

TITAN PHARMACEUTICALS INC

Form 10-K

March 15, 2006

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UNITED STATES

SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

FORM 10-K

FOR ANNUAL AND TRANSITION REPORTS PURSUANT TO

SECTIONS 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

(Mark One)

x ANNUAL REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

For the fiscal year ended December 31, 2005

Or

.. TRANSITION REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

Commission file number 001-13341

TITAN PHARMACEUTICALS, INC.

(Exact name of registrant as specified in its charter)

Delaware
(State or other jurisdiction of

incorporation or organization)

94-3171940
(I.R.S. Employer

identification number)

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400 Oyster Point Blvd., Suite 505,

South San Francisco, California
(Address of principal executive offices)

94080
(Zip code)

Registrant's telephone number, including area code: (650) 244-4990

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Name of each exchange on which registered
Common Stock, \$.001 par value	The American Stock Exchange

Securities registered pursuant to Section 12(g) of the Act: None

Indicate by check mark if the registrant is a well-known seasoned issuer as defined in Rule 405 of the Securities Act. Yes ☐ No ☒

Indicate by check mark if the registrant is not required to file reports pursuant to Section 13 or Section 15(d) of the Exchange Act. Yes ☐ No ☒

Indicate by check mark whether the registrant: (1) has filed all reports required to be filed by Section 13 or 15(d) of the Securities Exchange Act of 1934 during the preceding 12 months (or for such shorter period that the registrant was required to file such reports), and (2) has been subject to the filing requirements for the past 90 days. Yes ☒ No ☐

Indicate by check mark if disclosure of delinquent filers pursuant to Item 405 of Regulation S-K (§ 229.405 of this chapter) is not contained herein, and will not be contained, to the best of registrant's knowledge, in definitive proxy or information statements incorporated by reference in Part III of this Form 10-K or any amendment to this Form 10-K. ☐

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, or a non-accelerated filer. See definition or accelerated filer and large accelerated filer in Rule 12b-2 of the Exchange Act. (Check one):

Large accelerated filer ☐ Accelerated filer ☒ Non-accelerated filer ☐

Indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act). Yes ☐ No ☒

The aggregate market value of the 31,657,056 shares of voting and non-voting common equity held by non-affiliates of the registrant based on the closing price on June 30, 2005 was \$57.9 million.

As of March 1, 2006, 35,592,269 shares of common stock, \$.001 par value, of the registrant were issued and outstanding.

DOCUMENTS INCORPORATED BY REFERENCE: NONE

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PART I

Statements in this Form 10-K that are not descriptions of historical facts are forward-looking statements that are subject to risks and uncertainties. Actual results could differ materially from those currently anticipated due to a number of factors, including those set forth under Risk Factors including, but not limited to, the results of research and development efforts, the results of pre-clinical and clinical testing, the effect of regulation by the United States Food and Drug Administration (FDA) and other agencies, the impact of competitive products, product development, commercialization and technological difficulties, the results of financing efforts, the effect of our accounting policies, and other risks detailed in our Securities and Exchange Commission filings.

Probuphine®, Spheramine®, ProNeura and CCM are trademarks of Titan Pharmaceuticals, Inc. This Form 10-K also includes other trade names and trademarks of companies other than Titan Pharmaceuticals, Inc.

References herein to we, us, Titan, and our company refer to Titan Pharmaceuticals, Inc. and its subsidiaries unless the context otherwise requires.

Item 1. Business

(a) General Development of Business

We are a biopharmaceutical company developing proprietary therapeutics for the treatment of central nervous system (CNS) disorders, cardiovascular disease, bone disease and other disorders. Our product development programs focus primarily on large pharmaceutical markets with significant unmet medical needs and commercial potential. We are focused primarily on clinical development of the following products:

Probuphine: for the treatment of opioid dependence

Iloperidone: for the treatment of schizophrenia and related psychotic disorders (partnered with Vanda Pharmaceuticals, Inc.)

Spheramine: for the treatment of advanced Parkinson's disease (partnered with Schering AG)

DITPA: for the treatment of congestive heart failure and hyperlipidemia

Gallium maltolate: for the treatment of bone related diseases, chronic bacterial infections and cancer

We are directly developing our product candidates and also utilizing strategic partnerships. These partnerships help fund product development and enable us to retain significant economic interest in our products. In June 2004 we announced that Vanda Pharmaceuticals, Inc. (Vanda) had acquired from Novartis Pharma AG (Novartis) the worldwide rights to develop and commercialize iloperidone, our proprietary antipsychotic agent in Phase III clinical development for the treatment of schizophrenia and related psychotic disorders. Vanda is proceeding with and now funding the iloperidone Phase III development program. All of our rights and economic interests in iloperidone, including royalties on sales of iloperidone, remain essentially unchanged from our agreement with Novartis. Spheramine development is primarily funded by our corporate partner for Spheramine, Schering AG, Germany (Schering). Under the agreement, in exchange for exclusive, worldwide development, manufacturing and commercialization rights, Schering will provide clinical and manufacturing development funding, milestone payments, and a royalty to Titan on future product sales.

