

MEDICINES CO /DE  
Form 10-K  
March 01, 2007

# UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

## Form 10-K

(Mark One)

- ANNUAL REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934  
For the fiscal year ended: December 31, 2006
- or
- TRANSITION REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934  
For the transition period from \_\_\_\_\_ to \_\_\_\_\_.

Commission file number 000-31191

(Exact name of registrant as specified in its charter)

**Delaware**  
(State or other jurisdiction of  
incorporation or organization)  
**8 Campus Drive**  
**Parsippany, New Jersey**  
(Address of principal executive offices)

**04-3324394**  
(I.R.S. Employer  
Identification No.)  
**07054**  
(Zip Code)

Registrant's telephone number, including area code: (973) 656-1616

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 Per Share                                | NASDAQ Global Select Market               |
| <b>Securities registered pursuant to Section 12(g) of the Act: None</b> |   |

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 15(d) of the 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 such 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 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.

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Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, or a non-accelerated filer. See definition of 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 voting Common Stock held by non-affiliates of the registrant on June 30, 2006 was approximately \$985,386,079 based on the last reported sale price of the Common Stock on the Nasdaq National Market on June 30, 2006 of \$19.55 per share.

Number of shares of the registrant's class of Common Stock outstanding as of February 23, 2007: 51,561,027.

### **DOCUMENTS INCORPORATED BY REFERENCE**

The registrant intends to file a proxy statement pursuant to Regulation 14A within 120 days of the end of the fiscal year ended December 31, 2006. Portions of the proxy statement are incorporated herein by reference into the following parts of the Form 10-K:

Part III, Item 10, Directors, Executive Officers and Corporate Governance

Part III, Item 11, Executive Compensation;

Part III, Item 12, Security Ownership of Certain Beneficial Owners and Management and Related Stockholder Matters;

Part III, Item 13, Certain Relationships and Related Transactions, and Director Independence;

Part III, Item 14, Principal Accounting Fees and Services.

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**THE MEDICINES COMPANY**

**ANNUAL REPORT ON FORM 10-K**

**For the Fiscal Year Ended December 31, 2006**

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The Medicines Company® name and logo, Angiomax®, Angiox® and Cleviprex are either registered trademarks or trademarks of The Medicines Company in the United States and/or other countries. All other trademarks, service marks or tradenames appearing in this annual report on Form 10-K are the property of their respective owners. Except where otherwise indicated, or where the context may otherwise require, references to Angiomax in this annual report on Form 10-K mean Angiomax and Angiox, collectively.

This annual report on Form 10-K includes forward-looking statements within the meaning of Section 21E of the Securities Exchange Act of 1934, as amended, and Section 27A of the Securities Act of 1933, as amended. For this purpose, any statements contained herein regarding our strategy, future operations, financial position, future revenues, projected costs, prospects, plans and objectives of management, other than statements of historical facts, are forward-looking statements. The words anticipates, believes, estimates, expects, intends, may, plans, will, would and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. We cannot guarantee that we actually will achieve the plans, intentions or expectations expressed or implied in our forward-looking statements. There are a number of important factors that could cause actual results, levels of activity, performance or events to differ materially from those expressed or implied in the forward-looking statements we make. These important factors include our critical accounting estimates described in Item 7 of this annual report and the factors set forth under the caption Risk Factors in Item 1A of this annual report. Although we may elect to update forward-looking statements in the future, we specifically disclaim any obligation to do so, even if our estimates change, and readers should not rely on our forward-looking statements as representing our views as of any date subsequent to the date of this annual report.

**PART I**

**Item 1. Business**

**Our Company**

We are a pharmaceutical company committed to becoming a leading provider of innovative, cost effective acute care hospital products to the worldwide hospital marketplace. We have one marketed product, Angiomax® (bivalirudin), and two products in late-stage development, Cleviprex (clevidipine) and cangrelor, that we believe share common features valued by hospital practitioners, including a high level of pharmacological specificity, potency and predictability. We believe that Angiomax and our two product candidates possess favorable attributes that competitive products do not provide and can satisfy unmet medical needs in the acute care hospital product market and offer improved performance to hospital businesses.

Our first acute care hospital product, Angiomax, is an intravenous direct thrombin inhibitor approved for use as an anticoagulant in combination with aspirin in patients undergoing percutaneous coronary interventions, or PCI. PCI, which we also refer to as coronary angioplasty, is conducted to clear restricted blood flow in arteries around the heart. We are evaluating Angiomax for use in additional patient populations, including, in a Phase III trial, patients presenting with acute coronary syndromes, or ACS. Our revenues to date have been generated principally from sales of Angiomax in the United States. We reported net revenue of \$214.0 million and net income of \$63.7 million for the year ended December 31, 2006.

We are currently conducting Phase III clinical trials of Cleviprex and cangrelor as potential acute care hospital products. The first of these, Cleviprex, is an intravenous drug that is intended for the reduction and control of blood pressure in intensive care patients who require rapid and precise control of blood pressure. Our second product candidate, cangrelor, is an intravenous antiplatelet agent that is intended to prevent platelet activation and aggregation in the clotting process to reduce the risk of clot formation.

We currently focus our commercial and product development resources primarily on the U.S. acute care hospital market, which includes a concentration of hospitals that conduct a large percentage of acute care procedures in the United States. Our core strategy is to acquire, develop and commercialize products that we believe will help hospitals treat patients more efficiently by improving the effectiveness and safety of treatment while reducing cost. We believe that our ability to identify market needs and generate meaningful clinical data by investing aggressively in research and development enables us to successfully pursue this strategy. Our research and development investments are designed to provide clinical data that measure whether products:

- are effective, safe and predictable;
- enable shorter periods of treatment;
- are easier to use than current products;
- reduce the length of hospital stay; and
- lower hospital costs.

We believe that products with these attributes positively impact patient care and are attractive to the decision-makers who comprise our current and potential customers, including hospital management, physicians, hospital pharmacists, nurses and other care staff.

We have worldwide license rights to each of our products, except for specified Asian countries with respect to cangrelor. In the European Union and other foreign jurisdictions, we sell Angiomax to third party distributors that market and distribute the product to hospitals. We are currently evaluating the most

effective manner to develop, market and sell each of our products outside the United States, including the potential establishment of our own infrastructure outside the United States.

## **Angiomax**

### *Overview*

We exclusively licensed Angiomax from Biogen Idec, Inc. in 1997 and we have exclusive license rights to develop, market and sell Angiomax worldwide. We received our first marketing approval from the U.S. Food and Drug Administration, or the FDA, in December 2000 for Angiomax for use as an anticoagulant in combination with aspirin in patients with unstable angina undergoing percutaneous transluminal coronary angioplasty, or PTCA. In June 2005, the FDA approved new prescribing information for Angiomax to also include patients undergoing PCI, in addition to those undergoing PTCA. In November 2005, the FDA approved the expansion of the label to include PCI patients with or at risk of heparin-induced thrombocytopenia and thrombosis syndrome, a complication of heparin administration known as HIT/HITTS that can result in limb amputation, renal failure and death. In September 2004, we received authorization from the European Commission to market Angiomax under the name Angiox® (bivalirudin) in the member states of the European Union for use as an anticoagulant in combination with aspirin in patients undergoing PCI. Angiomax is also approved for sale in Australia, Canada, New Zealand and countries in Central America, South America and the Middle East for indications similar to those approved by the FDA.

We believe that Angiomax has the potential to replace heparin, an anticoagulant that historically has been used in the United States, in the treatment of arterial thrombosis, a condition involving the formation of blood clots in arteries. Arterial thrombosis is associated with life-threatening conditions, such as ischemic heart disease, peripheral vascular disease and stroke. There are three main areas of the hospital where heparin is used for acute treatment of arterial thrombosis: the cardiac catheterization laboratory, where coronary angioplasties are performed; the emergency department, where patients with ACS, including chest pain and heart attacks, are initially treated; and the operating room, where valve replacement surgery and coronary artery bypass graft surgery, or CABG surgery, a procedure in which surgeons bypass a blockage in the patient's artery by grafting a vein to the artery on both sides of the blockage to restore blood flow around the obstruction, are performed.

We have invested significantly in the development of clinical data on the clinical effects of Angiomax in the treatment of PCI and ACS patients. In almost all of our investigations to date, we have compared Angiomax to heparin or enoxaparin, a low-molecular weight heparin, which until relatively recently were the only injectable anticoagulants for use in coronary angioplasty, or combinations of drugs including heparin. In total, we have tested Angiomax against heparin or enoxaparin or combinations of drugs including heparin or enoxaparin in 12 comparative PCI and ACS trials. In the pivotal PCI and ACS trials, Angiomax use resulted in rates of complications, such as heart attack, also known as myocardial infarction, or MI, that were comparable to the comparator drugs in the trials while resulting in fewer bleeding events, including a reduction in the need for blood transfusion, as compared to the comparator drugs in the trials. In addition, in these trials, the therapeutic effects of Angiomax have been shown to be more predictable than heparin.

To date, we have concentrated our commercial efforts on replacing heparin in the cardiac catheterization laboratory, where the PCI procedures for which Angiomax is approved are performed. In evaluating our operating performance in the United States, we focus on use of Angiomax by existing hospital customers and penetration into new hospitals, both of which are critical elements of our ability to increase market share and revenue. We believe that Angiomax use has been growing consistently and that in the first half of 2006, Angiomax was used in approximately 33% of the coronary angioplasty procedures conducted in the United States.

We market Angiomax to interventional cardiology customers for its approved uses in PCI, including in patients with or at risk of HIT/HITTS. We market and sell Angiomax in the United States with a sales force of approximately 135 representatives and managers experienced in selling to hospital customers. In the European Union and other foreign jurisdictions, we sell Angiomax to third party-distributors that market and distribute the product to hospitals.

#### *Medical Need*

We are focused on developing Angiomax as an anticoagulation therapy for the cardiac catheterization laboratory, where coronary angioplasties are performed; the emergency department, where patients with ACS, including chest pain and heart attacks, are initially treated; and the operating room, where valve replacement surgery and CABG surgery are performed.

Coronary angioplasty procedures inherently increase the risk of clots forming in the coronary arteries or in other arteries of the body. Clots form as the body reacts to the manipulation of the artery as a result of, for example, the use of catheters and other devices in connection with the angioplasty procedure. Accordingly, anticoagulation therapy is routinely administered to patients undergoing angioplasty to slow the clotting process and avoid unwanted clotting in the coronary artery and the potential growth of clots or the movement of a clot or portions of it downstream in the blood vessels to new sites.

ACS patients are subject to chest pain that results from a range of conditions, from unstable angina to acute myocardial infarction, or AMI. Unstable angina is caused most often by a rupture of plaque on an arterial wall that results in clot formation and ultimately decreases coronary blood flow but does not cause complete blockage of the artery, and is often medically managed in the emergency department with anticoagulation therapy. AMI occurs when coronary arteries, which supply blood to the heart, become completely blocked by a clot. AMI patients are routinely treated with anticoagulants and are increasingly undergoing angioplasty as a primary treatment to unblock clogged arteries.

Many of the most severe ACS patients undergo CABG surgery. A high level of anticoagulation is necessary in on-pump cardiac surgery during the period of cardiopulmonary bypass in order to prevent clots from forming in the machine used in such surgery or in the patient's cardiovascular system. Anticoagulation is also necessary in off-pump cardiac surgery to prevent clots from forming in the patient's cardiovascular system as a result of the manipulation of coronary arteries and the heart.

Anticoagulation therapy attempts to modify actions of the components in the blood system that lead to the formation of blood clots and is usually started immediately after a diagnosis of blood clots, or after risk factors for clotting are identified. Anticoagulation therapy has typically involved the use of drugs to inhibit one or more components of the clotting process, thereby reducing the risk of clot formation. When anticoagulation is insufficient in patients being treated for ischemic heart disease, the consequences can include death, AMI, or revascularization. Revascularization occurs when a treated artery is blocked again and requires re-opening. In addition, because anticoagulation therapy reduces clotting, it also may cause excessive bleeding.

#### *Clinical Development*

We have invested significantly in the development of clinical data on the mode of action and clinical effects of Angiomax in procedures including coronary angioplasty and stenting. In almost all of our investigations to date, we have compared Angiomax to heparin, which until relatively recently was the only injectable anticoagulant for use in coronary angioplasty, or combinations of drugs including heparin.

We conducted the REPLACE-2 clinical trial in 2001 and 2002 to evaluate Angiomax as the foundation anticoagulant for angioplasty within the context of modern therapeutic products and technologies, including coronary stents. The trial, which involved 6,002 patients in 233 clinical sites, was designed to evaluate whether the use of Angiomax with provisional use of glycoprotein IIb/IIIa, or GP

Iib/IIIa inhibitors, provides clinical outcomes relating to rates of ischemic and bleeding events that are the same as, or non-inferior to, low-dose weight-adjusted heparin plus GP Iib/IIIa inhibitors. These outcomes were designed to be assessed using formal statistical tests for non-inferiority. The primary objective of REPLACE-2 was to demonstrate non-inferiority to heparin plus a GP Iib/IIIa inhibitor for the quadruple composite endpoint of death, MI, urgent revascularization and major bleeding. The secondary objectives of REPLACE-2 included non-inferiority to heparin plus a GP Iib/IIIa inhibitor for a triple composite endpoint of death, MI and urgent revascularization. Based on 30-day, 6-month and 12-month patient follow-up results, Angiomax met all primary and secondary objectives for the study. In addition, major hemorrhage was reported significantly less frequently in the Angiomax with provisional GP Iib/IIIa inhibitor arm compared to the heparin plus a GP Iib/IIIa inhibitor arm.

We are seeking to expand the indications for which we may market Angiomax beyond angioplasty. In December 2005, we completed patient enrollment in a 13,819 patient Phase III trial, called ACUITY, studying Angiomax use in patients presenting to the emergency department with ACS. We were testing whether Angiomax use is safe and effective in ACS patients when it is first administered in the emergency department at a lower dose than that which is currently used in PCI patients. If an emergency department ACS patient subsequently underwent PCI, the dose was increased to provide the usual anticoagulation during the procedure. Outcomes were also measured among ACS patients not undergoing PCI, namely, those medically managed or those who underwent CABG surgery. All of these emergency department ACS patients were randomized into one of three arms: a control arm, Arm A, providing for the administration of heparin or enoxaparin with GP Iib/IIIa inhibitors; a second arm, Arm B, providing for the administration of Angiomax with planned use of GP Iib/IIIa inhibitors; and a third arm, Arm C, providing for the administration of Angiomax alone and permitting use of GP Iib/IIIa inhibitors only in selected cases involving ischemic events during PCI.

The 30-day patient results, which were presented in March 2006 by the principal investigators, showed that Angiomax met all primary and secondary pre-specified 30-day objectives for the ACUITY study. Specifically, in Arm C, the Angiomax monotherapy arm, Angiomax was effective and reduced the risk of major bleeding by 47% compared to the control arm, Arm A. In the Angiomax combination arm, Arm B, the Angiomax and GP Iib/IIIa combination was as effective, with similar reductions in bleeding, as the control arm. These results were published in The New England Journal of Medicine in November 2006.

The investigators continued to conduct the ACUITY trial in 2006 as they collected 12-month patient follow-up results following the completion of enrollment in December 2005. We expect these results to be reported by the investigators in the first half of 2007. If these results are favorable, we expect to submit a supplemental New Drug Application, or sNDA, to the FDA in the second half of 2007 seeking an expansion of the Angiomax product label to include the trial results and information about ACUITY maintenance dosing regimen starting in the emergency department.

In December 2005, we submitted an application to the FDA for approval to market Angiomax in patients with or at risk of HIT/HITTS undergoing cardiac surgery after completing four studies in our Phase III clinical development program in cardiac surgery. In October 2006, we received a non-approvable letter from the FDA in connection with this application. In the letter, the FDA stated that it did not consider the data that we submitted in support of the application adequate to support approval for this indication because the FDA did not consider the evidence used to qualify patients for inclusion in the trials that formed the basis for our application as a persuasive indicator for the risk of HIT/HITTS. We have indicated to the FDA that we are evaluating potential next steps.

We also are preparing to study Angiomax in the pediatric setting and are working with the FDA to develop an appropriate study program. We are also supporting an investigator-initiated trial called HORIZONS to study Angiomax use in adult acute myocardial infarction, or AMI, patients. HORIZONS is designed to evaluate whether Angiomax with provisional use of GPIIb/IIIa inhibitors is as safe and



effective as heparin or enoxaparin with planned use of GPIIb/IIIa inhibitors in AMI patients. We believe that additional studies provide an important service by helping us to provide contemporary clinical data about the use of Angiomax, to answer specific questions about the use of Angiomax posed by the marketplace and to give us direction for future clinical trials.

## **Cleviprex**

### *Overview*

We are developing Cleviprex, an intravenous drug, for the short-term reduction and control of severe acute blood pressure when oral therapy is not desirable or feasible. We exclusively licensed Cleviprex in March 2003 from AstraZeneca AB. Under the terms of our agreement with AstraZeneca, we have exclusive license rights to develop, market and sell Cleviprex worldwide.

Cleviprex belongs to a well-known class of drugs, called IV calcium channel blockers, which are used to control severe acute high blood pressure. Cleviprex acts by selectively relaxing the smooth muscle cells that line small arteries, resulting in widening of the artery and reduction of blood pressure. We believe that Cleviprex may address an unmet need for rapid, precise control of severe acute blood pressure based on attributes demonstrated in clinical trials to date, namely rapid, titratable onset of effect on blood pressure, simple preparation and administration, arterial selectivity and metabolism independent of organ function.

We plan to submit a New Drug Application, or NDA, to the FDA in the first half of 2007 for approval to market Cleviprex in patients receiving an intravenous antihypertensive in the acute care setting when oral therapy is not desirable or feasible. We expect to submit an application for marketing approval in selected foreign markets following submission of the NDA in the United States.

### *Medical Need*

Increases in blood pressure, which are sometimes rapid and acute, often occur to patients treated in an acute care setting. Hospital physicians administer intravenous drugs to control high blood pressure, or hypertension, because prolonged severe hypertension is known to cause organ dysfunction, particularly in the brain, kidneys and liver. Low blood pressure is also known to cause organ dysfunction. As a result, physicians attempt to control blood pressure within a range to enable safe treatment of the patient.

In the United States in 2005, an estimated 3.1 million patients were treated with intravenous antihypertensives in an acute care setting. Approximately 1.8 million of these patients were treated for cardiovascular-related disease, approximately 238,000 were treated for neurological conditions and the remainder were treated for other conditions. Of the patients treated with an intravenous antihypertensive for cardiovascular-related disease:

- approximately 1.4 million patients were administered intravenous antihypertensives in connection with medicine and cardiology conditions,
- approximately 390,000 patients were treated with intravenous antihypertensives in cardiac and vascular surgery.

