
Pro forma net loss attributable to common stockholders	\$ (3,003,981)	\$ (9,494,107)	\$ (57,435,993)
Net loss per share: Basic and diluted, net loss attributable to common stockholders as reported	\$ (983.72)	\$ (3,137.18)	\$ (3,374.33)

The weighted average fair value of an option granted during the period from March 13, 2003 (inception) to December 31, 2003 and years ended December 31, 2004 and 2005 was \$1.06, \$3.97 and \$14.89 respectively. The fair value of each option grant is estimated on the date of the grant using the Black-Scholes option pricing model with the following assumptions for each year:

	Period from March 13, 2003 (inception) to December 31,	2003 ) to		
	2003	2004	2005	
Expected dividend yield	0%	0%	0%	
Expected volatility	0%	67%	67%-68%	
Expected term (years)	10	5	5	
Weighted average risk-free interest rate	3.65%	3.42%	4.00%	

Given the lack of an active public market for our common stock, the Company s board of directors determined the fair value of the Company s common stock for stock option awards and the Company did not employ a third party valuation firm to determine fair value. In establishing the Company s estimates of fair value, the Company considered the guidance set forth in the AICPA Practice Guide, *Valuation of Privately-Held-Company Equity Securities Issued as Compensation*, and made retrospective determinations of fair value. Information on stock

### Vanda Pharmaceuticals Inc. (A development stage company)

Notes to consolidated financial statements (continued)

option grants, net of forfeitures, during the previous two years ended December 31, 2005 is summarized as follows:

				Fair market value estimate	Intrinsic
Date of		Number of	Exercise	per common	value
Issuance	Type of equity issuance	options granted	price(1)	share	per share
06/15/04	Employee Options	3,443	\$ 0.33	\$ 3.21	\$ 2.88
09/01/04	Employee Options	91,668	0.33	4.07	3.74
12/06/04	Employee Options	777	0.33	5.69	5.36
02/10/05	Employee Options	209,893	0.33	10.52	10.19
04/05/05	Employee Options	27,974	0.33	15.99	15.66
08/15/05	Employee Options	15,559	0.33	16.85	16.52
09/28/05	Employee Options	620,973	0.33	16.85	16.52
10/03/05	Employee Options	906	0.33	17.18	16.85
11/14/05	Employee Options	83,087	0.83	17.18	16.35
12/29/05	Employee Options	358,847	4.73	17.18	12.45

(1) The Company s board of directors approved a modification to all outstanding stock option awards that were granted prior to February 10, 2005, repricing the options from their original exercise price of \$1.32 to \$0.33. According to FIN 44, the result of such a modification is to account for the modified stock option awards as variable from the date of the modification to the date the awards are exercised, forfeited, or cancelled. The Company remeasured the modified awards that were outstanding at the end of each quarter during the year ended December 31, 2005.

#### Stock warrants

The Company accounts for warrants granted to consultants and advisors under SFAS 123 and Emerging Issues Task Force Issue 96-18, *Accounting for Equity Investments that are Issued to Other Than Employees for Acquiring or in Conjunction with Selling Goods or Services*, (EITF 96-18). As such, warrants granted to non-employees are periodically re-measured and expense is incurred during their vesting terms.

#### **Income taxes**

The Company accounts for income taxes under the liability method in accordance with provisions of SFAS No. 109, *Accounting for Income Taxes*, (SFAS 109) which requires companies to account for deferred income taxes using the asset and liability method. Under the asset and liability method, current income tax expense or benefit is the amount of income taxes expected to be payable or refundable for the current year. A deferred income tax asset or liability is

recognized for future tax consequences attributable to differences between the financial statement carrying amounts of existing assets and liabilities and their respective tax bases and tax credits and loss carryforwards. Deferred tax assets are reduced by a valuation allowance when, in the opinion of management, it is more likely than not that some portion or all of the deferred tax assets will not be realized. Tax rate changes are reflected in income during the period such changes are enacted. Changes in ownership may limit the amount of net operating loss carryforwards that can be utilized in the future to offset taxable income.

#### Net loss per share

Net loss attributable to common stockholders per share is calculated in accordance with SFAS No. 128, *Earnings per Share*, and Staff Accounting Bulletin (SAB) No. 98. Basic earnings per share (EPS) is calculated by dividing the net income or loss attributable to common stockholders by the weighted average number of common shares outstanding, reduced by the weighted average unvested common shares subject to repurchase.

Diluted EPS is computed by dividing the net income or loss attributable to common stockholders by the weighted average number of other potential common stock outstanding for the period. Other potential common stock include Series A and B Preferred Stock, stock options and warrants but only to the extent that their inclusion is dilutive. The Company incurred a net loss in all periods presented, causing inclusion of any potentially dilutive securities to have an anti-dilutive affect, resulting in dilutive loss per share attributable to common stockholders and basic loss per share attributable to common stockholders being equivalent. The Company did not have any common shares issued for nominal consideration as defined under the terms of SAB No. 98, which would be included in EPS calculations.

Period from

March 13, 2003 (inception) to December 31, Year ended December 31. 2003 2004 2005 **Historical:** Numerator: Net loss \$ (2,970,821) \$ (9,474,286) \$ (23,884,301) Beneficial conversion feature deemed dividend to preferred stockholders (33,486,623)Net loss attributable to common stockholders \$ (2,970,821) \$ (9,474,286) \$ (57,370,924) Denominator: Weighted average common shares outstanding 3,020 3,020 30,346 Weighted average unvested common shares subject to repurchase (13,344)Denominator for basic and diluted net loss per share 3,020 3,020 17,002 Basic and diluted net loss per share attributable to common stockholders (983.72)\$ (3,137.18) (3,374.33)Historical outstanding anti-dilutive securities not included in diluted net loss per share calculation: Series A and B Preferred Stock(1) 7,565,703 3.021.368 15,794,632 Options to purchase common stock 236,204 314,961 1,532,540 Warrants to purchase common stock 13,626 50,335 50,335 7,930,999 3.271,198 17,377,507

(1) Common stock equivalents assuming conversion.

The unaudited pro forma shares used to compute basic and diluted net loss per share is the weighted average shares of common stock outstanding, reduced by the weighted average unvested common shares subject to repurchase, and includes the assumed conversion of the Series A and B Preferred Stock into shares of common stock as of January 1, 2005 or the actual date of issuance if later.

Year ended December 31,