We were incorporated in Delaware in February 1992 and have funded our operations through various sources, including an initial public offering in January 1996 and private placements of securities, as well as proceeds from warrant and option exercises, corporate licensing and collaborative agreements, and government-sponsored research grants.

Table of Contents**(b) Financial Information About Industry Segments**

We operate in only one business segment, the development of pharmaceutical products.

(c) Narrative Description of Business**Product Development Programs**

The following table provides summary status of our products in development:

Product	Potential Indication(s)	Phase of	
		Development	Marketing Rights
Probuphine	Opioid dependence	Phase III in preparation	Titan
Iloperidone	Schizophrenia, psychosis	Phase III	Vanda Pharmaceuticals, Inc.
Spheramine	Parkinson's disease	Phase IIb	Schering AG
DITPA	Congestive heart failure	Phase II	Titan
DITPA	Hyperlipidemia	Phase II	Titan
Gallium maltolate	Bone related disease, chronic bacterial infections, cancer	Phase I	Titan

Our products are at various stages of development and may not be successfully developed or commercialized. We do not currently have any products being commercially sold. Our proposed products will require significant further capital expenditures, development, testing, and regulatory clearances prior to commercialization. We may experience unanticipated problems relating to product development and cannot predict whether we will successfully develop and commercialize any products.

Probuphine

We are developing Prob

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Spheramine

Spheramine is a cell-based therapeutic being developed for potential treatment of advanced Parkinson's disease. It utilizes our proprietary cell-coated microcarrier (CCM) technology, which enables the development of cell-based therapies for minimally-invasive, site-specific delivery to the central nervous system of therapeutic factors precisely where they are needed.

Spheramine consists of microcarriers coated with human retinal pigment epithelial cells that directly enhance brain levels of dopamine, a neurotransmitter deficient in certain brain regions in Parkinson's disease, leading to movement disorders. Preclinical studies have demonstrated the preliminary efficacy and safety of Spheramine, including blinded studies in a primate model of Parkinson's disease. Positron emission tomography (PET) imaging studies in primates have confirmed the presence of increased dopamine signals in regions treated with Spheramine. A pilot clinical study of Spheramine performed by Titan in six patients with advanced Parkinson's disease demonstrated substantial improvement (average 48%) in motor function at one-year post treatment with no significant adverse events. These results were first reported at the American Academy of Neurology (AAN) annual meeting in 2002. In June 2005, Schering AG sponsored a symposium on Spheramine at the International Congress on Parkinson's Disease and Related Disorders in Berlin. In the keynote address, Ray Watts, M.D., Professor and Chairman, Department of Neurology, University of Alabama Birmingham, presented 48-month follow-up data for the six patients in our pilot clinical study of Spheramine. The data presented indicate that Spheramine is well tolerated and that patients continued to demonstrate 43% average improvement in motor function, four years after treatment.

Spheramine is currently being studied in a multicenter, randomized, blinded, placebo controlled clinical trial in Parkinson's disease. This Phase IIb clinical study will enroll 68 patients with advanced Parkinson's disease (Hoehn and Yahr Stages III and IV) to further evaluate the efficacy, safety, and tolerability of Spheramine. Fifty-five patients have been treated to date.

Schering AG, our corporate partner for worldwide development and commercialization of Spheramine, is funding the clinical development program for Spheramine. Under this agreement, Schering received exclusive, worldwide development, manufacturing and commercialization rights, and, in addition to the clinical and manufacturing development funding and milestone payments, Schering will pay us a royalty on future product sales. The Investigational New Drug application (IND) filed by Titan with the FDA was transferred to Schering in November 2005.

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demonstrated potentially beneficial effects in preliminary human testing. A double blind, placebo controlled Phase II study in 19 patients with moderately severe (New York Hospital Association (NYHA) Class II-III) congestive heart failure demonstrated an improvement in cardiac index, decrease in systemic vascular resistance, and no significant increase in heart rate. These study results also demonstrated a decrease in serum cholesterol and triglyceride levels.

We are evaluating DITPA as a potential treatment for CHF associated with low serum thyroid hormone (T3). There are approximately nine million people in the U.S. and Europe with CHF. In the U.S., approximately 25% of patients have moderate or severe symptoms (NYHA Class III or IV), and CHF is the most common hospital discharge diagnosis in the U.S. for patients over 65.