We have asked cardiologists, neurologists and cardiovascular surgeons to describe the features of an intravenous antihypertensive that they value, along with the benefits they would expect to achieve. The features these physicians valued were:

- a rapid onset and offset of antihypertensive effect;
- selective activity on arteries, not veins;
- drug clearance independent of organ function;
- no direct effect on a patient's heart rate; and

- no decrease in the ability of the heart to pump blood.

In this survey, physicians believed that a drug that had these features would be expected to achieve the following benefits:

- the ability to increase and decrease drug effect rapidly;
- the ability to control blood pressure within a range;
- the ability to be safely administered in patients with kidney or liver dysfunction; and
- the ability to be safely administered in patients with severe cardiovascular disease.

We believe, based on clinical data for Cleviprex, that Cleviprex has the potential for use in the acute care setting due to its rapid antihypertensive onset and offset effect, its selective activity on arteries and its ability to be cleared from the body independent of organ function.

#### *Clinical Development*

We are developing Cleviprex in a clinical trial program comprised of six Phase III clinical trials. We completed two Phase III efficacy clinical trials of Cleviprex, which we refer to as the ESCAPE trials. The ESCAPE trials were designed to evaluate the effectiveness of Cleviprex in controlling blood pressure before and after cardiac surgery compared to a placebo control. Results in both trials met the protocol-defined objective, as measured by rates of treatment success, which was defined as at least 15% reduction in blood pressure without the need to use an alternate drug. We have also completed three Phase III clinical trials, which we refer to as the ECLIPSE trials, to evaluate the safety of Cleviprex in approximately 1,600 patients in comparison to sodium nitroprusside, nicardipine and nitroglycerine, three leading blood pressure reducing agents, before, during and following cardiac surgery. Results in all three trials met the protocol-defined objectives, which included primary objectives measured by the incidences of death, stroke, myocardial infarction and renal dysfunction, and secondary objectives involving the evaluation of adverse experiences with Cleviprex and its blood pressure lowering effect. We completed enrollment of patients in our sixth Phase III clinical trial of Cleviprex in January 2007. In this trial, which we refer to as the VELOCITY trial, we are evaluating Cleviprex in 100 patients with acute severe hypertension in an acute care setting. We expect to review the results with investigators in the second half of 2007.

In 2007, we also intend to conduct Phase IIIb trials of Cleviprex in neurology and cardiology, along with a health economics study, and to support an observational study and clinical survey on treatment practices for acute severe hypertension conducted by third party researchers.

#### **Cangrelor**

##### *Overview*

We are developing cangrelor, a short-acting injectable antiplatelet agent, to prevent platelet activation and aggregation in the clotting process. Cangrelor is designed to bind directly to the P2Y<sub>12</sub> receptor, a receptor that has been implicated in platelet aggregation. We exclusively licensed cangrelor in December 2003 from AstraZeneca. Under the terms of our agreement with AstraZeneca, we have exclusive license rights to develop, market and sell cangrelor worldwide, excluding Japan, China, Korea, Taiwan and Thailand.

We are developing cangrelor for potential use as an intravenous antiplatelet agent in the acute care setting of the cardiac catheterization laboratory. Based on input from our hospital customers in the cardiac catheterization laboratory, we believe that the acute care limitations of current oral therapy, such as clopidogrel, the leading oral P2Y<sub>12</sub> receptor antiplatelet agent, which include delayed onset, prolonged effect and unpredictable effect, have created a need for an intravenous platelet inhibitor that acts quickly, is cleared from the bloodstream rapidly and enables rapid recovery of platelet function. We believe that

pre-clinical studies and clinical studies conducted in approximately 500 patients to date suggest that cangrelor has these attributes. These clinical studies consist of Phase II clinical trials of cangrelor conducted by AstraZeneca prior to licensing this product candidate to us, and a 40-person clinical trial that we conducted in healthy volunteers to identify a dosing strategy for use of cangrelor. Specifically, these studies suggest that cangrelor may have:

- an immediate inhibitory effect on platelets;
- an inhibitory effect on platelet activation and aggregation that is proportional to the dose administered;
- inhibitory effects that are sustainable through the period of infusion;
- a plasma half-life of less than five minutes; and
- platelet function recovery in less than an hour.

#### *Medical Need*

In the cardiac catheterization laboratory, the use of antiplatelet agents that block platelet aggregation is considered important therapy because several studies of oral platelet inhibitors have demonstrated better patient outcomes when these agents are administered before coronary angioplasty.

There is currently no intravenous drug that primarily inhibits platelet activation. One of the leading oral platelet inhibitors is clopidogrel, which, like cangrelor, blocks the adenosine diphosphate receptor and is one of the class of platelet inhibitors referred to as thienopyridines. Clopidogrel is commonly administered at a high dose by giving patients four to eight oral tablets before the angioplasty procedure. This practice is known as pre-loading. Although clopidogrel pre-loading has been shown to improve ischemic outcomes in coronary angioplasty, there are several safety and convenience issues with the use of this agent in acute care practice:

- Clopidogrel requires liver metabolism to form the active agent; therefore, the pre-loading dose may require up to six hours to achieve its full effect.
- There does not appear to be a clear relationship between increased dosage and intended effect that is consistent across different patient groups.
- The inhibition of platelet function is irreversible, meaning the agent remains bound to receptors for the life of the platelet, which is typically ten days. This may impede patient management and treatment flexibility, especially if a patient needs cardiac surgery, which is usually delayed for days awaiting the generation and release of new platelets from the bone marrow.
- Oral agents are difficult to administer in the acute care setting because they need to be swallowed by patients that may have received light anesthesia. This is especially true when there is a need to swallow multiple tablets in a restricted period of time.

Based on input from our hospital customers in the cardiac catheterization laboratory, we believe that the combination of the reduction in ischemic events through platelet inhibition and the acute care limitations of current oral therapy has created a need for an injectable platelet inhibitor that acts quickly and is cleared from the bloodstream rapidly.

In the operating room, surgeons have not had an approved agent at their disposal to control thrombosis during surgery by inhibiting platelets. The antiplatelet agents currently approved for use in coronary angioplasty, GP IIb/IIIa inhibitors, oral thienopyridines and aspirin, have not demonstrated feasibility in surgery due to bleeding concerns or the necessity of long infusions. We believe that cangrelor has potential for use in surgery due to its rapid effect in inhibiting platelets and the rapid recovery of platelet function following cessation of administration.

#### *Clinical Development*

We are currently evaluating cangrelor's effectiveness and safety in preventing ischemic events in patients who require PCI in two separate Phase III clinical trials. The larger trial, which we refer to as the CHAMPION-PCI trial and for which we commenced enrollment in March 2006, is an approximately 9,000-patient trial designed to evaluate whether use of intravenous cangrelor is superior to use of eight 75mg clopidogrel tablets in patients undergoing PCI. The primary composite endpoint of the CHAMPION-PCI trial will measure death, MI, or urgent revascularization at 48 hours after the procedure. Patients in this trial may be treated with other intravenous anticoagulants, such as Angiomax, heparin and GP IIb/IIIa inhibitors, at the investigator's discretion.

The second trial, which we refer to as the CHAMPION-PLATFORM trial and for which we commenced enrollment in October 2006, compares cangrelor plus usual care to placebo plus usual care in patients who require PCI. We currently expect to enroll approximately 6,500 patients in this trial. This trial will measure the composite endpoint of death, MI, or urgent revascularization at 48 hours after the procedure. The FDA has recommended that we use an alternative statistical design for this trial. We plan to discuss the statistical design with the FDA before we finalize the design and before we complete the first interim analysis of this trial.

There were approximately 2,000 patients enrolled in CHAMPION-PCI and 150 patients enrolled in CHAMPION-PLATFORM at the end of 2006. We plan to enroll in excess of 8,000 patients in these trials in 2007 and expect to complete patient enrollment in both trials in 2008. If we complete these trials on a timely basis and the results of these trials are favorable, we anticipate making submissions for marketing approvals in the United States in 2008 and in the European Union and selected markets thereafter.

#### **Sales**

We sell Angiomax in the United States using a hospital sales force of approximately 135 sales representatives and managers. In the summer of 2005, we expanded our sales force by approximately 50% to allow us to more effectively serve our existing customers and penetrate new hospitals. We also reconfigured our sales territories. Since we implemented this new configuration, we believe that the structure has provided us broader and more frequent access to our targeted accounts, and we expect the structure to drive future sales of Angiomax. Our sales force targets, as potential hospital customers, those hospitals with cardiac catheterization laboratories in the United States that perform approximately 200 or more coronary angioplasties per year. These hospitals conduct a significant percentage of the total number of the coronary angioplasties performed each year in the United States.

In support of sales efforts, we focus our Angiomax marketing in the United States on interventional cardiologists and other key clinical decision-makers in cardiac catheterization laboratories. We believe our ability to deliver relevant, advanced and reliable service and information to our concentrated customer base provides us with significant market presence in the United States even in the highly competitive sub-segments of the hospital market such as cardiology.

If Angiomax is approved for use in ACS or other indications, we intend to market Angiomax for those indications in the United States by supplementing our commercial organization.

We plan to expand our U.S. sales force by between approximately 50 and 100 persons commencing three to six months before the potential launch of Cleviprex. We believe that an expanded sales force would enable us to sell Cleviprex efficiently to hospital customers, including to Angiomax customers, if Cleviprex is approved by the FDA for sale in the United States.

We currently sell Angiomax in the United States to a limited number of domestic medical and pharmaceutical wholesalers with distribution centers located throughout the country. These wholesalers then sell and ship Angiomax to hospitals. In the United States, AmerisourceBergen Drug Corporation, McKesson Corporation and Cardinal Health accounted for more than 88% of our revenues for the year ended December 31, 2006. We are in the process of modifying our distribution system in the United States. As part of this modification, we plan to sell Angiomax to a third party who will ship Angiomax directly to our hospital customers. We anticipate that we will begin selling Angiomax under this revised distribution system in the second quarter and we expect that it will enable us to reduce our distribution costs in 2007 and provide us with improved data.

We sell Angiomax outside of the United States to third party distributors that market and distribute the product to hospital customers as Angiox. Nycomed Danmark A/S is our exclusive distributor of Angiox in all countries of the European Union other than Greece, Portugal and Spain, including those countries that we believe have the highest potential for Angiox sales. Upon execution of our sales, marketing and distribution agreement with Nycomed in 2002, Nycomed paid us a distributor fee of \$1.5 million and purchased from us common stock having an aggregate purchase price of \$1.0 million. Nycomed paid us an additional \$2.5 million under the agreement in 2004, upon Angiox receiving marketing authorization in the European Union for use as an anticoagulant in patients undergoing PCI. Our agreement requires Nycomed to make minimum purchases of Angiox following regulatory approval of Angiox for marketing and the term of the agreement continues on a country-by-country basis until the later of (1) the expiration of the last patent (and any extensions thereof) covering the product in that country, or (2) 10 years after launch of the product in the country. Either party may terminate the agreement for material breach upon notice to the other party, if the breach is not cured within the applicable cure period. Nycomed is currently selling Angiox in those countries in which packaging approval and any required pricing and reimbursement approval have been obtained.

We have an agreement with Oryx which distributes Angiomax in Canada and agreements with affiliates of Grupo Ferrer Internacional for the distribution of Angiox in Greece, Portugal and Spain and for countries in Central America and South America. Grupo Ferrer is currently selling Angiomax in Spain, Greece and selected countries in South America. We also have agreements with other third parties for other countries outside of the United States.

Our revenues from international sales, including the amortization of licensing fees and milestone payments under our Nycomed agreement, were \$11.3 million in 2006, \$9.5 million in 2005 and \$8.6 million in 2004.

### **Manufacturing**

Our product manufacturing operation is comprised of professionals with expertise in pharmaceutical manufacturing development and logistics and supply chain management. These professionals oversee the manufacturing and distribution of our products by third party companies. We do not have a manufacturing infrastructure and do not intend to develop one. We are party to agreements with contract manufacturers to supply bulk drug substance for our products and with other third parties to formulate, package and distribute our products.

*Angiomax*

In December 1999, we entered into a commercial development and supply agreement with UCB Bioproducts S.A., which was recently acquired by Lonza Ltd. and is now known as Lonza Braine, S.A., for the development and supply of Angiomax bulk drug substance. Together with UCB Bioproducts, we developed a second generation chemical synthesis process to improve the economics of manufacturing Angiomax bulk drug substance. This process, which was approved by the FDA in May 2003, is known as the Chemilog process.

We have agreed that until September 2010 we will purchase a substantial portion of our Angiomax bulk drug substance manufactured using the Chemilog process from Lonza Braine at agreed upon prices. Under the terms of our agreement, following the expiration of the agreement, which automatically renews for consecutive three-year periods unless either party provides notice of non-renewal within one year prior to the expiration of the initial term or any renewal term, or if we terminate the agreement prior to its expiration, Lonza Braine has agreed to transfer the development technology to us. We may only terminate the agreement prior to its expiration in the event of a material breach by Lonza Braine. If we engage a third party to manufacture Angiomax for us using this technology prior to bivalirudin becoming a generic drug in the United States, we will be obligated to pay Lonza Braine a royalty based on the amount paid by us to the third party manufacturer. In July 2004, we entered into a development and supply agreement with Lonza Ltd. for the development of an alternative method of manufacture and commercial supply of Angiomax. In August 2006, we terminated this agreement with Lonza Ltd. after Lonza Ltd. failed to successfully produce an alternative method of manufacturing Angiomax.

We have developed reproducible analytical methods and processes for the fill-finish of Angiomax drug product which have been conducted by Ben Venue Laboratories, Inc.

*Cleviprex*

Prior to our acquisition of Cleviprex, AstraZeneca manufactured all clevidipine bulk drug. We have transferred the manufacturing process for bulk drug to Johnson Matthey Pharma Services for scale-up and manufacture for Phase III clinical trials and commercial supply.

We are also a party to an agreement with Hospira, Inc., pursuant to which Hospira has agreed to use its proprietary formulation technology for scale up and manufacture for all finished drug product for all Phase III clinical trials of Cleviprex and, if and when Cleviprex is approved by the FDA, commercial supply, and to carry out release testing and clinical packaging. Together with our contract manufacturers, we have completed manufacturing development work for Cleviprex. We believe our contract manufacturers have the capability to manufacture and package Cleviprex on a commercial scale appropriate for launch of the drug if and when Cleviprex is approved for sale by the FDA.

*Cangrelor*

Prior to our acquisition of cangrelor, AstraZeneca manufactured all cangrelor bulk drug which, after testing and release, has been used in clinical trials. Following our acquisition of cangrelor, we transferred the manufacturing process for bulk drug to Johnson Matthey Pharma Services for scale-up and manufacture for Phase III clinical trials and commercial supply.

We have also entered into an agreement with Baxter Pharmaceutical Solutions LLC, a division of Baxter Healthcare Corporation, pursuant to which Baxter has agreed to manufacture all cangrelor finished drug product for all Phase III clinical trials and to carry out release testing. We have not entered into an agreement for commercial supply of cangrelor finished drug product, although we believe our contract manufacturers have the capability to manufacture and package cangrelor on a commercial scale appropriate for launch of the drug when and if cangrelor is approved for sale by the FDA.

## Business Development

We intend to continue building our acute care franchise of hospital products by selectively acquiring and developing clinical compound candidates or products approved for marketing. We believe that we have proven capability in developing and commercializing in-licensed or acquired acute care drug candidates. We believe that products may be acquired from pharmaceutical companies in the process of refining their own product portfolios and companies seeking specialist development or commercial collaborations.

In evaluating product acquisition candidates, we plan to continue to seek products that have the potential to provide reasonable evidence of safety and efficacy, together with the potential to reduce a patient's hospital stay. Our acquisition strategy is to acquire global rights for development compounds wherever possible. In the United States, we may acquire approved products that can be marketed in hospitals by our commercial organization.

### *License Agreements*

*Biogen Idec.* In March 1997, we entered into an agreement with Biogen, Inc., a predecessor of Biogen Idec, Inc., for the license of the anticoagulant pharmaceutical bivalirudin, which we have developed and marketed as Angiomax. Under the terms of the agreement, we acquired exclusive worldwide rights to the technology, patents, trademarks, inventories and know-how related to Angiomax. In exchange for the license, we paid \$2.0 million on the closing date and are obligated to pay up to an additional \$8.0 million upon the first commercial sales of Angiomax for the treatment of AMI in the United States and Europe. In addition, we are obligated to pay royalties on sales of Angiomax and on any sublicense royalties on a country-by-country basis earned until the later of (1) 12 years after the date of the first commercial sales of the product in a country or (2) the date on which the product or its manufacture, use or sale is no longer covered by a valid claim of the licensed patent rights in such country. Under the terms of the agreement, the royalty rate due to Biogen Idec on sales increases with growth in annual sales of Angiomax. The agreement also stipulates that we use commercially reasonable efforts to meet certain milestones related to the development and commercialization of Angiomax, including expending at least \$20 million for certain developmental and commercialization activities, which we met in 1998. The license and rights under the agreement remain in force until our obligation to pay royalties ceases. Either party may terminate the agreement for material breach by the other party, if the material breach is not cured within 90 days after written notice. In addition, we may terminate the agreement for any reason upon 90 days prior written notice. Through December 31, 2006, we have incurred a total of approximately \$59.0 million in royalties relating to Angiomax under our agreement with Biogen Idec.

*AstraZeneca.* In March 2003, we acquired from AstraZeneca exclusive worldwide license rights to Cleviprex for all countries other than Japan. Under the terms of the agreement, we have the rights to the patents, trademarks, inventories and know-how related to Cleviprex. In May 2006, we amended our license agreement with AstraZeneca to provide us with exclusive license rights in Japan in exchange for an upfront payment. We acquired this license after having studied Cleviprex under the study and exclusive option agreement with AstraZeneca that we entered into in March 2002. In exchange for the license, we paid \$1.0 million in 2003 upon entering into the license and agreed to pay up to an additional \$5.0 million upon reaching certain regulatory milestones, including a payment of \$1.5 million within 30 days of the acceptance for review of the NDA we plan to submit to the FDA for Cleviprex. Under the terms of the license agreement, we will be obligated to pay royalties on a country-by-country basis on future annual sales of Cleviprex, and on any sublicense royalties earned, until the later of (1) the duration of the licensed patent rights which are necessary to manufacture, use or sell Cleviprex in a country or (2) ten years from our first commercial sale of Cleviprex in such country. The licenses and rights under the agreement remain in force on a country-by-country basis until we cease selling Cleviprex in such country or the agreement is otherwise terminated. We may terminate the agreement upon 30 days written notice, unless AstraZeneca,





within 20 days of having received our notice, requests that we enter into good faith discussions to redress our concerns. If we cannot reach a mutually agreeable solution with AstraZeneca within three months of the commencement of such discussions, we may then terminate the agreement upon 90 days written notice. Either party may terminate the agreement for material breach upon 60 days prior written notice, if the breach is not cured within such 60 days.

In December 2003, we acquired from AstraZeneca exclusive license rights to cangrelor for all countries other than Japan, China, Korea, Taiwan and Thailand. Under the terms of the agreement, we have the rights to the patents, trademarks, inventories and know-how related to cangrelor.

In exchange for the license, we paid in January 2004 an upfront payment upon entering into the license and agreed to make additional payments upon reaching certain regulatory milestones. Under the terms of the license agreement, we will be obligated to pay royalties on a country-by-country basis on future annual sales of cangrelor, and on any sublicense royalties earned, until the later of (1) the duration of the licensed patent rights which are necessary to manufacture, use or sell cangrelor in a country or (2) ten years from our first commercial sale of cangrelor in such country. The licenses and rights under the agreement remain in force on a country-by-country basis until we cease selling cangrelor in such country or the agreement is otherwise terminated. We may terminate the agreement upon 30 days written notice, unless AstraZeneca, within 20 days of having received our notice, requests that we enter into good faith discussions to redress our concerns. If we cannot reach a mutually agreeable solution with AstraZeneca within three months of the commencement of such discussions, we may then terminate the agreement upon 90 days written notice. Either party may terminate the agreement for material breach upon 60 days prior written notice, if the breach is not cured within such 60 days.

### **Competition**

The development and commercialization of new drugs is competitive, and we face competition from major pharmaceutical companies, specialty pharmaceutical companies and biotechnology companies worldwide. Our competitors may develop or license products or other novel technologies that are more effective, safer or less costly than any that have been or are being developed by us, or may obtain FDA approval for their products more rapidly than we may obtain approval for ours.

The acquisition or licensing of pharmaceutical products is a competitive area, and a number of more established companies, which have acknowledged strategies to license or acquire products, may have competitive advantages, as may emerging companies taking similar or different approaches to product acquisition. These established companies may have a competitive advantage over us due to their size, cash flows and institutional experience.

Many of our competitors have substantially greater financial, technical and human resources than we have. Additional mergers and acquisitions in the pharmaceutical industry may result in even more resources being concentrated in our competitors. Competition may increase further as a result of advances made in the commercial applicability of technologies and greater availability of capital for investment in these fields. Our success will be based in part on our ability to build and actively manage a portfolio of drugs that addresses unmet medical needs and creates value in patient therapy.

We compete, in the case of Angiomax, and expect to compete, in the cases of Cleviprex and cangrelor, on the basis of efficacy, safety, ease of administration and economic value compared to drugs used in current practice or currently being developed.

#### *Angiomax*

Due to the incidence and severity of cardiovascular diseases, the market for anticoagulant therapies is large and competition is intense. We are seeking to expand the indications for which we may market Angiomax. We are evaluating Angiomax for additional uses including patients presenting with ACS. There

are a number of anticoagulant therapies currently on the market, awaiting regulatory approval or in development for these uses.

*Direct thrombin inhibitors.* Direct thrombin inhibitors act directly on thrombin, inhibiting the action of thrombin in the clotting process. Because thrombin activates platelets, direct thrombin inhibitors also prevent platelet aggregation. Direct thrombin inhibitors include Angiomax, Refludan from Berlex Laboratories and Argatroban from GlaxoSmithKline, Encysive Pharmaceuticals Inc. and Mitsubishi Chemical Corp. Both Refludan and Argatroban are approved for use in the treatment of patients with HIT/HITTS. Argatroban is also approved for use in patients with HIT/HITTS undergoing angioplasty.

*Indirect thrombin inhibitors.* Heparin is widely used in patients with ischemic heart disease. Heparin is manufactured and distributed by a number of companies as a generic product. Low molecular weight heparin products include Lovenox from Sanofi-Aventis and Fragmin from Pfizer Inc. Very short molecules of heparin, called pentasaccharide sequences, include Arixtra from Sanofi-Aventis. Low molecular weight heparins have been approved for use in the treatment of patients with unstable angina and are being developed for use in angioplasty and vascular surgery. Arixtra has been approved for use in the treatment and prevention of deep vein thrombosis and is being developed for arterial thrombosis.

*Platelet inhibitors.* Platelet inhibitors, such as GP IIb/IIIa inhibitors, block the aggregation of platelets. GP IIb/IIIa inhibitors include ReoPro from Eli Lilly and Company and Johnson & Johnson/Centocor, Inc., Integrilin from Schering-Plough Corporation, and Aggrastat from Merck & Co., Inc. and MediCure Inc. ReoPro is approved and marketed for angioplasty in a broad range of patients. Integrilin is approved and marketed for angioplasty and for the management of ACS. Aggrastat is approved for the management of ACS.

Although platelet inhibitors may be complementary to Angiomax, Angiomax may compete with platelet inhibitors for the use of hospital financial resources. For example, many U.S. hospitals receive a fixed reimbursement amount per procedure for the angioplasties and other treatment therapies they perform. Because this amount is not based on the actual expenses the hospital incurs, hospitals may choose to use either Angiomax or a platelet inhibitor but not necessarily several of the drugs together.

#### *Cleviprex*

We expect that Cleviprex will compete with a variety of parenteral antihypertensive agents in the acute care setting, many of which are generic. We also expect Cleviprex to compete with nitroglycerine, which is used for a variety of purposes in the acute care setting. We believe that the most commonly administered drugs used specifically for their intravenous antihypertensive effects are sodium nitroprusside, labetalol and Cardene IV.

#### *Cangrelor*

We expect that cangrelor will compete with oral platelet inhibitors that are used in acute care settings such as clopidogrel from Bristol Meyers Squibb/Sanofi Pharmaceuticals Partnership as well as prasugrel, an anti-platelet agent currently being developed by Eli Lilly and Company and Sankyo Co., Ltd.

### **Patents, Proprietary Rights and Licenses**

Our success will depend in part on our ability to protect the products we acquire or license by obtaining and maintaining patent protection both in the United States and in other countries. We rely upon trade secrets, know-how, continuing technological innovations, contractual restrictions and licensing opportunities to develop and maintain our competitive position. We plan to prosecute and defend any patents or patent applications we acquire or license.

In all, as of February 23, 2007, we exclusively licensed six issued United States patents, rights relating to eight issued United States patents and a broadly filed portfolio of corresponding foreign patents and patent applications. We have not yet filed any independent patent applications. The U.S. patents licensed by us are currently set to expire at various dates, including in the case of the principal patent for Angiomax, March 2010, in the case of the principal patent for Cleviprex, January 2016, and in the case of the principal patent for cangrelor, February 2014.

We have exclusively licensed from Biogen patents and applications for patents covering Angiomax and Angiomax analogs and other novel anticoagulants as compositions of matter, and processes for using Angiomax and Angiomax analogs and other novel anticoagulants. We are responsible for prosecuting and maintaining patents and patent applications relating to Angiomax. We have exclusively licensed from AstraZeneca rights to patents and patent applications covering Cleviprex as a composition of matter and covering formulations and uses of Cleviprex, and rights to patents and patent applications covering cangrelor as a composition of matter, and covering formulations and uses of cangrelor. Under both licenses, AstraZeneca is responsible for prosecuting and maintaining these patents and patent applications relating to Cleviprex and cangrelor, and we are required to reimburse AstraZeneca for expenses it incurs in connection with the prosecution and maintenance of the patents and patent applications.

The patent positions of pharmaceutical and biotechnology firms like us can be uncertain and involve complex legal, scientific and factual questions. In addition, the coverage claimed in a patent application can be significantly reduced before the patent is issued. Consequently, we do not know whether any of the applications we acquire or license will result in the issuance of patents or, if any patents are issued, whether they will provide significant proprietary protection or will be challenged, circumvented or invalidated. Because unissued U.S. patent applications filed prior to November 29, 2000 and patent applications filed within the last 18 months are maintained in secrecy until patents issue, and since publication of discoveries in the scientific or patent literature often lags behind actual discoveries, we cannot be certain of the priority of inventions covered by pending patent applications. Moreover, we may have to participate in interference proceedings declared by the United States Patent and Trademark Office to determine priority of invention, or in opposition proceedings in a foreign patent office, either of which could result in substantial cost to us, even if the eventual outcome is favorable to us. There can be no assurance that the patents, if issued, would be held valid by a court of competent jurisdiction. An adverse outcome could subject us to significant liabilities to third parties, require disputed rights to be licensed from third parties or require us to cease using such technology.

The development of acute care hospital products is intensely competitive. A number of pharmaceutical companies, biotechnology companies, universities and research institutions have filed patent applications or received patents in this field. Some of these applications could be competitive with applications we have acquired or licensed, or could conflict in certain respects with claims made under such applications. Such conflict could result in a significant reduction of the coverage of the patents we have acquired or licensed, if issued, which would have a material adverse effect on our business, financial condition and results of operations. In addition, if patents are issued to other companies that contain competitive or conflicting claims and such claims are ultimately determined to be valid, no assurance can be given that we would be able to obtain licenses to these patents at a reasonable cost, or develop or obtain alternative technology.

We also rely on trade secret protection for our confidential and proprietary information. No assurance can be given that others will not independently develop substantially equivalent proprietary information and techniques or otherwise gain access to our trade secrets or disclose such technology, or that we can meaningfully protect our trade secrets. We have a number of trademarks that we consider important to our business. The Medicines Company® name and logo, Angiomax®, Angiox® and Cleviprex name and logo are either our registered trademarks or our trademarks in the United States and/or other countries.

It is our policy to require our employees, consultants, outside scientific collaborators, sponsored researchers and other advisors to execute confidentiality agreements upon the commencement of employment or consulting relationships with us. These agreements generally provide that all confidential information developed or made known to the individual during the course of the individual's relationship with us is to be kept confidential and not disclosed to third parties except in specific circumstances. In the case of employees and consultants, the agreements provide that all inventions conceived by the individual shall be our exclusive property. There can be no assurance, however, that these agreements will provide meaningful protection or adequate remedies for our trade secrets in the event of unauthorized use or disclosure of such information.

### **Government Regulation**

Government authorities in the United States and other countries extensively regulate the testing, manufacturing, labeling, safety advertising, promotion, storage, sales, distribution, export and marketing, among other things, of our products and product candidates. In the United States, the FDA regulates drugs under the Federal Food, Drug, and Cosmetic Act and implementing regulations. We cannot market a drug until we have submitted an application for marketing authorization to the FDA, and the FDA has approved it. Both before and after approval is obtained, violations of regulatory requirements may result in various adverse consequences, including, among other things, warning letters, the FDA's delay in approving or refusal to approve a product, product recall or seizure, suspension or withdrawal of an approved product from the market, interruption of production, operating restrictions, injunctions and the imposition of civil or criminal penalties. The steps required before a drug may be approved by the FDA and marketed in the United States include:

- pre-clinical laboratory tests, animal studies and formulation studies;
- submission to the FDA of an investigational new drug exemption, or IND, for human clinical testing, which must become effective before human clinical trials may begin;
- adequate and well-controlled clinical trials to establish the safety and efficacy of the drug for each indication;
- submission to the FDA of a new drug application, or an NDA;
- satisfactory completion of an FDA inspection of the manufacturing facility or facilities at which the drug is produced to assess compliance with current good manufacturing practices, or cGMP; and
- FDA review and approval of the NDA.

Pre-clinical tests include laboratory evaluations of product chemistry, toxicity and formulation, as well as animal studies. The results of the pre-clinical tests, together with manufacturing information and analytical data, are submitted to the FDA as part of an IND, which must become effective before human clinical trials may begin. An IND will automatically become effective 30 days after receipt by the FDA, unless before that time the FDA raises concerns or questions about issues such as the conduct of the trials as outlined in the IND. In such a case, the IND sponsor and the FDA must resolve any outstanding FDA concerns or questions before clinical trials can proceed. Submission of an IND does not necessarily result in the FDA allowing clinical trials to commence.

Clinical trials involve the administration of the investigational drug to human subjects under the supervision of qualified investigators. Clinical trials are conducted under protocols detailing the objectives of the study, the parameters to be used in monitoring subject safety, and the effectiveness criteria, or endpoints, to be evaluated. Each protocol must be submitted to the FDA as part of the IND and the FDA may or may not allow that trial to proceed.

Clinical trials typically are conducted in three sequential phases, but the phases may overlap or be combined. Each trial must be reviewed and approved by an independent Institutional Review Board before it can begin. Phase I usually involves the initial introduction of the investigational drug into people to evaluate its safety, dosage tolerance, pharmacodynamics, and, if possible, to gain an early indication of its effectiveness. Phase II usually involves trials in a limited patient population to:

- evaluate dosage tolerance and appropriate dosage;
- identify possible adverse effects and safety risks; and
- evaluate preliminarily the efficacy of the drug for specific indications.

Phase III trials usually further evaluate clinical efficacy and test further for safety by administering the drug in its final form in an expanded patient population. We cannot guarantee that Phase I, Phase II or Phase III testing will be completed successfully within any specified period of time, if at all. Furthermore, we or the FDA may suspend clinical trials at any time on various grounds, including a finding that the subjects or patients are being exposed to an unacceptable health risk.

Assuming successful completion of the required clinical testing, the results of the pre-clinical studies and of the clinical studies, together with other detailed information, including information on the manufacture and composition of the drug, are submitted to the FDA in the form of an NDA requesting approval to market the product for one or more indications. Before approving an application, the FDA usually will inspect the facility or the facilities at which the drug is manufactured, and will not approve the product unless cGMP compliance is satisfactory. If the FDA determines the application or manufacturing facilities are not acceptable, the FDA may outline the deficiencies in the submission and often will request additional testing or information. Notwithstanding the submission of any requested additional information, the FDA ultimately may decide that the application does not satisfy the regulatory criteria for approval. After approval, certain changes to the approved product, such as adding new indications, manufacturing changes, or additional labeling claims, are subject to further FDA review and approval before the changes can be implemented. The testing and approval process requires substantial time, effort and financial resources, and we cannot be sure that any approval will be granted on a timely basis, if at all. As a condition of approval of an application, the FDA may require postmarket testing and surveillance to monitor the drug's safety or efficacy.

After the FDA approves a product, we and our contract manufacturers must comply with a number of post-approval requirements. For example, holders of an approved NDA are required to report certain adverse reactions and production problems, if any, to the FDA, and to comply with certain requirements concerning advertising and promotional labeling for their products. Also, quality control and manufacturing procedures must continue to conform to cGMP after approval, and the FDA periodically inspects manufacturing facilities to assess compliance with cGMP. Accordingly, we and our contract manufacturers must continue to expend time, money, and effort to maintain compliance with cGMP and other aspects of regulatory compliance.

We use and will continue to use third party manufacturers to produce our products in clinical and commercial quantities, and we cannot be sure that future FDA inspections will not identify compliance issues 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 may result in restrictions on a product, manufacturer, or holder of an approved NDA, including withdrawal of the product from the market. Also, new government requirements may be established that could delay or prevent regulatory approval of our products under development.

We are also subject to foreign regulatory requirements governing human clinical trials and marketing approval for pharmaceutical products which we sell outside the United States. The requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement vary widely from country to

country. Whether or not we obtain FDA approval, we must obtain approval of a product by the comparable regulatory authorities of foreign countries before manufacturing or marketing the product in those countries. The approval process varies from country to country and the time required for these approvals may differ substantially from that required for FDA approval. Clinical trials in one country may not be accepted by other countries, and approval in one country may not result in approval in any other country. For clinical trials conducted outside the United States, the clinical stages generally are comparable to the phases of clinical development established by the FDA.

### **Research and Development**

Our research and development expenses totaled \$63.5 million in 2006, \$64.4 million in 2005 and \$49.3 million in 2004.

### **Employees**

We believe that our success depends greatly on our ability to identify, attract and retain capable employees. We have assembled a management team with significant experience in drug development and commercialization.

As of February 23, 2007, we employed 289 persons. Our employees are not represented by any collective bargaining unit, and we believe our relations with our employees are good.

### **Available Information**

Our Internet address is <http://www.themedicinescompany.com>. The contents of our website are not part of this annual report on Form 10-K, and our Internet address is included in this document as an inactive textual reference only. We make our annual reports on Form 10-K, quarterly reports on Form 10-Q, current reports on Form 8-K and all amendments to those reports available free of charge on our website as soon as reasonably practicable after we file such reports with, or furnish such reports to, the Securities and Exchange Commission, or SEC.

### **Item 1A. Risk Factors**

*Investing in our common stock involves a high degree of risk. You should carefully consider the risks and uncertainties described below in addition to the other information included or incorporated by reference in this annual report. If any of the following risks actually occur, our business, financial condition or results of operations would likely suffer. In that case, the trading price of our common stock could fall.*

#### **Risks Related to Our Financial Results**

*We have a history of net losses and may not maintain profitability on an annual basis*

Except for 2004 and 2006, we have incurred net losses on an annual basis since our inception. As of December 31, 2006, we had an accumulated deficit of approximately \$241.2 million. We expect to make substantial expenditures to further develop and commercialize our products, including costs and expenses associated with clinical trials, regulatory approvals and commercialization. Although we achieved profitability in 2004 and in 2006, and expect to be profitable in 2007, we were not profitable in 2005 and will likely need to generate significantly greater revenue in future periods to achieve and maintain profitability in light of our planned expenditures. We may not achieve profitability in future periods or at all, and we may not be able to maintain profitability for any substantial period of time. If we fail to achieve profitability or maintain profitability on a quarterly or annual basis within the time frame expected by investors or securities analysts, the market price of our common stock may decline.

***Our business is very dependent on the commercial success of Angiomax***

Angiomax is our only commercial product. We expect Angiomax will account for almost all of our revenue for at least 2007. The commercial success of Angiomax will depend upon:

- its continued acceptance by regulators, physicians, patients and other key decision-makers as a safe, therapeutic and cost-effective alternative to heparin and other products used in current practice or currently being developed;
- our ability to expand the indications for which we can market Angiomax and the clinical data we generate to support expansion of the product label; and
- the extent to which we and our international distributors are successful in marketing Angiomax.