The important role of thyroid hormone in maintaining heart and blood vessel function, and the association of low T3 and increased mortality in CHF suggest a potential role for DITPA as a thyroid hormone replacement therapy in CHF.

Patient enrollment in a double blind, placebo controlled Phase IIb clinical study with DITPA in NYHA Class III and Class IV CHF patients with low thyroid hormone (T3) levels is ongoing. The study will evaluate clinical and laboratory parameters related to severity of CHF, including change in global clinical status, echocardiographic parameters, B-type Natriuretic Peptide (BNP) levels, exercise testing and quality of life measurements in addition to safety.

DITPA is also currently being evaluated in a second randomized, double blind, placebo controlled Phase II study with NYHA Class II-IV CHF, sponsored by the Department of Veterans Affairs Cooperative Studies Program and funded by a \$3.8 million grant.

We are also developing DITPA as a potential treatment for hyperlipidemia. Recent data have indicated that further lowering of target LDL cholesterol level guidelines may be important in reducing the incidence of heart disease and stroke. Accordingly, the National Cholesterol Education Program (NCEP) has recently updated its guidelines to recommend that doctors target lower LDL cholesterol levels in their patients.

In March 2006, we initiated a Phase II, randomized, double-blind, placebo-controlled study to evaluate DITPA in individuals receiving standard lipid-lowering therapy, whose LDL cholesterol levels are above NCEP guidelines. The primary objective of the study is to evaluate DITPA as a cholesterol lowering agent in combination with standard therapy in patients with LDL cholesterol levels greater than the NCEP goals.

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In 2005, we completed a dose ranging Phase I clinical study of gallium maltolate in cancer patients. Significant blood levels of gallium were achieved, and a maximum tolerated dose level was not reached in this study. We are currently completing development of an optimal formulation of gallium maltolate with increased bioavailability, and subsequent clinical trials are planned to use this new formulation of gallium maltolate.

ProNeura Continuous Drug Delivery Technology

Our ProNeura continuous drug delivery system consists of a small, solid rod made from a mixture of ethylene-vinyl acetate (EVA) and a drug substance. The resulting product is a solid matrix that is placed subcutaneously, normally in the upper arm in a simple 15-minute office procedure, and is removed in a similar manner at the end of the treatment period. The drug substance is released slowly, at continuous levels, through the process of diffusion. This results in a constant rate of release similar to intravenous administration. We believe that such long-term, linear release characteristics are desirable by avoiding peak and trough level dosing that poses problems for many CNS and other therapeutic agents.

In addition to Probuphine, which is our first product in clinical testing to utilize our proprietary ProNeura long term drug delivery technology, we are developing our ProNeura sustained drug delivery technology for other potential treatment applications in which conventional treatment is limited by variability in blood drug levels and poor patient compliance. ProNeura technology was developed to address the need for a simple, practical method to achieve continuous long-term drug delivery, and potentially can provide controlled drug release on an outpatient basis over extended periods up to 6 - 12 months.

Sponsored Research and License Agreements

We are a party to several agreements with research institutions, companies, universities and other entities for the performance of research and development activities and for the acquisition of licenses relating to such activities.

Iloperidone

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In January 2000, we entered into a sublicense agreement with Schering granting Schering exclusive worldwide commercialization rights to Spheramine. Under the agreement, we will collaborate with Schering on manufacturing and clinical development of cell therapy for the treatment of Parkinson's disease. We will receive funding for development activities, as well as potential reimbursement of certain prior research and development expenses. Schering will fully fund, and manage in collaboration with us, all future pilot and pivotal clinical studies, and manufacturing and development activities. Schering will pay us a royalty on net sales of Spheramine. Schering may terminate this sublicense for any reason by providing us 90 days notice in advance.

DITPA

In October 2003, through the acquisition of Developmental Therapeutics, Inc. (DTI), we acquired an exclusive worldwide license to an issued U.S. patent and pending international patent applications covering DITPA. Under this license agreement, we made an initial stock payment of 1,187,500 shares of our common stock and a cash payment of \$171,250 to The University of Arizona, the licensor of the technology, and will also make an additional payment of 712,500 shares of our common stock upon the achievement of positive pivotal study results or certain other substantial milestones within five years. A cash payment of \$102,750 or, alternatively, an additional payment of 37,500 shares of our common stock, will also be made to the licensor of the technology upon achievement of such study results or such other substantial milestones within five years. Also under this agreement, we are required to make royalty payments to the licensor based on net sales of products and processes incorporating the licensed technology, subject to minimum annual amounts commencing in the first year following the commercial sale of the product, as well as a percentage of any income derived from any sublicense of the licensed technology. In addition, we are required to make milestone payments to the licensor upon the achievement of certain clinical or regulatory milestones.

Gallium Complexes

In August 2000, through the acquisition of GeoMed, Inc., we acquired an exclusive worldwide license to make, use and sell products developed under the patent rights to the compositions and application of gallium complexes. Under this license agreement, we are required to make an annual license payment to Dr. Lawrence Bernstein, technology inventor, of \$50,000, as well as royalty payments based on future net sales of products and processes incorporating the licensed technology. We must also pay all costs and expenses incurred in patent prosecution and maintenance.

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ProNeura Long-term Drug Delivery System

In October 1995, we acquired from the Massachusetts Institute of Technology (MIT) an exclusive worldwide license to certain U.S. and foreign patents relating to the long-term drug delivery system. The exclusive nature of the MIT license is subject to certain conditions regarding timely performance of product development activities. We must also satisfy certain other usual terms and conditions set forth in the MIT license in order to retain our license rights, including payments of royalties based on sales of products and processes incorporating the licensed technology, as well as a percentage of income derived from sublicenses of the licensed technology.

Patents and Proprietary Rights

We have obtained rights to certain patents and patent applications relating to our proposed products and may, in the future, seek rights from third parties to additional patents and patent applications. We also rely on trade secrets and proprietary know-how, which we seek to protect, in part, by confidentiality agreements with employees, consultants, advisors, and others. For risks we face with respect to patents and proprietary rights, see Risk Factors. We may be unable to protect our patents and proprietary rights.

Iloperidone

We hold a license from Aventis under one issued U.S. patent and certain foreign patents relating to iloperidone and its methods of use. Our license is exclusive for use in the treatment of psychiatric disorders, psychotic disorders and analgesia. The term of the U.S. patent that covers certain aspects of our iloperidone product runs through 2011. This does not include possible term extensions. Prosecution of various divisional and continuation applications and their foreign counterparts continues satisfactorily, although it is uncertain whether additional patents will be granted.

Spheramine and Other Cell Therapy Products

We are the exclusive licensee under a license agreement with NYU of certain U.S. and foreign

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Gallium Complexes

We are the exclusive licensee under the license agreement with Dr. Lawrence Bernstein of certain U.S. and foreign patents and patent applications relating to the gallium complexes. 10 U.S. patents and several foreign patents have issued that cover pharmaceutical compositions and methods of use for gallium complexes. Prosecution of other U.S. and foreign patent applications relating to this technology continues satisfactorily, although it is uncertain whether additional patents will be granted. Patents in these families have patent terms running through 2009. This date does not include any possible patent term extensions available under 35 U.S.C. § 156. We are also the exclusive licensee under a license agreement with Ohio State University of certain issued U.S. and foreign patents related to the use of gallium compounds to treat rheumatoid arthritis. The U.S. patent term runs through 2009. In addition, we are licensees of certain issued U.S. and foreign patents and patent applications licensed from the University of Iowa, relating to methods of use to inhibit the growth of *P. aeruginosa*, and to treat infections caused by intracellular pathogens and pathogens causing chronic pulmonary infections, and human immunodeficiency virus infections. The two issued U.S. patents have terms running through 2019. We have also filed additional patent applications covering the use of gallium complexes in treating infection by intracellular prokaryotes and DNA viruses, treating inflammatory arthritis, and treatment and prevention of adverse liver conditions.

ProNeura Long-term Drug Delivery System

We are the exclusive licensee under the MIT license to three U.S. patents relating to a long-term drug delivery system, with patent terms running through 2007 and 2009, and certain European patents with patent terms running through 2008 and 2010. These dates do not include possible term extensions available under 35 U.S.C. § 156. Three additional patent applications have been filed which incorporate the use of specific compounds with the ProNeura technology, including an application related to Probuphine.

Competition

The pharmaceutical and biotechnology industries are characterized by rapidly evolving technology and intense competition. Many companies of all sizes, including major

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Spheramine

Several new treatments for Parkinson's disease are in pre-clinical and clinical development. In addition, several public and private companies, including StemCells, Inc., are actively pursuing alternative cell transplant technologies. Deep brain stimulation, also known as subthalamic stimulation is also a competing therapy for patients with advanced Parkinson's disease. The FDA has approved a stimulator device (Activa) manufactured by Medtronic, Inc., which is marketed in the U.S. We believe Spheramine may have potential competitive advantages to this therapy.