We plan to continue in 2007 to seek to expand the indications for which we may market Angiomax. Even if we are successful in expanding the Angiomax label, we cannot assure you that the expanded label will result in higher revenue or income on a continuing basis. If Angiomax is not commercially successful, we will have to find additional sources of funding or curtail operations. As of December 31, 2006, our inventory was \$41.6 million. In addition, we have inventory-related purchase commitments to Lonza Braine totaling \$14.8 million during 2007 and \$4.3 million during 2008 for Angiomax bulk drug substance and \$2.0 million in remaining Angiomax-related filling, finishing and packaging commitments during 2007. If sales of Angiomax were to decline, we could be required to make an allowance for excess or obsolete inventory or increase our accrual for product returns.

***Our revenue has been substantially dependent on a limited number of domestic wholesalers and international distributors to which we sell Angiomax, and such revenue may fluctuate from quarter to quarter based on the buying patterns of these wholesalers and distribution partners and the levels of inventory they maintain***

We currently sell Angiomax to a limited number of national medical and pharmaceutical wholesalers with distribution centers located throughout the United States. Outside of the United States, we sell Angiomax to several international distributors. These wholesalers and distributors then sell Angiomax to hospitals. During the year ended December 31, 2006, revenue from the sale of Angiomax to our three largest U.S. wholesalers totaled approximately 88% of our net revenue and sales to one of our international distributors totaled approximately 3% of our net revenue. Our reliance on a small number of wholesalers and distributors could cause our revenue to fluctuate from quarter to quarter based on the buying patterns of these wholesalers and distributors, regardless of underlying hospital demand. For instance, because an order from Nycomed, one of our European distributors, was not recognized in the quarter ended March 31, 2006 due to a delay in Nycomed's acceptance of the order, our revenue for the first quarter of 2006 was reduced. Under our revised distribution model, our revenue from sales of Angiomax in the United States will be almost exclusively from sales to one third party beginning in the second quarter of 2007, and will continue to be subject to fluctuation from quarter to quarter based on the buying pattern of this third party.

If inventory levels at this third party or at our international distributors become too high, they may seek to reduce their inventory levels by reducing purchases from us. In 2005, we agreed with our largest wholesalers to enter into fee-for-service arrangements. As a result, we estimate that our three largest wholesalers reduced aggregate Angiomax inventory to an average of four to six weeks during the last two quarters of 2005 and the first quarter of 2006. In implementing the inventory reduction to reach this level during this period, we estimate that our three largest wholesalers reduced their aggregate inventories of Angiomax by approximately \$39.0 million which had an adverse effect on our revenue.

***Failure to achieve our revenue targets or raise additional funds in the future may require us to delay, reduce the scope of, or eliminate one or more of our planned activities***

We will need to generate significantly greater revenue to achieve and maintain profitability on an annual basis. The development of Angiomax for additional indications, the development of Cleviprex and cangrelor, including clinical trials, manufacturing development and regulatory approvals, and the acquisition and development of additional product candidates by us will require a commitment of substantial funds. Our future funding requirements, which may be significantly greater than we expect, will depend upon many factors, including:

- the extent to which Angiomax is commercially successful in the United States;
- the extent to which our international distributors, including Nycomed, are commercially successful;
- the progress, level and timing of our research and development activities related to our clinical trials with respect to Angiomax, Cleviprex and cangrelor;
- the cost and outcomes of regulatory submissions and reviews;
- the continuation or termination of third party manufacturing or sales and marketing arrangements;
- the size, cost and effectiveness of our sales and marketing programs;
- the status of competitive products;
- our ability to defend and enforce our intellectual property rights;
- our decision whether to establish an infrastructure outside the United States; and
- the establishment of additional strategic or licensing arrangements with other companies, or acquisitions.

If our existing resources are insufficient to satisfy our liquidity requirements due to slower than anticipated sales of Angiomax or otherwise, or if we acquire additional product candidates or businesses, or if we otherwise believe that raising additional capital would be in our interest and the interests of our stockholders, we may sell equity or debt securities or seek additional financing through other arrangements. Any sale of additional equity or debt securities may result in dilution to our stockholders, and debt financing may involve covenants limiting or restricting our ability to take specific actions, such as incurring additional debt or making capital expenditures. We cannot be certain that public or private financing will be available in amounts or on terms acceptable to us, if at all. If we seek to raise funds through collaboration or licensing arrangements with third parties, we may be required to relinquish rights to products, product candidates or technologies that we would not otherwise relinquish or grant licenses on terms that may not be favorable to us. If we are unable to obtain additional financing, we may be required to delay, reduce the scope of, or eliminate one or more of our planned research, development and commercialization activities, which could harm our financial condition and operating results.

#### **Risks Related to Commercialization**

##### ***Angiomax competes with all categories of anticoagulant drugs, which may limit the use of Angiomax***

Because different anticoagulant drugs act on different components of the clotting process, we believe that continued clinical work will be necessary to determine the best combination of drugs for clinical use. We recognize that Angiomax competes with other anticoagulant drugs to the extent Angiomax and any of these anticoagulant drugs are approved for the same or similar indications.

In addition, other anticoagulant drugs may compete with Angiomax for hospital financial resources. For example, many U.S. hospitals receive a fixed reimbursement amount per procedure for the angioplasties and other treatment therapies they perform. Because this amount is not based on the actual expenses the hospital incurs, hospitals may choose to use either Angiomax or other anticoagulant drugs, but not necessarily several of the drugs together.





***Because the market for thrombin inhibitors is competitive, our product may not obtain widespread use***

We have positioned Angiomax as a replacement for heparin, which is a widely used, inexpensive, generic drug used in patients with arterial thrombosis. Because heparin is inexpensive and has been widely used for many years, physicians and medical decision-makers may be hesitant to adopt Angiomax. In addition, due to the high incidence and severity of cardiovascular diseases, competition in the market for thrombin inhibitors is intense and growing. We cannot assure you that the rate of Angiomax sales growth will not slow or decline in 2007 and future years. There are a number of direct and indirect thrombin inhibitors currently on the market, awaiting regulatory approval and in development, including orally administered agents. The thrombin inhibitors on the market include products for use in the treatment of patients with HIT/HITTS, patients with unstable angina and patients with deep vein thrombosis.

***We face substantial competition, which may result in others discovering, developing or commercializing competing products before or more successfully than we do***

Our industry is highly competitive. Our success will depend on our ability to acquire and develop products and apply technology, and our ability to establish and maintain markets for our products. Potential competitors in the United States and other countries include major pharmaceutical and chemical companies, specialized pharmaceutical companies and biotechnology firms, universities and other research institutions. Many of our competitors have substantially greater research and development capabilities and experience, and greater manufacturing, marketing and financial resources, than we do. Accordingly, our competitors may develop or license products or other novel technologies that are more effective, safer, more convenient or less costly than existing products or technologies or products or technologies that are being developed by us or may obtain regulatory approvals for products more rapidly than we are able. Technological development by others may render our products or product candidates noncompetitive. We may not be successful in establishing or maintaining technological competitiveness.

***Near-term growth in our sales of Angiomax is dependent on acceptance by physicians, patients and other key decision-makers of Angiomax clinical data***

In the fall of 2002, we completed a 6,002 patient post-marketing Phase IIIb/IV clinical trial of Angiomax in coronary angioplasty called REPLACE-2. In November 2002, the principal investigators of the REPLACE-2 trial announced that, based on 30-day patient follow-up results, Angiomax met all of the primary and secondary objectives of the trial. In March 2003, we released the results of the detailed cost analysis study to examine per-patient total hospital resource consumption at U.S. clinical trial sites. In September 2003, the principal investigators of the clinical trial announced that, based on six-month patient follow-up results, Angiomax again met all of the primary and secondary objectives of the trial. In November 2003, the principal investigators presented one-year follow-up mortality data from the trial, which confirmed the 30-day and six-month mortality results. In December 2005, we completed enrollment in a 13,819 patient Phase III clinical trial studying Angiomax use in patients presenting to the emergency department with acute coronary syndromes called the ACUITY trial. In March 2006, the principal investigators of the ACUITY trial announced that ACUITY had met its objectives in favor of Angiomax based on 30-day patient results. The investigators for the ACUITY trial have continued to conduct the trial as they collect 12-month patient follow-up results. We expect these results to be reported by the investigators in the first half of 2007. These results may not meet the trial objectives or be consistent with the 30-day results.

We believe that the near-term commercial success of Angiomax will depend upon the extent to which physicians, patients and other key decision-makers accept the results of the Angiomax clinical trials. For example, since the original results of REPLACE-2 were announced in 2002, additional hospitals have granted Angiomax formulary approval and hospital demand for the product has increased. We cannot be certain, however, that these trends will continue. Some commentators have challenged various aspects of

the trial design of REPLACE-2, the conduct of the study and the analysis and interpretation of the results from the study, including how we define bleeding and the clinical relevance of types of ischemic events. The FDA has noted that in its view, statistical non-inferiority was not demonstrated as compared to the heparin plus a GP IIb/IIIa inhibitor arm of the trial for the 30-day ischemic endpoint in the REPLACE-2 trial. Similarly, we cannot be certain of the extent to which physicians, patients and other key decision-makers will accept the results of the ACUITY trial. If physicians, patients and other key decision-makers do not accept the REPLACE-2 and ACUITY trial results, adoption of Angiomax may suffer, and our business will be materially adversely affected.

***Our ability to generate future revenue from products will be affected by reimbursement and drug pricing***

Acceptable levels of reimbursement of drug treatments by government authorities, private health insurers and other organizations will have an effect on our ability to successfully commercialize, and attract collaborative partners to invest in the development of, product candidates. We cannot be sure that reimbursement in the United States or elsewhere will be available for any products we may develop or, if already available, will not be decreased in the future. If reimbursement is not available or is available only to limited levels, we may not be able to commercialize our products, or may not be able to obtain a satisfactory financial return on our products.

In certain countries, particularly the countries of the European Union, the pricing of prescription pharmaceuticals and the level of reimbursement are subject to governmental control. In some countries, it can take an extended period of time to establish and obtain reimbursement, and reimbursement approval may be required at the individual patient level, which can lead to further delays.

Third party payers increasingly are challenging prices charged for medical products and services. Also, the trend toward managed health care in the United States and the changes in health insurance programs, as well as legislative proposals, may result in lower prices for pharmaceutical products, including any products that may be offered by us. Cost-cutting measures that health care providers are instituting, and the effect of any health care reform, could materially adversely affect our ability to sell any products that are successfully developed by us and approved by regulators. Moreover, we are unable to predict what additional legislation or regulation, if any, relating to the health care industry or third party coverage and reimbursement may be enacted in the future or what effect such legislation or regulation would have on our business.

***We must comply with federal, state and foreign laws and regulations relating to the health care business, and, if we do not fully comply with such laws and regulations, we could face substantial penalties.***

We and our customers are subject to extensive regulation by the federal government, the states and foreign countries in which we may conduct our business. The laws that directly or indirectly affect our ability to operate our business include the following:

- the federal Medicare and Medicaid Anti-Kickback law, which prohibits persons from knowingly and willfully soliciting, offering, receiving or providing remuneration, directly or indirectly, in cash or in kind, to induce either the referral of an individual or furnishing or arranging for a good or service, for which payment may be made under federal health care programs such as Medicare and Medicaid;
- other Medicare laws and regulations that prescribe the requirements for coverage and payment for services performed by our customers, including the amount of such payment;
- the Federal False Claims Act, which imposes civil and criminal liability on individuals and entities who submit, or cause to be submitted, false or fraudulent claims for payment to the government; and

- the Federal False Statements Act, which prohibits knowingly and willfully falsifying, concealing or covering up a material fact or making any materially false statement in connection with delivery of or payment for health care benefits, items or services;

If our operations are found to be in violation of any of the laws and regulations described above or any other law or governmental regulation to which we or our customers are or will be subject, we may be subject to civil and criminal penalties, damages, fines exclusion from the Medicare and Medicaid programs and the curtailment or restructuring of our operations. Similarly, if our customers are found to be non-compliant with applicable laws, they may be subject to sanctions, which could also have a negative impact on us. Any penalties, damages, fines, curtailment or restructuring of our operations would adversely affect our ability to operate our business and our financial results. Any action against us for violation of these laws, even if we successfully defend against it, could cause us to incur significant legal expenses, divert our management's attention from the operation of our business and damage our reputation.

***We could be exposed to significant liability if we are unable to obtain insurance at acceptable costs and adequate levels or otherwise protect ourselves against potential product liability claims***

Our business exposes us to potential product liability risks which are inherent in the testing, manufacturing, marketing and sale of human healthcare products. Product liability claims might be made by patients in clinical trials, consumers, health care providers or pharmaceutical companies or others that sell our products. These claims may be made even with respect to those products that are manufactured in licensed and regulated facilities or otherwise possess regulatory approval for commercial sale.

These claims could expose us to significant liabilities that could prevent or interfere with the development or commercialization of our products. Product liability claims could require us to spend significant time and money in litigation or pay significant damages. With respect to our commercial sales and our clinical trials, we are covered by products liability insurance in the amount of \$20.0 million per occurrence and \$20.0 million annually in the aggregate on a claims-made basis. This coverage may not be adequate to cover any product liability claims.

As we continue to commercialize our products, we may wish to increase our product liability insurance. Product liability coverage is expensive. In the future, we may not be able to maintain or obtain such product liability insurance on reasonable terms, at a reasonable cost or in sufficient amounts to protect us against losses due to product liability claims.

#### **Risks Related to Regulatory Matters**

***If we do not obtain regulatory approvals for our product candidates we will not be able to market our product candidates and our ability to generate additional revenue could be materially impaired***

Except for Angiomax, which has been approved for sale in the United States for use as an anticoagulant in combination with aspirin in patients with unstable angina undergoing PCI and patients undergoing PCI with or at risk of HIT/HITTS, and which has been approved for sale in the European Union and in other countries for indications similar to those approved by the FDA, we do not have any other product approved for sale in the United States or any foreign market. We must obtain approval from the FDA in order to sell our product candidates in the United States and from foreign regulatory authorities in order to sell our product candidates in other countries. Obtaining FDA approval is uncertain, time-consuming and expensive. Any regulatory approval we ultimately obtain may be limited or subject to restrictions or post-approval commitments that render the product commercially non-viable. Securing regulatory approval requires the submission of extensive pre-clinical and clinical data, information about product manufacturing processes and inspection of facilities and supporting information to the regulatory authorities for each therapeutic indication to establish the product's safety and efficacy. If we are unable to submit the necessary data and information, for example, because the results of clinical

trials are not favorable, or if the FDA delays reviewing or does not approve our applications, we will be unable to obtain regulatory approval. Delays in obtaining or failure to obtain regulatory approvals may:

- delay or prevent the successful commercialization of any of our product candidates;
- diminish our competitive advantage; and
- defer or decrease our receipt of revenue.

The regulatory review and approval process to obtain marketing approval for a new drug or indications takes many years and requires the expenditure of substantial resources. This process can vary substantially based on the type, complexity, novelty and indication of the product candidate involved. The regulatory authorities have substantial discretion in the approval process and may refuse to accept any application or may decide that data is insufficient for approval and require additional pre-clinical, clinical or other studies. In addition, varying interpretations of the data obtained from pre-clinical and clinical testing could delay, limit or prevent regulatory approval of a product candidate.

***We cannot expand the indications for which we are marketing Angiomax unless we receive FDA approval for each additional indication. Failure to expand these indications will limit the size of the commercial market for Angiomax***

The FDA has approved Angiomax for use as an anticoagulant in combination with aspirin in patients with unstable angina undergoing PCI and patients undergoing PCI with or at risk of HIT/HITTS. One of our key objectives is to expand the indications for which Angiomax is approved for marketing by the FDA. In order to market Angiomax for expanded indications, we will need to conduct appropriate clinical trials, obtain positive results from those trials and obtain FDA approval for such proposed indications. Obtaining FDA approval is uncertain, time-consuming and expensive. The regulatory review and approval process to obtain marketing approval for a new indication can take many years and require the expenditure of substantial resources. This process can vary substantially based on the type, complexity, novelty and indication of the product candidate involved. The regulatory authorities have substantial discretion in the approval process and may refuse to accept any application or may decide that any data submitted is insufficient for approval and require additional pre-clinical, clinical or other studies. In addition, varying interpretations of the data obtained from pre-clinical and clinical testing could delay, limit or prevent regulatory approval of a new indication product candidate. For example, we recently received a non-approvable letter from the FDA in connection with our application to market Angiomax in patients with or at risk of HIT/HITTS undergoing cardiac surgery. While we plan to discuss the matter with the FDA, the FDA may require additional studies which may require the expenditure of substantial resources. Even if any such studies are undertaken, we can provide no assurance that we will be successful in obtaining regulatory approval for this indication in a timely manner or at all. If the one-year results of the ACUITY trial are favorable, we currently anticipate submitting an application with the FDA in 2007 for an expansion of the Angiomax product label to include information about ACUITY maintenance dosing regimen starting in the emergency department and the trial results. If the one-year results are not favorable, however, we are unlikely to be able to seek or obtain FDA approval to expand the Angiomax product label. If we are unsuccessful in expanding the Angiomax product label, the size of the commercial market for Angiomax will be limited.

***Clinical trials of product candidates are expensive and time-consuming, and the results of these trials are uncertain***

Before we can obtain regulatory approvals to market any product for a particular indication, we will be required to complete pre-clinical studies and extensive clinical trials in humans to demonstrate the safety and efficacy of such product for such indication.

Clinical testing is expensive, difficult to design and implement, can take many years to complete and is uncertain as to outcome. Success in pre-clinical testing or early clinical trials does not ensure that later clinical trials will be successful, and interim results of a clinical trial do not necessarily predict final results. An unexpected result in one or more of our clinical trials can occur at any stage of testing. We may experience numerous unforeseen events during, or as a result of, the clinical trial process that could delay or prevent us from receiving regulatory approval or commercializing our products, including:

- our clinical trials may produce negative or inconclusive results, and we may decide, or regulators may require us, to conduct additional clinical trials which even if undertaken cannot ensure we will gain approval;
- data obtained from pre-clinical testing and clinical trials may be subject to varying interpretations, which could result in the FDA or other regulatory authorities deciding not to approve a product in a timely fashion, or at all;
- the cost of clinical trials may be greater than we currently anticipate;
- regulators or institutional review boards may not authorize us to commence a clinical trial or conduct a clinical trial at a prospective trial site;
- we, or the FDA, might suspend or terminate a clinical trial at any time on various grounds, including a finding that participating patients are being exposed to unacceptable health risks. For example, in March 2005, we voluntarily suspended enrollment in one of our clinical trials for Cleviprex until December 2005 to review an interim analysis of safety data from the trial; and
- the effects of our product candidates may not be the desired effects or may include undesirable side effects or the product candidates may have other unexpected characteristics.