DITPA

Several other companies which are currently marketing drugs such as beta blockers, ace inhibitors and inotropes, which may be used for the treatment of heart failure. These companies include Abbott, AstraZeneca, Aventis, Johnson & Johnson, Pfizer and Sanofi-Synthelabo. In addition, companies such as Bristol-Myers Squibb, Merck and OSI Pharmaceuticals are developing new drugs which may be used to treat heart failure. Although DITPA represents a potential new class of agents for the treatment of CHF, these products may compete with DITPA.

Statins, alone or in combination, form the mainstay of therapy of hyperlipidemia. Several large pharmaceutical companies such as Pfizer, Merck, Astra Zeneca, Schering Plough and Bristol Myers Squibb currently market statins. Next generations of statins are also under development by companies such as Novartis and Bristol Myers Squibb. In addition, several smaller companies (e.g. Metabasis Therapeutics, Forbes Medi Tech and Avant Immunotherapeutics) are also attempting to develop novel therapeutics for this indication.

Gallium Complexes

Intravenously administered gallium nitrate is approved to treat hypercalcemia related to malignancy and may have potential for treatment of certain cancers. Other intravenous products, including the bisphosphonates, are available or are in development in the U

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Human clinical trials on drugs are typically conducted in three sequential phases, although the phases may overlap. Phase I trials typically consist of testing the product in a small number of healthy volunteers or patients, primarily for safety in one or more doses. During Phase II, in addition to safety, dose selection and efficacy of the product is evaluated in up to several hundred patients and sometimes more. Phase III trials typically involve additional testing for safety and confirmation of efficacy in an expanded patient population at multiple test sites. The FDA may order the temporary or permanent discontinuation of a clinical trial at any time.

The results of the pre-clinical and clinical testing on new drugs, if successful, are submitted to the FDA in the form of a New Drug Application, or NDA. The NDA approval process requires substantial time and effort and there can be no assurance that any approval will be granted on a timely basis, if at all. The FDA may refuse to approve an NDA if applicable regulatory requirements are not satisfied. Product approvals, if granted, may be withdrawn if compliance with regulatory standards is not maintained or problems occur following initial marketing.

The FDA may also require post-marketing testing and surveillance of approved products, or place other conditions on their approvals. These requirements could cause it to be more difficult or expensive to sell the products, and could therefore restrict the commercial applications of such products. Product approvals may be withdrawn if compliance with regulatory standards is not maintained or if problems occur following initial marketing. With respect to patented products or technologies, delays imposed by the governmental approval process may materially reduce the period during which we will have the exclusive right to exploit such technologies.

We believe we are in compliance with all material applicable regulatory requirements. However, see **Risk Factors** We must comply with extensive government regulations for additional risks we face regarding regulatory requirements and compliance.

Foreign Regulatory Issues

Sales of pharmaceutical products outside the United States are subject to foreign regulatory requirements that vary widely from country to country. Whether or not FDA approval has been obtained, approval of a product by a comparable regulatory authority of a foreign country must generally be obtained prior to the commencement of marketing in that country. Although the time required to obtain such approval may be longer or shorter than that required for FDA approval, the requirements for FDA approval are among the most detailed in the world and FDA approval generally takes longer than foreign regulatory approvals.

Employees

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Item 1A. Risk Factors

Our business is subject to numerous risks.

We have a history of operating losses and may never be profitable.

From our inception through December 31, 2005, we had an accumulated deficit of approximately \$208.2 million. We will continue to incur losses for the foreseeable future as a result of the various costs associated with our research, development, financial, administrative, regulatory and management activities. We may never achieve or sustain profitability.

Our products are at various stages of development and may not be successfully developed or commercialized.

We do not currently have any products being sold on the commercial market. Our proposed products are at various stages of development, but all will require significant further capital expenditures, development, testing, and regulatory clearances prior to commercialization. Of the large number of drugs in development, only a small percentage successfully complete the U.S. Food and Drug Administration (FDA) regulatory approval process and are commercialized. We are subject to the risk that some or all of our proposed products:

will be found to be ineffective or unsafe;

will not receive necessary regulatory clearances;

will be unable to get to market in a timely manner;

will not be capable of being produced in commercial quantities at reasonable costs;

will not be successfully marketed; or

will not be widely accepted by the physician community.

To date, we have experienced setbacks in some of our product development efforts. The results of a study evaluating the EKG profile of patients taking iloperidone, for example, found that iloperidone appeared

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We face many uncertainties relating to our human clinical trial strategy and results.

In order to obtain the regulatory approvals that we need to commercialize any of our product candidates, we must demonstrate that each product candidate is safe and effective for use in humans for each target indication. The results of preclinical and Phase I and Phase II clinical studies are not necessarily indicative of whether a product will demonstrate safety and efficacy in large patient populations. Although two of our product candidates have reached Phase III human clinical trials, results from the studies have not supported a regulatory filing. Several other product candidates are currently advancing into Phase II human clinical trials. We may not be able to demonstrate that any of our product candidates will be safe or effective in advanced trials that involve larger numbers of patients. Clinical trials are subject to oversight by institutional review boards and the FDA and:

must be conducted in conformance with the FDA's good laboratory practice regulations;

must meet requirements for institutional review board oversight;

must meet requirements for informed consent;

must meet requirements for good clinical practices;

are subject to continuing FDA oversight; and

may require large numbers of test subjects.

As described above in , Our products are at various stages of development and may not be successfully developed or commercialized, our product development programs have in the past been and may in the future be curtailed, redirected or eliminated at any time for some or all of the following reasons:

unanticipated, negative or ambiguous results;

undesirable side effects which delay or extend the trials;

our inability to locate, recruit and qualify a sufficient number of patients for our trials;

regulatory delays or other regulatory actions;

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reevaluation of our clinical development strategy.

Accordingly, our clinical trials may not proceed as anticipated or otherwise adequately support our applications for regulatory approval.

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We face risks associated with clinical trial liability claims in the event that the use or misuse of our product candidates results in personal injury or death.

We face an inherent risk of clinical trial liability claims in the event that the use or misuse of our product candidates results in personal injury or death. Our clinical liability insurance coverage may not be sufficient to cover claims that may be made against us. Any claims against us, regardless of their merit, could severely harm our financial condition, strain our management and other resources or destroy the prospects for commercialization of the product which is the subject of any such claim.

We may be unable to protect our patents and proprietary rights.

Our future success will depend to a significant extent on our ability to:

obtain and keep patent protection for our products and technologies on an international basis;

enforce our patents to prevent others from using our inventions;

maintain and prevent others from using our trade secrets; and

operate and commercialize products without infringing on the patents or proprietary rights of others.

We cannot assure you that our patent rights will afford any competitive advantages, and these rights may be challenged or circumvented by third parties. Further, patents may not be issued on any of our pending patent applications in the U.S. or abroad. Because of the extensive time required for development, testing and regulatory review of a potential product, it is possible that before a potential product can be commercialized, any related patent may expire or remain in existence for only a short period following commercialization, reducing or eliminating any advantage of the patent. If we sue others for infringing our patents, a court may determine that such patents are invalid or unenforceable. Even if the validity of our patent rights is upheld by a court, a court may not prevent the alleged infringement of our patent rights on the grounds that such activity is not covered by our patent claims.

In addition, third parties may sue us for infringing their patents. In the event of a successful claim of infringement against us, we may be required

management from other important tasks.

As is commonplace in the biotechnology and pharmaceutical industry, we employ individuals who were previously employed at other biotechnology or pharmaceutical companies, including our competitors or potential competitors. To the extent our employees are involved in research areas which are similar to those areas in which they were involved at their former employers, we may be subject to claims that such employees and/or we have inadvertently or otherwise used or disclosed the alleged trade secrets or other proprietary information of the former employers. Litigation may be necessary to defend against such claims, which could result in substantial costs and be a distraction to management even if we are successful in defending such claims.

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price, the more shares of common stock we will have to issue under the Standby Equity Distribution Agreement to draw-down the full amount. If our stock price is lower, then our existing stockholders would experience greater dilution.

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Item 2. Properties

We have a five-year operating lease, expiring in June 2010, for approximately 22,595 square feet of office space in South San Francisco, California. We also have a five-year lease, expiring in January 2007, for approximately 4,200 square feet of office and laboratory space in Somerville, New Jersey.

Item 3. Legal Proceedings

In March 2005, Dr. Bernard Sabel initiated an appraisal proceeding in the Court of Chancery of the State of Delaware relating to the merger of Titan's subsidiary ProNeura, Inc. into Titan. The complaint indicates that Mr. Sabel wants the court to appraise the value of the 108,800 shares of the common stock of ProNeura owned by him. The complaint does not specify an amount that Mr. Sabel considers the fair value of the shares. Discovery is proceeding in connection with this appraisal proceeding.

Item 4. Submission of Matters to a Vote of Security Holders

Not applicable

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Our common stock trades on the American Stock Exchange under the symbol TTP. The table below sets forth the high and low sales prices of our common stock as reported by the American Stock Exchange for the periods indicated.

	High	Low
Fiscal Year Ended December 31, 2005:		
First Quarter	\$ 3.24	\$ 2.20
Second Quarter	\$ 2.47	\$ 1.80
Third Quarter	\$ 2.31	\$ 1.73
Fourth Quarter	\$ 1.89	\$ 1.19
Fiscal Year Ended December 31, 2004:		
First Quarter	\$ 5.89	\$ 2.80
Second Quarter	\$ 5.15	\$ 2.43
Third Quarter	\$ 2.84	\$ 1.80
Fourth Quarter	\$ 3.39	\$ 1.94

(b) Approximate Number of Equity Security Holders

The number of record holders of our common stock as of March 1, 2006 was approximately 154. Based on the last ADP search, we believe there are in excess of 10,000 beneficial holders of our common stock.