The rate of completion of clinical trials depends in part upon the rate of enrollment of patients. Patient enrollment is a function of many factors, including the size of the patient population, the proximity of patients to clinical sites, the eligibility criteria for the trial, the existence of competing clinical trials and the availability of alternative or new treatments. In particular, the patient population targeted by some of our clinical trials may be small. Delays in patient enrollment in any of our current or future clinical trials may result in increased costs and program delays.

***If we or our contract manufacturers fail to comply with the extensive regulatory requirements to which we, our contract manufacturers and our products are subject, our products could be subject to restrictions or withdrawal from the market and we could be subject to penalties***

The testing, manufacturing, labeling, safety, advertising, promotion, storage, sales, distribution, export and marketing, among other things, of our products, both before and after approval, are subject to extensive regulation by governmental authorities in the United States, Europe and elsewhere throughout the world. Both before and after approval of a product, quality control and manufacturing procedures must conform to current good manufacturing practice, or cGMP. Regulatory authorities, including the FDA, periodically inspect manufacturing facilities to assess compliance with cGMP. Our failure or the failure of our contract manufacturers to comply with the laws administered by the FDA, the European Medicines Agency, or other governmental authorities could result in, among other things, any of the following:

- delay in approving or refusal to approve a product;
- product recall or seizure;
- suspension or withdrawal of an approved product from the market;
- interruption of production;



- operating restrictions;
- warning letters;
- injunctions;
- fines and other monetary penalties;
- criminal prosecutions; and
- unanticipated expenditures.

**Risks Related to our Dependence on Third Parties for Manufacturing, Research and Development, and Distribution Activities**

*We depend on single suppliers for the production of Angiomax, Cleviprex and cangrelor bulk drug substance and different single suppliers to carry out all fill-finish activities*

We do not manufacture any of our products and do not plan to develop any capacity to manufacture them. We currently obtain all of our Angiomax bulk drug substance from one manufacturer, Lonza Braine, and rely on another manufacturer, Ben Venue Laboratories, to carry out all fill-finish activities for Angiomax, which includes final formulation and transfer of the drug into vials where it is then freeze-dried and sealed. The terms of our agreement with Lonza Braine require us to purchase from Lonza Braine a substantial portion of our Angiomax bulk drug product manufactured using the Chemilog process, a chemical synthesis process that we developed with UCB Bioproducts S.A., the predecessor to Lonza Braine.

We currently obtain all of our Cleviprex bulk drug substance from one manufacturer, Johnson Matthey Pharma Services. We rely on a different single supplier, Hospira, Inc., and its proprietary formulation technology, for the manufacture of all finished Cleviprex product, as well as for release testing and clinical packaging.

We have transferred the manufacturing process for all of our cangrelor bulk drug substance from AstraZeneca to Johnson Matthey Pharma Services for scale up and manufacture for Phase III clinical trials and commercial supplies. We will also rely on a different single supplier, Baxter Pharmaceutical Solutions LLC, for the manufacture of all finished cangrelor drug product for all Phase III clinical trials and to carry out release testing.

There are a limited number of manufacturers capable of manufacturing Angiomax, Cleviprex and cangrelor. We do not currently have alternative sources for production of bulk drug substance or to carry out fill-finish activities. Consolidation within the pharmaceutical manufacturing industry could further reduce the number of manufacturers capable of producing our products, or otherwise affect our existing contractual relationships.

In the event that Lonza Braine, Johnson Matthey, Hospira, Ben Venue or Baxter is unable or unwilling to carry out their respective manufacturing obligations or terminate or refuse to renew their respective arrangements with us, we may be unable to obtain alternative manufacturing, or obtain such manufacturing on commercially reasonable terms or on a timely basis. If we were required to transfer manufacturing processes to other third party manufacturers, we would be required to satisfy various regulatory requirements, which could cause us to experience significant delays in receiving an adequate supply of Angiomax, Cleviprex or cangrelor. Moreover, we may not be able to transfer processes that are proprietary to the manufacturer. Any delays in the manufacturing process may adversely impact our ability to meet commercial demands for Angiomax on a timely basis and supply product for clinical trials of Angiomax, Cleviprex or cangrelor.



***The development and commercialization of our products may be terminated or delayed, and the costs of development and commercialization may increase, if third parties on whom we rely to manufacture and support the development and commercialization of our products do not fulfill their obligations***

Our development and commercialization strategy involves entering into arrangements with corporate and academic collaborators, contract research organizations, distributors, third party manufacturers, licensors, licensees and others to conduct development work, manage or conduct our clinical trials, manufacture our products and market and sell our products outside of the United States. We do not have the expertise or the resources to conduct such activities on our own and, as a result, are particularly dependent on third parties in most areas.

We may not be able to maintain our existing arrangements with respect to the commercialization or manufacture of Angiomax or establish and maintain arrangements to develop and commercialize Cleviprex, cangrelor or any additional product candidates or products we may acquire on terms that are acceptable to us. Any current or future arrangements for development and commercialization may not be successful. If we are not able to establish or maintain agreements relating to Angiomax, Cleviprex, cangrelor or any additional products we may acquire on terms that we deem favorable, our results of operations would be materially adversely affected.

Third parties may not perform their obligations as expected. The amount and timing of resources that third parties devote to developing, manufacturing and commercializing our products are not within our control. Our collaborators may develop, manufacture or commercialize, either alone or with others, products and services that are similar to or competitive with the products that are the subject of the collaboration with us. Furthermore, our interests may differ from those of third parties that manufacture or commercialize our products. Our collaborators may re-evaluate their priorities from time to time, including following mergers and consolidations, and change the focus of their development, manufacturing or commercialization efforts. Disagreements that may arise with these third parties could delay or lead to the termination of the development or commercialization of our product candidates, or result in litigation or arbitration, which would be time consuming and expensive.

If any third party that manufactures or supports the development or commercialization of our products breaches or terminates its agreement with us, or fails to commit sufficient resources to our collaboration or conduct its activities in a timely manner, or fails to comply with regulatory requirements, such breach, termination or failure could:

- delay or otherwise adversely impact the manufacturing, development or commercialization of Angiomax, Cleviprex, cangrelor or any additional products that we may acquire or develop;
- require us to seek a new collaborator or undertake unforeseen additional responsibilities or devote unforeseen additional resources to the manufacturing, development or commercialization of our products; or
- result in the termination of the development or commercialization of our products.

*Use of third party manufacturers may increase the risk that we will not have appropriate supplies of our product candidates*

Reliance on third party manufacturers entails risks to which we would not be subject if we manufactured product candidates or products ourselves, including:

- reliance on the third party for regulatory compliance and quality assurance;
- the possible breach of the manufacturing agreement by the third party; and
- the possible termination or nonrenewal of the agreement by the third party, based on its own business priorities, at a time that is costly or inconvenient for us.

Angiomax and our product candidates may compete with products and product candidates of third parties for access to manufacturing facilities. If we are not able to obtain adequate supplies of Angiomax, Cleviprex and cangrelor, it will be more difficult for us to compete effectively and develop our product candidates.

Our contract manufacturers are subject to ongoing, periodic, unannounced inspection by the FDA and corresponding state and foreign agencies or their designees to evaluate compliance with the FDA's cGMP, regulations and other governmental regulations and corresponding foreign standards. We cannot be certain that our present or future manufacturers will be able to comply with cGMP regulations and other FDA regulatory requirements or similar regulatory requirements outside the United States. We do not control compliance by our contract manufacturers with these regulations and standards. Failure of our third party manufacturers or us to comply with applicable regulations could result in sanctions being imposed on us, including, fines and other monetary penalties, injunctions, civil penalties, failure of regulatory authorities to grant marketing approval of our product candidates, delays, suspension or withdrawal of approvals, license revocation, seizures or recalls of product candidates or products, interruption of production, warning letters, operating restrictions and criminal prosecutions, any of which could significantly and adversely affect supplies of Angiomax and our product candidates.

#### **Risks Related to our Intellectual Property**

*A breach of any of the agreements under which we license commercialization rights to products or technology from others could cause us to lose license rights that are important to our business or subject us to claims by our licensors*

We license rights to products and technology that are important to our business, and we expect to enter into additional licenses in the future. For instance, we have exclusively licensed patents and patent applications relating to Angiomax from Biogen Idec and Health Research Inc. and relating to Cleviprex and cangrelor from AstraZeneca. Under these agreements, we are subject to commercialization and

development, sublicensing, royalty, patent prosecution and maintenance, insurance and other obligations. For instance, we are required under our license of Cleviprex to submit an NDA for Cleviprex by September 30, 2007 and under our license of cangrelor to submit an NDA for cangrelor by December 31, 2008. Any failure by us to comply with any of these obligations or any other breach by us of our license agreements could give the licensor the right to terminate the license in whole, terminate the exclusive nature of the license or bring a claim against us for damages. Any such termination or claim, particularly relating to our agreements with respect to Angiomax, could have a material adverse effect on our business. We have entered into an agreement with Biogen Idec that suspends the statute of limitations relating to any claims for damages and/or license termination that they may bring in the event that a dispute arises between us and Biogen Idec relating to the late filing of our application under the Hatch Waxman Act for an extension of the term of the principal patent that covers Angiomax. Even if we contest any such termination or claim and are ultimately successful, our stock price could suffer. In addition, upon any termination of a license agreement, we may be required to license to the licensor any related intellectual property that we developed.

***If we are unable to obtain or maintain patent protection for the intellectual property relating to our products, the value of our products will be adversely affected***

The patent positions of pharmaceutical companies like us are generally uncertain and involve complex legal, scientific and factual issues. Our success depends significantly on our ability to:

- obtain and maintain U.S. and foreign patents, including defending those patents against adverse claims;
- protect trade secrets;
- operate without infringing the proprietary rights of others; and
- prevent others from infringing our proprietary rights.

We may not have any additional patents issued from any patent applications that we own or license. If additional patents are granted, the claims allowed may not be sufficiently broad to protect our technology. In addition, issued patents that we own or license may be challenged, narrowed, invalidated or circumvented, which could limit our ability to stop competitors from marketing similar products or limit the length of term of patent protection we may have for our products. Changes in patent laws or in interpretations of patent laws in the United States and other countries may diminish the value of our intellectual property or narrow the scope of our patent protection.

Our patents also may not afford us protection against competitors with similar technology. Because patent applications in the United States and many foreign jurisdictions are typically not published until eighteen months after filing, or in some cases not at all, and because publications of discoveries in the scientific literature often lag behind actual discoveries, neither we nor our licensors can be certain that others have not filed or maintained patent applications for technology used by us or covered by our pending patent applications without our being aware of these applications.

We exclusively license U.S. patents, patent applications and patent rights and corresponding foreign patents, patent applications and patent rights relating to Angiomax, Cleviprex and cangrelor. We exclusively license six issued U.S. patents relating to Angiomax, the rights relating to Cleviprex under three issued U.S. patents and the rights relating to cangrelor under five issued U.S. patents. We have not yet filed any independent patent applications.

The principal U.S. patent that covers Angiomax expires in 2010. The U.S. Patent and Trademark Office, or PTO, has rejected our application under the Hatch Waxman Act for an extension of the term of the patent beyond 2010 because the application was not filed on time by our counsel. In October 2002, we

filed a request with the PTO for reconsideration of the denial of the application. In addition, we have been exploring other alternatives to extend the term of the patent.

On December 6, 2006, the United States House of Representatives passed a bill that, if enacted, would have provided the PTO with discretion to consider patent extension applications filed late unintentionally under the Hatch Waxman Act. On December 9, 2006, the United States Senate adjourned without considering this bill. While we are hopeful that, in the current session, Congress will consider legislation similar to that passed by the House in December 2006, we can provide no assurance that a bill will be introduced or enacted or that, if it is enacted, the PTO will consider our application.

In January 2007, having been advised that the PTO might take some near term administrative activity on our request for reconsideration, we filed a petition requesting the PTO to stay any action on the application. The PTO granted a stay of 30 days from February 12, 2007. We can provide no assurance that, at the end of that 30 day period or thereafter, the PTO will not take action that could adversely affect our efforts to extend the term of the patent. We continue to explore alternatives to extend the term of the patent but we can provide no assurance that we will be successful in doing so.

We have entered into agreements with the counsel involved in the late filing that suspend the statute of limitations on our claims against them for failing to make a timely filing. We have entered into a similar agreement with Biogen Idec relating to any claims for damages and/or license termination they may bring in the event that a dispute arises between us and Biogen Idec relating to the late filing. These agreements may be terminated by either party upon 30 days notice. We cannot assure you that Biogen Idec will not terminate this agreement.

***We may be unable to utilize the Chemilog process if Lonza Braine breaches our agreement***

Our agreement with Lonza Braine for the supply of Angiomax bulk drug substance requires that Lonza Braine transfer the technology that was used to develop the Chemilog process to a secondary supplier of Angiomax bulk drug substance or to us or an alternate supplier at the expiration of the agreement. If Lonza Braine fails or is unable to transfer successfully this technology, we would be unable to employ the Chemilog process to manufacture our Angiomax bulk drug substance, which could cause us to experience delays in the manufacturing process and increase our manufacturing costs in the future.

***If we are not able to keep our trade secrets confidential, our technology and information may be used by others to compete against us***

We rely significantly upon unpatented proprietary technology, information, processes and know-how. We seek to protect this information by confidentiality agreements with our employees, consultants and other third party contractors, as well as through other security measures. We may not have adequate remedies for any breach by a party to these confidentiality agreements. In addition, our competitors may learn or independently develop our trade secrets. If our confidential information or trade secrets become publicly known, they may lose their value to us.

***If we infringe or are alleged to infringe intellectual property rights of third parties, it will adversely affect our business***

Our research, development and commercialization activities, as well as any product candidates or products resulting from these activities, may infringe or be claimed to infringe patents or patent applications under which we do not hold licenses or other rights. Third parties may own or control these patents and patent applications in the United States and abroad. These third parties could bring claims against us or our collaborators that would cause us to incur substantial expenses and, if successful against us, could cause us to pay substantial damages. Further, if a patent infringement suit were brought against

us or our collaborators, we or they could be forced to stop or delay research, development, manufacturing or sales of the product or product candidate that is the subject of the suit.

As a result of patent infringement claims, or in order to avoid potential claims, we or our collaborators may choose or be required to seek a license from the third party and be required to pay license fees or royalties or both. These licenses may not be available on acceptable terms, or at all. Even if we or our collaborators were able to obtain a license, the rights may be nonexclusive, which could result in our competitors gaining access to the same intellectual property. Ultimately, we could be prevented from commercializing a product, or be forced to cease some aspect of our business operations, if, as a result of actual or threatened patent infringement claims, we or our collaborators are unable to enter into licenses on acceptable terms. This could harm our business significantly.

There has been substantial litigation and other proceedings regarding patent and other intellectual property rights in the pharmaceutical and biotechnology industries. In addition to infringement claims against us, we may become a party to other patent litigation and other proceedings, including interference proceedings declared by the PTO and opposition proceedings in the European Patent Office, regarding intellectual property rights with respect to our products and technology. The cost to us of any patent litigation or other proceeding, even if resolved in our favor, could be substantial. Some of our competitors may be able to sustain the costs of such litigation or proceedings more effectively than we can because of their substantially greater financial resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could have a material adverse effect on our ability to compete in the marketplace. Patent litigation and other proceedings may also absorb significant management time.

### **Risks Related to Growth and Employees**

#### ***If we fail to acquire and develop additional product candidates or approved products it will impair our ability to grow***

We have a single product, Angiomax, approved for marketing. In order to generate additional revenue, we intend to acquire and develop additional product candidates or approved products. The success of this growth strategy depends upon our ability to identify, select and acquire pharmaceutical products that meet the criteria we have established. Because we neither have, nor intend to establish, internal scientific research capabilities, we are dependent upon pharmaceutical and biotechnology companies and other researchers to sell or license product candidates to us. We will be required to integrate any acquired products into our existing operations. Managing the development of a new product entails numerous financial and operational risks, including difficulties in attracting qualified employees to develop the product.

Any product candidate we acquire will require additional research and development efforts prior to commercial sale, including extensive pre-clinical and/or clinical testing and approval by the FDA and corresponding foreign regulatory authorities. All product candidates are prone to the risks of failure inherent in pharmaceutical product development, including the possibility that the product candidate will not be safe and effective or approved by regulatory authorities.

In addition, we cannot assure you that any approved products that we develop or acquire will be:

- manufactured or produced economically;
- successfully commercialized; or
- widely accepted in the marketplace.

We have previously acquired rights to products and, after having conducted development activities, determined not to devote further resources to those products. We cannot assure you that any additional products that we acquire will be successfully developed.

In addition, proposing, negotiating and implementing an economically viable acquisition is a lengthy and complex process. Other companies, including those with substantially greater financial, marketing and sales resources, may compete with us for the acquisition of product candidates and approved products. We may not be able to acquire the rights to additional product candidates and approved products on terms that we find acceptable, or at all.

*We may undertake strategic acquisitions in the future and any difficulties from integrating such acquisitions could damage our ability to attain or maintain profitability*

We may acquire additional businesses and products that complement or augment our existing business. Integrating any newly acquired business or product could be expensive and time-consuming. We may not be able to integrate any acquired business or product successfully or operate any acquired business profitably. Moreover, we may need to raise additional funds through public or private debt or equity financing to acquire any businesses or products, which may result in dilution for stockholders or the incurrence of indebtedness.

*We may not be able to manage our business effectively if we are unable to attract and retain key personnel and consultants*

Our industry has experienced a high rate of turnover of management personnel in recent years. We are highly dependent on our ability to attract and retain qualified personnel for the acquisition, development and commercialization activities we conduct or sponsor. If we lose one or more of the members of our senior management, including our Chairman and Chief Executive Officer, Clive A. Meanwell, or our President and Chief Operating Officer, John P. Kelley, or other key employees or consultants, our ability to implement successfully our business strategy could be seriously harmed. Our ability to replace these key employees may be difficult and may take an extended period of time because of the limited number of individuals in our industry with the breadth of skills and experience required to acquire, develop and commercialize products successfully. Competition to hire from this limited pool is intense, and we may be unable to hire, train, retain or motivate such additional personnel.

#### **Risks Related to Our Common Stock**

*Fluctuations in our operating results could affect the price of our common stock*

Our operating results may vary from period to period based on factors including the amount and timing of sales of Angiomax, underlying hospital demand for Angiomax, our customers' buying patterns, the timing, expenses and results of clinical trials, announcements regarding clinical trial results and product introductions by us or our competitors, the availability and timing of third party reimbursement, including in Europe, sales and marketing expenses and the timing of regulatory approvals. If our operating results do not meet the expectations of securities analysts and investors as a result of these or other factors, the trading price of our common stock will likely decrease.