(c) Dividends

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The selected financial data presented below summarizes certain financial data which has been derived from and should be read in conjunction with our consolidated financial statements and notes thereto included in the section beginning on page F-1. See also Item 7. Management's Discussion and Analysis of Financial Condition and Results of Operations.

	2005	Year Ended December 31, (in thousands, except per share data)				2001
		2004	2003	2002		
Statements of Operations Data:						
Total revenue(1)	\$ 89	\$ 31	\$ 89	\$ 2,892	\$ 4,572	
Operating expenses:						
Research and development	17,770	20,415	22,258	29,819	23,339	
Acquired/in-process research and development(2)		759	3,896			
General and administrative	5,370	5,237	5,109	5,076	5,383	
Other income, net	589	376	1,285	3,821	6,686	
Net loss	\$ (22,462)	\$ (26,004)	\$ (29,889)	\$ (28,		

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Iloperidone	Schizophrenia, psychosis	Phase III	Vanda Pharmaceuticals, Inc.
Spheramine	Parkinson s disease	Phase IIb	Schering AG
DITPA	Congestive heart failure	Phase II	Titan
DITPA	Hyperlipidemia	Phase II	Titan
Gallium maltolate	Bone related disease, chronic bacterial infections, cancer	Phase I	Titan

Table of Contents**Summary Compensation Table**

Name and Principal Position	Year	Annual Compensation Salary	Bonus	Other Compensation
Louis R. Bucalo, M.D. President and Chief Executive Officer	2005	\$ 366,325		
	2004	\$ 357,042		
	2003	\$ 348,038		
Sunil Bhonsle Executive Vice President and Chief Operating Officer	2005	\$ 279,208		
	2004	\$ 272,125		
	2003	\$ 265,276		
Richard C. Allen, Ph.D. Executive Vice President, Cell Therapy	2005	\$ 244,383		\$ 33,688(1)
	2004	\$ 238,200		
	2003	\$ 232,230		
Robert E. Farrell, J.D. Executive Vice President and Chief Financial Officer	2005	\$ 233,108		
	2004	\$ 227,217		
	2003	\$ 221,447		

(1)

Name	Number of Securities					
	Shares		Underlying		Value of Unexercised	
	Acquired on	Value	Unexercised Options at		in-the-Money	
			FY-End		Options at FY-End(1)	
	Exercise	Realized	Exercisable	Unexercisable	Exercisable	Unexercisable
Louis R. Bucalo		\$	1,650,272	116,147	\$ 37,167	\$
Sunil Bhonsle		\$	747,905	70,000	\$	\$
Richard C. Allen	83,523	\$ 94,911	516,161	30,000	\$	\$
Robert E. Farrell		\$	317,052	45,000	\$	\$

(1) Based on the fair market value of our common stock at year-end, \$1.43 per share, less the exercise price payable for such shares.

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Equity Compensation Plan Information

The following table sets forth aggregate information regarding our equity compensation plans in effect as of December 31, 2005:

Plan category	Number of securities to be issued upon exercise of outstanding options	Weighted- average exercise price of outstanding options	Number of securities remaining available for future issuance under equity compensation plans
		(b)	(c)
Equity compensation plans approved by security holders	(a) 4,471,808	\$ 7.52	1,934,646
Equity compensation plans not approved by security holders(1)(2)	2,026,379	\$ 7.62	331,387
Total	6,498,187	\$ 7.56	2,266,033

- (1) In August 2002, we amended our 2001 Employee Non-Qualified Stock Option Plan. Pursuant to this amendment, a total of 1,750,000 shares of common stock were reserved and authorized for issuance for option grants to employees and consultants who are not officers or directors of Titan.
- (2) In November 1999 and in connection with the redemption of warrants, we granted 813,000 non-qualified stock options outside of our stock option plans to our executive officers, at an exercise price of \$12.69, vesting equally over 36 months from the date of grant.

Item 13. Certain Relationships and Related Transactions.

Not applicable

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Audit Fees This category includes aggregate fees billed by our independent auditors for the audit of Titan's annual financial statements, audit of management's assessment and effectiveness of internal controls over financial reporting, review of financial statements included in our quarterly reports on Form 10-Q and services that are normally provided by the auditor in connection with statutory and regulatory filings for those fiscal years.