*Our stock price has been and may in the future be volatile. This volatility may make it difficult for you to sell common stock when you want or at attractive prices*

Our common stock has been and in the future may be subject to substantial price volatility. From January 1, 2005 to February 16, 2007, the last reported sale price of our common stock ranged from a high of \$36.18 per share to a low of \$15.50 per share. The value of your investment could decline due to the effect of any of the following factors upon the market price of our common stock:

- changes in securities analysts' estimates of our financial performance;
- changes in valuations of similar companies;

- variations in our operating results;
- acquisitions and strategic partnerships;
- announcements of technological innovations or new commercial products by us or our competitors;
- disclosure of results of clinical testing or regulatory proceedings by us or our competitors;
- the timing, amount and receipt of revenue from sales of our products and margins on sales of our products;
- governmental regulation and approvals;
- developments in patent rights or other proprietary rights;
- changes in our management; and
- general market conditions.

In addition, the stock market has experienced significant price and volume fluctuations, and the market prices of specialty pharmaceutical companies have been highly volatile. Moreover, broad market and industry fluctuations that are not within our control may adversely affect the trading price of our common stock. You must be willing to bear the risk of fluctuations in the price of our common stock and the risk that the value of your investment in our securities could decline.

*Our corporate governance structure, including provisions in our certificate of incorporation and by-laws and Delaware law, may prevent a change in control or management that security holders may consider desirable*

Section 203 of the General Corporation Law of the State of Delaware and our certificate of incorporation and by-laws contain provisions that might enable our management to resist a takeover of our company or discourage a third party from attempting to take over our company. These provisions include the inability of stockholders to act by written consent or to call special meetings, a classified board of directors and the ability of our board of directors to designate the terms of and issue new series of preferred stock without stockholder approval.

These provisions could have the effect of delaying, deferring, or preventing a change in control of us or a change in our management that stockholders may consider favorable or beneficial. These provisions could also discourage proxy contests and make it more difficult for stockholders to elect directors and take other corporate actions. These provisions could also limit the price that investors might be willing to pay in the future for shares of our common stock or our other securities.

**Item 1B.** Unresolved Staff Comments

None.

**Item 2.** Properties

We currently occupy approximately 52,128 square feet of office space in Parsippany, New Jersey under a lease expiring in January 2013. In addition, we lease approximately 5,700 square feet of office space in Waltham, Massachusetts under a lease expiring in December 2008. We also have offices in Milton Park, Abingdon, United Kingdom. We believe our current arrangements will be sufficient to meet our needs for the foreseeable future and that any required additional space will be available on commercially reasonable terms to meet space requirements if they arise.

**Item 3. Legal Proceedings**

From time to time, we are subject to legal proceedings and claims that arise in the ordinary course of our business. We believe that no matters currently pending would, in the event of an adverse outcome, have a material impact on our consolidated financial position, results of operations, or liquidity.

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

None.

**PART II**

**Item 5. Market for Registrant's Common Equity, Related Stockholder Matters and Issuer Purchases of Equity Securities**

**Market Information and Holders**

Our common stock trades on the NASDAQ Global Select Market under the symbol MDCO. The following table reflects the range of the high and low sale price per share of our common stock, as reported on the NASDAQ Global Select Market or its predecessor, the NASDAQ National Market, for the periods indicated. These prices reflect inter-dealer prices, without retail mark-up, mark-down or commission and may not necessarily represent actual transactions.

|                                     | Common Stock Price |          |
|-------------------------------------|--------------------|----------|
|                                     | High               | Low      |
| <b>Year Ended December 31, 2005</b> |                    |          |
| First Quarter                       | \$ 29.95           | \$ 20.70 |
| Second Quarter                      | 24.95              | 20.83    |
| Third Quarter                       | 24.55              | 20.13    |
| Fourth Quarter                      | 23.70              | 15.50    |
| <b>Year Ended December 31, 2006</b> |                    |          |
| First Quarter                       | 22.00              | 16.54    |
| Second Quarter                      | 21.34              | 16.81    |
| Third Quarter                       | 23.25              | 18.28    |
| Fourth Quarter                      | 36.18              | 22.05    |

American Stock Transfer & Trust Company is the transfer agent and registrar for our common stock. As of the close of business on February 23, 2007, we had 193 holders of record of our common stock.

**Dividends**

We have never declared or paid cash dividends on our common stock. We anticipate that we will retain all of our future earnings, if any, for use in the expansion and operation of our business and do not anticipate paying cash dividends in the foreseeable future. Payment of future dividends, if any, will be at the discretion of our board of directors.



**Item 6. Selected Financial Data**

In the table below, we provide you with our selected consolidated financial data. We have prepared this information using our audited consolidated financial statements for the years ended December 31, 2006, 2005, 2004, 2003 and 2002. In 2006 and 2004, we computed diluted earnings per share by giving effect to options and warrants outstanding at December 31, 2006 and 2004, respectively. We have not included options or warrants in the computation of diluted net loss per share for any other periods, as their effects would have been anti-dilutive. For further discussion of the computation of basic and diluted earnings/(loss) per share, please see note 9 to our consolidated financial statements.

You should read the following selected consolidated financial data in conjunction with our consolidated financial statements and related notes and Item 7 - Management's Discussion and Analysis of Financial Condition and Results of Operations.

|   | <b>Year Ended December 31,</b>               |             |             |             |             |
|---|--|-------------|-------------|-------------|-------------|
|   | <b>2006</b>                                  | <b>2005</b> | <b>2004</b> | <b>2003</b> | <b>2002</b> |
|   | <b>(in thousands, except per share data)</b> |             |             |             |             |
| <b>Statements of Operations Data</b>                              |  |             |             |             |             |
| Net revenue   | \$ 213,952                                   | \$ 150,207  | \$ 144,251  | \$ 85,591   | \$ 38,301   |
| Operating expenses  |  |             |             |             |             |
| Cost of revenue   | 51,812                                       | 34,762      | 29,123      | 22,749      | 10,284      |
| Research and development  | 63,536                                       | 64,389      | 49,290      | 35,905      | 37,951      |
| Selling, general and administrative                               | 88,265                                       | 63,053      | 50,275      | 45,082      | 36,808      |
| Total operating expenses  | 203,613                                      | 162,204     | 128,688     | 103,736     | 85,043      |
| Income/(loss) from operations                                     | 10,339                                       | (11,997 )   | 15,563      | (18,145 )   | (46,742 )   |
| Other income/(expense), net                                       | 7,319  | 4,344       | 2,126       | 1,403       | 911         |
| Benefit from/(provision for) income taxes                         | 46,068                                       | (100 )      | (690 )      | (128 )      |             |
| Net income/(loss)   | 63,726                                       | (7,753 )    | 16,999      | (16,870 )   | (45,831 )   |
| Basic earnings/(loss) per common share                            | \$ 1.27                                      | \$ (0.16 )  | \$ 0.36     | \$ (0.37 )  | \$ (1.23 )  |
| Shares used in computing basic earnings/(loss) per common share   | 50,300                                       | 49,443      | 47,855      | 45,624      | 37,210      |
| Diluted earnings/(loss) per common share                          | \$ 1.25                                      | \$ (0.16 )  | \$ 0.34     | \$ (0.37 )  | \$ (1.23 )  |
| Shares used in computing diluted earnings/(loss) per common share | 51,034                                       | 49,443      | 49,772      | 45,624      | 37,210      |

|  | <b>As of December 31,</b> |             |             |             |             |
|--|---------------------------|-------------|-------------|-------------|-------------|
|  | <b>2006</b>               | <b>2005</b> | <b>2004</b> | <b>2003</b> | <b>2002</b> |
|  | <b>(in thousands)</b>     |             |             |             |             |
| <b>Balance Sheet Data</b>  |                           |             |             |             |             |
| Cash and cash equivalents, available for sale securities and accrued interest receivable | \$ 198,231                | \$ 141,012  | \$ 161,224  | \$ 136,855  | \$ 43,638   |
| Working capital  | 228,523                   | 169,912     | 173,349     | 139,725     | 54,172      |
| Total assets   | 318,568                   | 208,707     | 210,044     | 166,662     | 74,714      |
| Accumulated deficit  | (241,172 )                | (304,898 )  | (297,145 )  | (314,144 )  | (297,274 )  |
| Total stockholders' equity   | 269,951                   | 170,899     | 171,671     | 140,165     | 53,934      |

Effective January 1, 2006, we adopted the fair value recognition provisions of SFAS 123(R), Share-Based Payment (SFAS 123(R)), using the accelerated expense attribution method specified in FASB Interpretation No. (FIN) 28, Accounting for Stock Appreciation Rights and Other Variable Stock Option or Award Plans (FIN 28). SFAS 123(R) requires us to recognize compensation expense in an amount equal to the fair value of all share-based awards granted to employees resulting in \$8.5 million in share-based compensation expense during 2006.

**Item 7. 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 financial statements and accompanying notes included elsewhere in this annual report. In addition to the historical information, the discussion in this annual report contains certain forward-looking statements that involve risks and uncertainties. Our actual results could differ materially from those anticipated by the forward-looking statements due to our critical accounting estimates discussed below and important factors set forth in this annual report, including under "Risk Factors" in Item 1A of this annual report.*

**Overview**

We are a pharmaceutical company providing innovative, cost effective acute care hospital products to the worldwide hospital marketplace. We have one marketed product, Angiomax® (bivalirudin), and two products in late-stage development, Cleviprex (clevidipine) and cangrelor. We market Angiomax to interventional cardiology customers for its approved uses in PCI, including in patients with HIT/HITTS. We market and sell Angiomax in the United States with a sales force of approximately 135 representatives and managers experienced in selling to hospital customers. In the European Union and other foreign jurisdictions, we sell Angiomax to third party distributors that market and distribute the product to hospitals. Our revenues to date have been generated principally from sales of Angiomax in the United States. We reported net revenue of \$214.0 million and net income of \$63.7 million for the year ended December 31, 2006, which includes a net income tax benefit of \$46.6 million recognized through the reduction of a portion of our valuation allowances associated with deferred tax assets.

In evaluating our operating performance, we focus on use of Angiomax by existing hospital customers, as well as penetration to new hospitals, which are critical elements of our ability to increase revenues. In 2005, we expanded our sales force and increased our marketing capabilities. We believe that our improved sales and marketing capabilities, and the expansion of our product label, has and will continue to allow us to more effectively serve our existing customers and penetrate new hospitals.

Research and development expenses represent costs incurred for product acquisition, clinical trials, activities relating to regulatory filings and manufacturing development efforts. We outsource much of our clinical trials and all of our manufacturing development activities to third parties to maximize efficiency and minimize our internal overhead. We expense our research and development costs as they are incurred. Selling, general and administrative expenses consist primarily of salaries and related expenses, general corporate activities and costs associated with marketing and promotional activities. Research and development expense and selling, general and administrative expense also include stock-based compensation expense, which we allocate based on the responsibilities of the recipients of the stock-based compensation.

We currently sell Angiomax in the United States to a limited number of domestic medical and pharmaceutical wholesalers with distribution centers located throughout the country. These wholesalers then sell and ship Angiomax to hospitals. In the United States, AmerisourceBergen Drug Corporation, McKesson Corporation and Cardinal Health accounted for more than 88% of our revenues for the year ended December 31, 2006. We are in the process of modifying our distribution system in the United States. As part of this modification, we plan to sell Angiomax to a third party who will ship Angiomax directly to our hospital customers. We anticipate that we will begin selling Angiomax under this revised distribution system in the second quarter and we expect that it will enable us to reduce our distribution costs in 2007 and provide us with improved data.

In 2005, we entered into fee-for-service arrangements with our largest wholesalers. We believe that these arrangements resulted in reductions in wholesaler inventories, improved margins, more predictable buying patterns and more frequent data on wholesaler inventory levels and hospital demand. In order to

reduce inventory level to a four to six week level, these wholesalers reduced their aggregate inventory by approximately \$39 million in the last two quarters of 2005 and in the first quarter of 2006.

Except for 2004 and 2006, we have incurred losses on an annual basis since inception. We expect to continue to spend significant amounts on the development of our products. We plan to continue to invest in clinical studies to expand the approved indications for Angiomax and to continue to develop Cleviprex and cangrelor. We also plan to continue our sales and marketing programs to promote Angiomax, and to support programs to educate and inform physicians, nurses, pharmacists and other medical decision makers about the benefits of Angiomax. In light of these activities, our intention to expand our sales force in preparation for the launch of Cleviprex, and our plan to continue to evaluate possible acquisitions of development-stage products, approved products, or businesses that fit within our growth strategy, we will likely need to generate greater revenues to maintain profitability.

We have accrued for U.S. and state income taxes, state taxes based on net worth and for a certain amount of income tax in international jurisdictions in our financial statements to the extent these taxes apply. At December 31, 2006, net operating losses available to offset future taxable income for federal income tax purposes were approximately \$225.0 million. If not utilized, federal net operating loss carryforwards will expire at various dates beginning in 2012 and ending in 2025. We have reduced a portion of our valuation allowance associated with the deferred tax assets because we now consider the realization of these assets to be more likely than not. The future utilization of net operating losses and credits may be subject to limitation based upon changes in ownership under the rules of the Internal Revenue Code (IRC). We experienced changes in ownership as defined by Section 382 of the IRC during the years ended December 31, 1998 and 2002. Based on the market value of our common stock at the time of those changes, we believe there will be no impact on our ability to utilize our net operating losses and credits. Of the \$225.0 million of our federal net operating losses, \$188.4 million is subject to limitations through 2010.

#### **Application of Critical Accounting Estimates**

The discussion and analysis of our financial condition and results of operations is based on our consolidated financial statements, which have been prepared in accordance with U.S. generally accepted accounting principles. The preparation of these financial statements requires us to make estimates and judgments that affect our reported assets and liabilities, revenues and expenses, and other financial information. Actual results may differ significantly from these estimates under different assumptions and conditions. In addition, our reported financial condition and results of operations could vary due to a change in the application of a particular accounting standard.

We regard an accounting estimate or assumption underlying our financial statements as a critical accounting estimate where:

- the nature of the estimate or assumption is material due to the level of subjectivity and judgment necessary to account for highly uncertain matters or the susceptibility of such matters to change; and
- the impact of the estimates and assumptions on financial condition or operating performance is material.

Our significant accounting policies are more fully described in note 2 to our consolidated financial statements included in this annual report on Form 10-K. Not all of these significant accounting policies, however, require that we make estimates and assumptions that we believe are critical accounting estimates. We have discussed our accounting policies with the audit committee of our board of directors, and we believe that our estimates relating to revenue recognition, inventory, stock-based compensation and income taxes described below are critical accounting estimates.

### ***Revenue Recognition***

*Product Sales.* We do not recognize revenue from product sales until there is persuasive evidence of an arrangement, delivery has occurred, the price is fixed and determinable, the buyer is obligated to pay us, the obligation to pay is not contingent on resale of the product, the buyer has economic substance apart from us, we have no obligation to bring about sale of the product, the amount of returns can be reasonably estimated and collectibility is reasonably assured.

#### *Domestic Sales*

We record allowances for chargebacks and other discounts, and accruals for product returns, rebates, and fee-for-service charges, at the time of sale, and report revenue net of such amounts. In determining the amounts of certain of these allowances and accruals, we must make significant judgments and estimates. For example, in determining these amounts, we estimate hospital demand, buying patterns by hospitals and group purchasing organizations from wholesalers and the levels of inventory held by wholesalers. Making these determinations involves estimating whether trends in past buying patterns by hospitals and group purchasing organizations will predict future product sales. Under the terms of our fee-for-service arrangements with our largest wholesalers, these wholesalers provided us with frequent data on wholesaler inventory levels and hospital purchases. We apply this data in determining the amounts of certain of these allowances and accruals.

The nature of our allowances and accruals requiring critical accounting estimates, and the specific considerations we use in estimating their amounts, are as follows:

- *Product returns.* Our customers have the right to return any unopened product during the 18-month period beginning six months prior to the labeled expiration date and ending 12 months after the labeled expiration date. As a result, in calculating the accrual for product returns, we must estimate the likelihood that product sold to customers might remain in their inventory to within six months of expiration and analyze the likelihood that such product will be returned within 12 months after expiration.

In estimating the likelihood of product remaining in our wholesalers' inventory, we have relied on information from our wholesalers regarding their inventory levels, measured hospital demand as reported by third party sources and on internal sales data. We also considered our wholesalers' past buying patterns, estimated remaining shelf life of product previously shipped and the expiration dates of product currently being shipped.

In estimating the likelihood of product return, we rely primarily on historic patterns of returns and estimated remaining shelf life of product previously shipped. During 2006, \$0.2 million of Angiomax was returned to us, representing less than 0.1% of the total revenue from Angiomax sales. During 2005, \$0.1 million of Angiomax was returned to us, representing approximately 0.1% of the total revenue from Angiomax sales.

At December 31, 2006 and 2005, our accrual for product returns was \$0.4 million and \$0.2 million, respectively. A 10% change in our accrual for product returns in 2006 would not have had a material effect on our reported revenue in 2006.

As we implement our revised distribution system, we plan to monitor inventory levels and analyze its buying and product return patterns.

- *Chargebacks and rebates.* Although we have sold Angiomax primarily to wholesalers, we typically enter into agreements with hospitals, either directly or through group purchasing organizations acting on behalf of their hospital members, in connection with the hospitals' purchases of Angiomax from the wholesalers. Based on the terms of these agreements, most of our hospital customers have the right to receive a discounted price and volume-based rebate on product purchases. We provide



a credit to the wholesaler, or a chargeback, representing the difference between the wholesaler's acquisition list price and the discounted price.

As a result of these contracts, at the time of product shipment, we must estimate the likelihood that Angiomax sold to wholesalers might be ultimately sold to a contracting hospital or group purchasing organization. We must also estimate the contracting hospital's or group purchasing organization's volume of purchases.

We base our estimates on the historic chargeback data we receive from wholesalers, which detail historic buying patterns and sales mix for particular hospitals and group purchasing organizations, and the applicable customer chargeback rates and rebate thresholds.

Under our revised distribution model, we do not anticipate that our expenses related to chargebacks and rebates will be impacted.

At December 31, 2006 and 2005, our allowance for chargebacks was \$0.3 million and \$0.5 million, respectively. A 10% change in our accrual for chargebacks in 2006 would not have had a material effect on our reported revenue in 2006. Our accrual for rebates was \$0.8 million at December 31, 2006 and \$1.5 million at December 31, 2005. A 10% change in our allowance for rebates would have had an approximate \$0.1 million effect on our reported revenue in 2006.

We have adjusted our allowances for chargebacks and our accruals for product returns and rebates in the past based on our actual sales experience, and we will likely be required to make adjustments to these allowances and accruals in the future. We continually monitor our allowances and accruals and make adjustments when we believe actual experience may differ from our estimates. The allowances included in the table below reflect these adjustments.

The following table provides a summary of activity with respect to our allowances and accruals over the course of 2006, 2005 and 2004 (amounts in thousands):

|   | Returns | Chargebacks | Rebates  |
|---|---------|-------------|----------|
| <b>Balance at December 31, 2003</b>         | 1,102   | 1,772       | 1,828    |
| Allowances for sales during 2004            | 121     | 5,978       | 2,663    |
| Actual credits issued for prior years sales | (617 )  | (1,852 )    | (1,913 ) |
| Actual credits issued for sales during 2004 | (3 )    | (2,795 )    | (954 )   |
| <b>Balance at December 31, 2004</b>         | 603     | 3,103       | 1,624    |
| Allowances for sales during 2005            | (240 )  | 1,776       | 2,334    |
| Actual credits issued for prior years sales | (146 )  | (2,895 )    | (1,317 ) |
| Actual credits issued for sales during 2005 |         | (1,478 )    | (1,187 ) |
| <b>Balance at December 31, 2005</b>         | \$ 217  | \$ 506      | \$ 1,454 |
| Allowances for sales during 2006            | 404     | 4,240       | 2,247    |
| Actual credits issued for prior years sales | (212 )  | (737 )      | (1,318 ) |
| Actual credits issued for sales during 2006 | (8 )    | (3,681 )    | (1,549 ) |
| <b>Balance at December 31, 2006</b>         | \$ 401  | \$ 328      | \$ 834   |

Because our three largest U.S. wholesalers accounted for approximately 88% of our net revenue in 2006, it is critical that these wholesalers have adequate inventory to meet product demand. Since we began selling Angiomax in the United States in 2001, wholesaler buying patterns have sometimes been unpredictable. In addition, product demand has generally not grown at a uniform rate. For example, we experienced an accelerated rate of demand for Angiomax following commercial launch and following the announcement of REPLACE-2 trial results in November 2002. As a result, we offered incentives to wholesalers from time to time in the past to ensure that they had what we believed to be appropriate stocks of product to meet expected increased demand as a result of events such as clinical trials, regulatory

approvals and competitive developments, or to ensure that they were stocking within a normal range of inventory for a product like Angiomax. We have not offered any such incentives in 2006 since we entered into the fee for service arrangements with wholesalers in 2005.

*International Distributors.* Under our arrangements with Nycomed, we sell our product at a fixed transfer price. The established transfer price is typically determined twice in each year, prior to the first and last shipment of Angiomax to the distributor each year. If not agreed upon by the parties prior to such shipments, the price is determined by taking an average of the transfer price for the preceding three shipments, subject to certain minimum pricing. The transfer price is denominated in Euros, which are then converted to U.S. Dollars, payable by our distributors, at the exchange rate between the two currencies, as quoted by the European Central Bank, just prior to shipment.

Under our agreements with our other international distributors, we sell our product at a percentage of the distributor's established net price. The established net price is typically determined in the quarter in which we sell our products to these distributors based on the distributor's net selling price to its customers. In those situations, usually prior to product launch, where product is sold prior to the establishment of the distributor's selling price, we initially record revenue at minimum prices outlined in these agreements and later adjust our selling price to the distributor once the distributor's net selling price is determined. In accordance with the terms of these agreements, under no circumstances would the subsequent adjustment result in the net selling price being lower than the minimum price.

Revenue from the sale of distribution rights includes amortization of milestone payments. We record these milestone payments as deferred revenue until contractual performance obligations have been satisfied, and we then recognize these payments ratably over the term of these agreements. When the period of deferral cannot be specifically identified from the contract, we must estimate the period based upon other critical factors contained within the contract. We review these estimates at least annually, which could result in a change in the deferral period.

*Reimbursement Revenue.* During 2006, in collaboration with a third party, we paid fees for services rendered by a research organization and other out-of-pocket costs for which we were reimbursed at cost, without mark-up or profit. The reimbursements received were reported as part of net revenue in our consolidated statements of operations and the fees for the services rendered and the out-of-pocket costs were included in research and development expenses. We have not incurred any fees under this arrangement in 2007 and do not expect to incur any additional fees under this arrangement.

### ***Inventory***

We record inventory upon the transfer of title from our vendors. Inventory is stated at the lower of cost or market value and valued using first-in, first-out methodology. Angiomax bulk product is classified as raw materials, and its costs are determined using acquisition costs from our contract manufacturers. We record work-in-progress costs of filling, finishing and packaging against specific product batches. Prior to FDA approval of any product, we expense all inventory costs as research and development. Following FDA approval we plan to record as inventory any bulk product manufactured.

We review our inventory for slow moving or obsolete amounts based on expected revenues. As of December 31, 2006 we had not recorded an allowance for slow moving or obsolete amounts of inventory. If actual revenues are less than expected, we may be required to make allowances for excess amounts of inventory in the future.

### ***Stock-Based Compensation***

We have established equity compensation plans for our employees, directors and certain other individuals. All grants and terms are authorized by our Board of Directors. We may grant non-qualified stock options and restricted stock awards under our 2004 Stock Incentive Plan (the "2004 Plan"). Options and restricted stock awards generally become exercisable or vest over four years from the grant date, and options must be exercised within ten years of the grant date.

Prior to the January 1, 2006 adoption of FASB Statement No. 123(R), Share Based Payment ( SFAS 123(R) ), we accounted for stock option plans and restricted stock award plans in accordance with Accounting Principles Board Opinion ( APB ) No. 25, Accounting for Stock Issued to Employees. Accordingly, no compensation expense had been recognized for stock options since all options granted had an exercise price equal to the market value of the underlying stock on the grant date. As permitted by SFAS No. 123, Accounting for Stock-Based Compensation , stock-based compensation was presented as a pro forma disclosure in the notes to the consolidated financial statements.

Effective January 1, 2006, we adopted the fair value recognition provisions of SFAS 123(R). We have elected to use the modified prospective transition method and, therefore, adjustments to prior periods are not required as a result of adopting SFAS 123(R). Under this method, the provisions of SFAS 123(R) apply to all awards granted after January 1, 2006, and to any unrecognized expense of awards unvested at the date of adoption based on the grant date fair value. We will recognize expense over the vesting periods using the accelerated expense attribution method expense over the vesting periods using the accelerated expense attribution method specified in FASB Interpretation No. 28, Accounting for Stock Appreciation Rights and Other Variable Stock Option or Award Plans (FIN 28). We record expense associated with restricted stock awards as compensation cost over the requisite vesting periods based on the market value on the date of grant.

We estimate the fair value of each option on the date of grant using the Black-Scholes closed-form option-pricing model based on assumptions for the expected term of the stock options, expected volatility of our common stock, and prevailing interest rates. SFAS 123(R) also requires us to estimate forfeitures in calculating the expense relating to stock-based compensation as opposed to only recognizing forfeitures and the corresponding reduction in expense as they occur.

We have based our assumptions on the following:

| <b>Assumption</b>  | <b>Method of estimating</b>   |
|--|---|
| <ul style="list-style-type: none"> <li>• Estimated expected term of options</li> </ul> | <ul style="list-style-type: none"> <li>• Employees' historical exercise experience and we have also made estimates of future exercises of unexercised options based on the midpoint between the vesting date and end of the contractual term</li> </ul> |
| <ul style="list-style-type: none"> <li>• Expected volatility</li> </ul>                | <ul style="list-style-type: none"> <li>• Historic price of our common stock and the historic prices of the stock of our peer group</li> </ul>   |
| <ul style="list-style-type: none"> <li>• Risk-free interest rate</li> </ul>            | <ul style="list-style-type: none"> <li>• Yields of U.S. Treasury securities corresponding with the expected life of option grants</li> </ul>  |
| <ul style="list-style-type: none"> <li>• Forfeiture rates</li> </ul>                   | <ul style="list-style-type: none"> <li>• Historical forfeiture data</li> </ul>  |

Of these assumptions, the expected term of the option and expected volatility of our common stock are the most difficult to estimate since they are based on the exercise behavior of the employees and expected performance of our common stock. Increases in the term and the volatility of our common stock will generally cause an increase in compensation expense.

### ***Income Taxes***

Our annual effective tax rate is based on pre-tax earnings, existing statutory tax rates, limitations on the use of tax credits and net operating loss carryforwards and tax planning opportunities available in the jurisdictions in which we operate. Significant judgment is required in evaluating our tax position.

On a periodic basis, we evaluate the realizability of our deferred tax assets and liabilities and will adjust such amounts in light of changing facts and circumstances, including but not limited to future projections of taxable income, tax legislation, rulings by relevant tax authorities, tax planning strategies and the progress of ongoing tax audits. Settlement of filing positions that may be challenged by tax authorities could impact the income tax position in the year of resolution. Our current tax liability is presented in the consolidated balance sheet within accrued expenses.



At December 31, 2006, we had \$103.9 million of gross deferred tax assets, which included the effects of federal net operating loss carryforwards of \$225.0 million, research and development credits of \$15.7 million and other items of \$11.6 million. These assets are offset by a \$54.7 million valuation allowance since the realization of these future benefits is not considered more likely than not as our ability to estimate long-term future taxable income with a high level of certainty is limited. In assessing the realizability of deferred tax assets, we consider whether it is more likely than not that some portion or all of the deferred tax assets will not be realized. The ultimate realization of deferred tax assets is dependent upon the generation of future taxable income during the period in which those temporary differences become deductible or the NOLs and credit carryforwards can be utilized. When considering the reversal of the valuation allowance, we consider the level of past and anticipated future taxable income and the utilization of the carryforwards. Based upon these considerations, we reduced our valuation allowance by \$49.2 million in the fourth quarter of 2006 because we believe it is more likely than not that we will realize the benefits of a portion of our deferred tax assets. This valuation adjustment resulted in a benefit from income taxes in 2006. We expect that future periods will include income taxes at a higher effective rate. Revisions to the estimated net realizable value of the deferred tax asset could cause our provision for income taxes to vary significantly from period to period. Factors that could significantly impact our valuation allowance include the regulatory approval of products currently under development, extension of the patent rights relating to Angiomax or failure to achieve future anticipated revenues. Should we further reduce or increase the valuation allowance on deferred tax assets, a current year tax benefit or expense would be recognized and future periods would then include income taxes at a higher or lower rate than the effective rate in the period that the adjustment is made.

## Results of Operations

### Years Ended December 31, 2006 and 2005

*Net Revenue.* Net revenue increased 42% to \$214.0 million for 2006 as compared to \$150.2 million for 2005. In 2006, we derived approximately \$200.7 million of net revenue from U.S. sales of Angiomax and approximately \$11.3 million of net revenue from international sales of Angiomax. In 2005, we derived approximately \$140.7 million of net revenue from U.S. sales of Angiomax and approximately \$9.5 million of net revenue from international sales of Angiomax. We believe that the increase in U.S. sales in 2006 was due primarily to increased purchases of Angiomax by existing hospital customers, adoption of Angiomax by new hospital customers and the effects of higher prices as a result of a 7% price increase to our wholesalers in February 2006. The increase in U.S. sales also partly reflects the impact of reduced purchases by wholesalers in connection with our fee-for-service arrangements that we entered into with wholesalers in 2005 and 2006. We estimate that in implementing a planned inventory reduction, our wholesalers reduced their aggregate inventories by approximately \$13.0 million in the first quarter of 2006 and approximately \$26.0 million in the last two quarters of 2005. Our international revenue during 2006, while higher than our international revenue in 2005, reflected an increase in sales to our Canadian distributors. Nycomed sales remained at the same level.

### Net Revenue

| Net Revenue              | Year Ended December 31, |                       |                        |                       |
|--------------------------|-------------------------|-----------------------|------------------------|-----------------------|
|                          | 2006<br>(in thousands)  | % of Total<br>Revenue | 2005<br>(in thousands) | % of Total<br>Revenue |
| <b>Angiomax</b>          |                         |                       |                        |                       |
| United States Sales      | \$ 200,727              | 94 %                  | \$ 140,721             | 94 %                  |
| International Sales      | 11,277                  | 5 %                   | 9,486                  | 6 %                   |
| <b>Reimbursement</b>     | 1,948                   | 1 %                   |                        |                       |
| <b>Total Net Revenue</b> | \$ 213,952              | 100 %                 | \$ 150,207             | 100 %                 |

In 2006 and 2005, we recognized \$0.3 million and \$0.4 million, respectively, of international revenue from the amortization of milestone payments related to the \$2.5 million and \$1.5 million in non-refundable fees received from Nycomed. These milestone payments were recorded as deferred revenue in 2004 and 2002, respectively, and are being recognized ratably over the estimated term of our agreement with Nycomed.

*Cost of Revenue.* As shown in the table below, cost of revenue in 2006 was \$51.8 million, or 24% of net revenue, compared to \$34.8 million, or 23% of net revenue, in 2005. Cost of revenue consisted of expenses in connection with the manufacture of Angiomax sold, royalty expenses under our agreement with Biogen Idec and the logistics costs of selling Angiomax, such as distribution, storage, and handling.

#### Cost of Revenue

| Cost of Revenue              | Year Ended December 31, |                 |                        |                 |
|------------------------------|-------------------------|-----------------|------------------------|-----------------|
|                              | 2006<br>(in thousands)  | % of Total Cost | 2005<br>(in thousands) | % of Total Cost |
| Manufacturing                | \$ 18,508               | 36 %            | \$ 14,223              | 41 %            |
| Royalty                      | 27,216                  | 52 %            | 16,142                 | 46 %            |
| Logistics                    | 6,088                   | 12 %            | 4,397                  | 13 %            |
| <b>Total Cost of Revenue</b> | <b>\$ 51,812</b>        | <b>100 %</b>    | <b>\$ 34,762</b>       | <b>100 %</b>    |

The increase in cost of revenue for 2006 compared to 2005 resulted from an increase in manufacturing costs, logistics costs and royalty expenses due to higher sales volume and a higher effective royalty rate under our agreement with Biogen Idec. We expect our cost of revenue to increase in 2007 as a result of increased royalty rates on higher anticipated sales.

*Research and Development Expenses.* Research and development expenses decreased 1.3% to \$63.5 million for 2006, from \$64.4 million for 2005. The decrease in research and development expenses resulted primarily from a decrease in spending relating to AMI resulting from the completion of patient enrollment in 2005. This decrease was partially offset by increased investment in our Cleviprex and cangrelor development programs, increased investment in other research and development expenses, including \$1.9 million of expenses that we incurred in collaboration with a third party vendor under a contract research agreement, increased investment in statistics and data management for the analysis of the ACUITY trial data, and stock-based compensation expense of \$1.5 million.

The following table identifies, for each of our major research and development projects, our spending for 2006 and 2005. Spending for past periods is not necessarily indicative of spending in future periods.

## Research and Development Spending

| Research and Development           | Year Ended December 31, |                   |                        |                   |
|------------------------------------|-------------------------|-------------------|------------------------|-------------------|
|                                    | 2006<br>(in thousands)  | % of<br>Total R&D | 2005<br>(in thousands) | % of<br>Total R&D |
| <b>Angiomax</b>                    |                         |                   |                        |                   |
| Clinical trials                    | \$ 14,954               | 24 %              | \$ 37,377              | 58 %              |
| Manufacturing development          | 1,331                   | 2 %               | 936                    | 1 %               |
| Administrative and headcount costs | 2,695                   | 4 %               | 5,928                  | 9 %               |
| <b>Total Angiomax</b>              | <b>18,980</b>           | <b>30 %</b>       | <b>\$ 44,241</b>       | <b>68 %</b>       |
| <b>Cleviprex</b>                   |                         |                   |                        |                   |
| Clinical trials                    | 9,870                   | 16 %              | 7,535                  | 12 %              |
| Manufacturing development          | 1,108                   | 2 %               | 568                    | 1 %               |
| Administrative and headcount costs | 4,512                   | 7 %               | 856                    | 1 %               |
| <b>Total Cleviprex</b>             | <b>15,490</b>           | <b>25 %</b>       | <b>8,959</b>           | <b>14 %</b>       |
| <b>Cangrelor</b>                   |                         |                   |                        |                   |
| Clinical trials                    | 14,222                  | 22 %              | 1,090                  | 2 %               |
| Manufacturing development          | 2,153                   | 3 %               | 1,867                  | 3 %               |
| Administrative and headcount costs | 3,579                   | 6 %               | 700                    | 1 %               |
| <b>Total Cangrelor</b>             | <b>19,954</b>           | <b>31 %</b>       | <b>3,657</b>           | <b>6 %</b>        |
| <b>Other</b>                       | <b>9,112</b>            | <b>14 %</b>       | <b>7,532</b>           | <b>12 %</b>       |
| <b>Total</b>                       | <b>\$ 63,536</b>        | <b>100 %</b>      | <b>\$ 64,389</b>       | <b>100 %</b>      |

*Angiomax:*

Research and development spending in 2006 related to Angiomax decreased significantly due to a decrease in clinical trial expenses reflecting the completion in 2005 of enrollment in two clinical trial programs, including our 13,819 patient Phase III ACUITY trial. We continued to have research and development expenses during 2006 for ACUITY relating primarily to data analysis, but at significantly reduced rates. The investigators continued to conduct the ACUITY trial in 2006 as they collected 12-month patient follow-up results.

We also continued to incur research and development expense relating to Angiomax in connection with our efforts to expand the indications for which Angiomax is approved. In October 2006, we received a non-approvable letter from the FDA in connection with our application to market Angiomax in patients with or at risk of heparin-induced thrombocytopenia and thrombosis syndrome, or HIT/HITTS, undergoing cardiac surgery. In the letter, the FDA stated that it does not consider the data we submitted in support of the application adequate to support approval for this indication because it did not consider the evidence used to qualify patients for inclusion in the trials as a persuasive indicator for the risk of HIT/HITTS. We have indicated to the FDA that we are evaluating potential next steps.

We are preparing to study Angiomax in the pediatric setting and are working with the FDA to develop an appropriate study program. We are also supporting an investigator-initiated trial called HORIZONS to study Angiomax use in adult acute myocardial infarction, or AMI, patients. HORIZONS is designed to evaluate whether Angiomax with provisional use of GPIIb/IIIa inhibitors is as safe and effective as heparin or enoxaparin with planned use of GPIIb/IIIa inhibitors in AMI patients.

We expect spending for Angiomax to continue to decrease as a percentage of our research and development expense.