Audit-Related Fees This category consists of services by our independent auditors that, including accounting consultations on transaction related matters, are reasonably related to the performance of the audit or review of Titan's financial statements and are not reported above under Audit Fees.

whether the service places the auditor in the position of auditing his or her own work;

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whether the service results in the auditor acting as management or an employee of the Company; and

whether the service places the auditor in a position of being an advocate for the Company.

10.23* Employment Agreement between Registrant and Robert E. Farrell dated August 9, 1996.(4)

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- (6) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 1997.
- (7) Incorporated by reference from the Registrant's Definitive Proxy Statement filed on July 28, 2000.
- (8) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 1999.
- (9) Incorporated by reference from the Registrant's Quarterly Report on Form 10-Q for the period ended September 30, 2000.
- (10) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 2001.
- (11) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 2002.
- (12) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 2003.
- (13) Incorporated by reference from the Registrant's Annual Report on Form 10-K for the year ended December 31, 2004.
- (14) Incorporated by reference from the Registrant's Current Report on Form 8-K dated September 28, 2005.

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TITAN PHARMACEUTICALS, INC.

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We have also audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States), the consolidated balance sheet of Titan Pharmaceuticals, Inc. and its subsidiaries as of December 31, 2005, and the related consolidated statements of operations, stockholders' equity, and cash flows for the year then ended and our report dated February 27, 2006 expressed an unqualified opinion thereon.

/s/ Odenberg Ullakko Muranishi & Co. LLP

San Francisco, California

February 27, 2006

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REPORT OF ERNST & YOUNG LLP, INDEPENDENT REGISTERED PUBLIC ACCOUNTING FIRM

The Board of Directors and Stockholders

Titan Pharmaceuticals, Inc.

We have audited the accompanying consolidated statements of operations, stockholders' equity, and cash flows of Titan Pharmaceuticals, Inc. for the year ended December 31, 2003. 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 audit.

We conducted our audit 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. An audit also includes assessing the accounting principles used and significant estimates made by management, as well as evaluating the overall financial statement presentation. We believe that our audit provides a reasonable basis for our opinion.

In our opinion, the consolidated financial statements referred to above present fairly, in all material respects, the consolidated results of operations and cash flows of Titan Pharmaceuticals, Inc. for the year ended December 31, 2003, in conformity with U.S. generally accepted accounting principles.

/s/ Ernst & Young LLP

Palo Alto, California

February 20, 2004

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Table of Contents**TITAN PHARMACEUTICALS, INC.****CONSOLIDATED STATEMENTS OF OPERATIONS**

	Years ended December 31,		
	2005	2004	2003
	(in thousands, except per share amount)		
Revenue:			
Contract revenue	\$	\$	\$
License revenue	89	31	61
Total revenue	89	31	89
Operating expenses:			
Research and development	17,770	20,415	22,258
Acquired research and development		759	3,896
General and administrative	5,370	5,237	5,109
Total operating expenses	23,140	26,411	31,263
Loss from operations	(23,051)	(26,380)	(31,174)
Other income (expense):			
Interest income	570	673	1,278
Other income (expense)	19	(297)	7
Other income, net	589	376	1,285
Net loss	\$ (22,4		

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Basic and diluted net loss per share, as reported	\$ (0.69)	\$ (0.83)	\$ (1.07)
Pro forma basic and diluted net loss per share	\$ (0.72)	\$ (0.86)	\$ (1.14)

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In 1997, we entered into an exclusive license agreement with Aventis SA (formerly Hoechst Marion Roussel, Inc.). The agreement gave us a worldwide license to the patent rights and know-how related to the antipsychotic agent iloperidone, including the ability to develop, use, sublicense, manufacture and sell products

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Table of Contents**15. Quarterly Financial Data (Unaudited)**

	First	Second	Third	Fourth
	Quarter	Quarter	Quarter	Quarter
	(in thousands, except per share amount)			
2005				
Total revenue	\$ 14	13	1	\$ 61
Net loss	\$ (6,296)	\$ (5,742)	\$ (6,378)	\$ (4,046)
Basic and diluted net loss per share	\$ (0.19)	\$ (0.18)	\$ (0.20)	\$ (0.12)
2004				
Total revenue	\$ 1			\$ 30
Net loss	\$ (6,381)	\$ (5,555)	\$ (6,270)	\$ (7,798)
Basic and diluted net loss per share	\$ (0.22)	\$ (0.17)	\$ (0.20)	\$ (0.24)

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Konrad M. Weis, Ph.D.

/s/ ROBERT E. FARRELL

Robert E. Farrell, J.D.

Executive Vice President and Chief
Financial Officer (principal financial and
accounting officer)

March 14, 2006

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