*Cleviprex*

*Research and development expenditures for Cleviprex increased during 2006 as we continued evaluating Cleviprex in anticipation of submitting an NDA with the FDA in the first half of 2007. During 2006 we continued development of Cleviprex through the following trials:*

- We completed three Phase III 500-patient clinical trials known as the ECLIPSE trials to evaluate the safety of Cleviprex in comparison to sodium nitroprusside, nicardipine and nitroglycerine during and following cardiac surgery. We had voluntarily suspended enrollment in these trials in March 2005 after a planned interim analysis of approximately half of the study population showed more frequent atrial fibrillation among patients randomized to Cleviprex than patients randomized to comparator drugs. After completing our interim review of the results of the safety studies, we found no significant differences in interim safety results between the clevidipine and the comparator arms. We resumed enrolling patients in December 2005 and completed enrollment in July 2006.
- We completed enrollment of our sixth Phase III clinical trial of Cleviprex in 100 patents with severe hypertension, known as the VELOCITY trial. We commenced enrollment in this trial in September 2006 and completed enrollment in January 2007.

*We expect research and development expenses for Cleviprex in 2007 primarily will include costs associated with the preparation of submissions for marketing approval for Cleviprex, manufacturing, Phase IIIb trials of Cleviprex in neurology and cardiology, along with a health economics study, and an observational study and clinical survey on treatment practices for acute severe hypertension conducted by third party researchers.*

*Cangrelor*

*Research and development expenditures related to cangrelor increased as a result of two separate Phase III clinical trials for the evaluation of cangrelor's effectiveness and safety in preventing ischemic events in patients who require PCI. In March 2006, we commenced enrollment of approximately 9,000-patients in our CHAMPION-PCI trial which we designed to evaluate whether use of intravenous cangrelor is superior to use of clopidogrel tablets in patients undergoing PCI. We commenced enrollment in October 2006 of a second trial, called CHAMPION-PLATFORM, which compares cangrelor plus usual care to placebo plus usual care in patients who require PCI. We currently expect to enroll approximately 6,500 patients in this trial.*

*We had enrolled approximately 2,000 patients in CHAMPION-PCI and approximately 150 patients in CHAMPION-PLATFORM at the end of 2006. We plan to enroll in excess of 8,000 patients in the aggregate in these trials in 2007 and expect to complete patient enrollment in both trials in 2008.*

*Other.* Spending in this category consists of infrastructure costs in support of our product development efforts which includes expenses for data management, statistical analysis and product safety as well as expenses related to business development activities. Increases in 2006 were primarily driven by an increase in personnel costs to support regulatory compliance medical writing, in addition to expenses related to stock-based compensation. Additionally, we incur business development expenses in connection with our efforts to evaluate early stage compounds and evaluations of strategic opportunities for the development and commercialization.

In order to support the continued development of Angiomax, Cleviprex and cangrelor, we expect our research and development expense to increase in 2007 from 2006 levels. We expect this increase in our research and development expenses to be primarily attributable to costs associated with enrollment of our ongoing Phase III clinical trials for cangrelor, the 9,000 patient CHAMPION-PCI trial and the 6,500 patient CHAMPION-PLATFORM trial, and additional manufacturing development costs for Cleviprex

and cangrelor. We also anticipate that stock based compensation expense included in research and development expenses will increase in 2007 as a result of a higher stock price as compared to 2006 stock option grants and anticipated stock option grants to new and current employees.

Our success in expanding the approved indications for Angiomax, or developing and obtaining marketing approval for Cleviprex and cangrelor, is highly uncertain. In particular, estimating our future levels of spending on development of Angiomax is uncertain following completion of the ACUITY trial. We cannot predict expenses associated with ongoing data analysis or regulatory submissions, if any. Nor can we reasonably estimate or know the nature, timing and estimated costs of the efforts necessary to complete the development of, or the period in which material net cash inflows are expected to commence from, either Cleviprex or cangrelor due to the numerous risks and uncertainties associated with developing drugs, including the uncertainty of:

- the scope, rate of progress and cost of our clinical trials and other research and development activities;
- future clinical trial results;
- the terms and timing of any collaborative, licensing and other arrangements that we may establish;
- the cost and timing of regulatory approvals;
- the cost and timing of establishing sales, marketing and distribution capabilities;
- the cost of establishing clinical and commercial supplies of our product candidates;
- the effect of competing technological and market developments; and
- the cost of filing, prosecuting, defending and enforcing any patent claims and other intellectual property rights.

*Selling, General and Administrative Expenses.* Selling, general and administrative expenses increased by 40% to \$88.3 million for 2006, from \$63.1 million for 2005. The increase in selling, general and administrative expenses of \$25.2 million was primarily due to an increase in Angiomax selling and promotional expenses, increases in educational grants, Cleviprex market research expenses, increased infrastructure costs and \$6.6 million of stock-based compensation.

We expect sales, general and administrative expenses to increase in 2007 from 2006 anticipated levels primarily due to Cleviprex-related pre-launch expenditures, continued promotional spending on Angiomax and increased sales force compensation.

We also expect total stock-based compensation expense included in selling, general and administrative expenses to increase in 2007 as a result of a higher stock price as compared to 2006 grants and anticipated stock option grants to new and current employees.

*Other Income.* Other income, which is primarily comprised of interest income, increased approximately 69% to \$7.3 million for 2006, from \$4.3 million for 2005. The increase in other income of \$3.0 million was primarily due to higher rates of return on our available for sale securities in 2006, combined with higher levels of cash to invest as a result of our generation of operating and investing cash flows.

*Benefit from/(Provision for) Income Tax.* The tax benefit for 2006 was \$46.1 million as compared to a \$0.1 million provision for 2005. The benefit for 2006 was a result of our decision to reduce approximately \$49.2 million of our valuation allowance against our deferred tax assets because we believe it is more likely than not that we will realize a benefit from these assets. This was partially offset by a provision for U.S. alternative minimum taxes, which can not entirely be offset with our NOL carryforwards, and state taxes based on net worth, while the provision for 2005 was primarily comprised of state taxes based on net worth.

We currently do not anticipate income tax benefits of this magnitude in the foreseeable future. We expect that future periods will include taxes at a higher rate than the effective rate for 2006. Due to the recognition of a portion of the deferred tax assets in 2006, we expect an effective income tax rate of 35-38% in 2007.

We will continue to evaluate the realizability of our deferred tax assets and liabilities on a periodic basis, and will adjust such amounts in light of changing facts and circumstances, including but not limited to future projections of taxable income, tax legislation, rulings by relevant tax authorities and the progress of ongoing tax audits. If we further reduce or increase the valuation allowance of deferred tax assets in future years, we would recognize a tax benefit or expense.

## Results of Operations

### Years Ended December 31, 2005 and 2004

*Net Revenue.* Net revenue increased 4% to \$150.2 million for 2005 as compared to \$144.3 million for 2004. In 2005, we derived approximately \$140.7 million of net revenue from U.S. sales of Angiomax and approximately \$9.5 million of net revenue from international sales. In 2004, we derived approximately \$135.7 million of net revenue from U.S. sales of Angiomax and approximately \$8.6 million of net revenue from international sales. We believe that the increase in U.S. sales in 2005 was due primarily to increased purchases of Angiomax by existing hospital customers, adoption of Angiomax by new hospital customers and the effects of higher prices as a result of a 10% price increase to our wholesalers in December 2004, partly offset by the impact of reduced purchases by wholesalers in connection with the fee-for-service arrangements that we entered into with wholesalers in 2005. We estimate that in implementing the planned inventory reduction, our wholesalers reduced their aggregate inventories by approximately \$26.0 million over the last two quarters of 2005. One of our European distributors, Nycomed, began stocking Angiox in the second half of 2004 in advance of the expected product launch in Europe, increasing our international revenue in 2004. Our international revenue during 2005, while higher than our international revenue in 2004, reflected lower rates of purchases by Nycomed following its initial stocking of the product.

### Net Revenue

| Net Revenue              | Year Ended December 31, |                       |                        |                       |
|--------------------------|-------------------------|-----------------------|------------------------|-----------------------|
|                          | 2005<br>(in thousands)  | % of Total<br>Revenue | 2004<br>(in thousands) | % of Total<br>Revenue |
| <b>Angiomax</b>          |                         |                       |                        |                       |
| United States Sales      | \$ 140,721              | 94 %                  | \$ 135,666             | 94 %                  |
| International Sales      | 9,486                   | 6 %                   | 8,585                  | 6 %                   |
| <b>Total Net Revenue</b> | \$ 150,207              | 100 %                 | \$ 144,251             | 100 %                 |

In 2005 and 2004, we recognized \$0.4 million and \$0.3 million, respectively, of international revenue from the amortization of milestone payments related to the \$2.5 million and \$1.5 million in non-refundable fees received from Nycomed. These milestone payments were recorded as deferred revenue in 2004 and 2002, respectively, and are being recognized ratably over the estimated term of our agreement with Nycomed.

*Cost of Revenue.* As shown in the table below, cost of revenue in 2005 was \$34.8 million, or 23% of net revenue, compared to \$29.1 million, or 20% of net revenue, in 2004. Cost of revenue consisted of expenses in connection with the manufacture of Angiomax sold, royalty expenses under our agreement with Biogen Idec and the logistics costs of selling Angiomax, such as distribution, storage, and handling.

#### Cost of Revenue

| Cost of Revenue              | Year Ended December 31, |                 | 2004<br>(in thousands) | % of Total Cost |
|------------------------------|-------------------------|-----------------|------------------------|-----------------|
|                              | 2005<br>(in thousands)  | % of Total Cost |                        |                 |
| Manufacturing                | \$ 14,223               | 41 %            | \$ 14,136              | 49 %            |
| Royalty                      | 16,142                  | 46 %            | 10,785                 | 37 %            |
| Logistics                    | 4,397                   | 13 %            | 4,202                  | 14 %            |
| <b>Total Cost of Revenue</b> | <b>\$ 34,762</b>        | <b>100 %</b>    | <b>\$ 29,123</b>       | <b>100 %</b>    |

The increase in cost of revenue as a percentage of net revenue was primarily a result of higher royalties based on a higher effective royalty rate under our licensing agreement with Biogen Idec offset partially by Angiomax sales price increases in December 2004.

*Research and Development Expenses.* Research and development expenses increased 31% to \$64.4 million for 2005, from \$49.3 million for 2004. The increase in research and development expenses was primarily due to a net \$12.3 million increase of Angiomax clinical trial costs in 2005, including a \$14.0 million increase in costs for ACUITY and a \$2.4 million increase in costs for a study of Angiomax in AMI patients, offset by a \$2.1 million decrease in expenses related to a Phase III trial program that we conducted in 2004 studying the use of Angiomax as an anticoagulant in patients undergoing cardiac surgery and in the treatment of patients with HIT/HITTS who were undergoing cardiac surgery, and a \$2.0 million decrease in other Angiomax studies. Angiomax manufacturing and development costs decreased by \$1.2 million in 2005 compared to 2004 due to lower expenses related to the development of an alternative method of manufacture and commercial supply with Lonza. The overall increase in research and development expenses for 2005 was also due to an increase of \$0.4 million in costs related to cangrelor development, \$2.1 million of increased infrastructure costs for data management, statistical analysis and product safety related costs and a \$2.6 million increase in business development activities.

The following table identifies for each of our major research and development projects, our spending for 2005 and 2004. Spending for past periods is not necessarily indicative of spending in future periods.

#### Research and Development Spending

| Research and Development           | Year Ended December 31, |                | 2004<br>(in thousands) | % of Total R&D |
|------------------------------------|-------------------------|----------------|------------------------|----------------|
|                                    | 2005<br>(in thousands)  | % of Total R&D |                        |                |
| <b>Angiomax</b>                    |                         |                |                        |                |
| Clinical trials                    | \$ 37,377               | 58 %           | \$ 25,069              | 51 %           |
| Manufacturing development          | 936                     | 1 %            | 2,088                  | 4 %            |
| Administrative and headcount costs | 5,928                   | 9 %            | 6,950                  | 14 %           |
| <b>Total Angiomax</b>              | <b>\$ 44,241</b>        | <b>68 %</b>    | <b>\$ 34,107</b>       | <b>69 %</b>    |
| <b>Cleviprex</b>                   | 8,959                   | 14 %           | 9,101                  | 18 %           |
| <b>Cangrelor</b>                   | 3,657                   | 6 %            | 3,210                  | 7 %            |
| <b>Other</b>                       | 7,532                   | 12 %           | 2,872                  | 6 %            |
|                                    | <b>\$ 64,389</b>        | <b>100 %</b>   | <b>\$ 49,290</b>       | <b>100 %</b>   |





*Selling, General and Administrative Expenses.* Selling, general and administrative expenses increased 25% to \$63.1 million for 2005, from \$50.3 million for 2004. The increase in selling, general and administrative expenses of \$12.8 million was due to a \$7.5 million increase in employee-related expense in connection with our sales force expansion in 2005, which resulted in an increase in the size of our sales force by approximately 50%, and a \$5.3 million increase in other expenses related to Angiomax promotion, headcount additions and legal costs.

*Non-cash Stock Compensation.* We recorded amortization expense for deferred compensation of approximately \$0.7 million for the year ended December 31, 2004. The amortization expense was included in our operating expenses in the consolidated statements of operations and related to deferred stock compensation that was recorded in 2000 and amortized over the respective vesting periods of the individual stock options. As of December 31, 2004 there was no additional deferred stock compensation expense to be amortized.

In September 2003, we amended the terms of fully-vested options to purchase 10,000 shares of common stock that were granted to a non-employee consultant in May 2001. In May 2003, we granted options to a non-employee consultant to purchase 50,000 shares at an exercise price based on the fair market value of our common stock on the date of the consulting agreement in April 2003. In addition, in November and December of 2005 we granted options to a non-employee consultant to purchase 7,100 shares at an exercise price based on the fair market value of our common stock. In each case, these options were valued utilizing the Black-Scholes option-pricing model. We recorded the \$35,200 and \$0.1 million in non-cash stock compensation expense associated with these options during 2005 and 2004, respectively. All of these options have now fully vested and we will not be required to record any additional associated non-cash stock compensation expense.

*Other Income.* Other income, which is comprised almost completely of interest income, increased approximately 105% to \$4.3 million for 2005, from \$2.1 million for 2004. The increase in interest income of \$2.2 million was primarily due to higher rates of return on our available for sale securities in 2005.

*Provision for Income Tax.* Tax expense for 2005 was \$0.1 million as compared to \$0.7 million for 2004. The provision for 2005 mostly reflected state taxes based on net worth, and the provision for 2004 mostly reflected U.S. alternative minimum taxes based on our first full year of profitability and state taxes based on net worth.

## **Liquidity and Capital Resources**

*Sources of Liquidity.* Since our inception, we have financed our operations principally through the sale of common and preferred stock, sales of convertible promissory notes and warrants, interest income and revenues from sales of Angiomax. Except for 2006 and 2004, we have incurred losses on an annual basis since our inception. We had \$196.8 million in cash, cash equivalents and available for sale securities as of December 31, 2006.

*Cash Flows.* As of December 31, 2006, we had \$75.5 million in cash and cash equivalents, as compared to \$25.7 million as of December 31, 2005. Our primary sources of cash during 2006 included net cash provided by operating activities of \$32.1 million, which was partially offset by \$6.3 million in net cash used in investing activities, and \$24.0 million in net cash provided by financing activities.

During 2006, we generated cash through operating activities of \$32.1 million, primarily through net profit of \$63.7 million, which was offset by non-cash items of \$37.5 million, consisting mainly of a deferred tax benefit of \$46.6 million and stock compensation expense of \$8.5 million, and an increase to certain working capital items of \$6.3 million.

During 2006, we used \$6.3 million in net cash in investing activities, which consisted of the purchase of \$149.8 million of available for sale securities, offset by \$144.3 million in proceeds from the maturation and

sale of available for sale securities, and purchases of \$0.8 million of fixed assets, mostly related to leasehold improvements and computer equipment for IT infrastructure.

During 2006, we received \$24.0 million in cash provided by financing activities, which consisted of net proceeds to us related to purchases of stock pursuant to option exercises and our employee stock purchase plan.

*Funding Requirements.* We expect to devote substantial resources to our research and development efforts and to our sales, marketing and manufacturing programs associated with the commercialization of our products. Our funding requirements will depend on numerous factors including:

- the extent to which Angiomax is commercially successful in the United States;
- the extent to which our international partners, including Nycomed, are commercially successful;
- the progress, level and timing of our research and development activities related to our clinical trials with respect to Angiomax, Cleviprex and cangrelor;
- the cost and outcomes of regulatory submissions and reviews;
- the continuation or termination of third party manufacturing or sales and marketing arrangements;
- the size, cost and effectiveness of our sales and marketing programs;
- the status of competitive products;
- our ability to defend and enforce our intellectual property rights;
- our decision whether to establish an infrastructure outside the United States; and
- the establishment of additional strategic or licensing arrangements with other companies, or acquisitions.

If our existing resources are insufficient to satisfy our liquidity requirements due to slower than anticipated revenues from Angiomax or otherwise, or if we acquire additional product candidates, or if we otherwise believe that raising additional capital would be in our interests and the interests of our stockholders, we may sell equity or debt securities or seek financing through other arrangements. Any sale of additional equity or debt securities may result in dilution to our stockholders, and debt financing may involve covenants limiting or restricting our ability to take specific actions, such as incurring additional debt or making capital expenditures. We cannot be certain that public or private financing will be available in amounts or on terms acceptable to us, if at all. If we seek to raise funds through collaboration or licensing arrangements with third parties, we may be required to relinquish rights to products, product candidates or technologies that we would not otherwise relinquish or grant licenses on terms that may not be favorable to us. If we are unable to obtain additional financing, we may be required to delay, reduce the scope of, or eliminate one or more of our planned research, development and commercialization activities, which could harm our financial condition and operating results.

In January 2007 we announced our intention to commence an offering of 6,000,000 shares of our common stock pursuant to an effective shelf registration in an underwritten public offering. However, we subsequently determined not to proceed with the public offering of common stock at that time.

#### **Contractual Obligations**

Our long-term contractual obligations include commitments and estimated purchase obligations entered into in the normal course of business. These include commitments related to purchases of inventory of our products, research and development service agreements, operating leases and selling, general and administrative obligations.



Future estimated contractual obligations as of December 31, 2006 are:

| Contractual Obligations (in thousands) | Total     | Less than 1 year | 1 - 3 Years | 3 - 5 Years | More than 5 years |
|--|-----------|------------------|-------------|-------------|-------------------|
| Inventory related commitments          | \$ 21,160 | \$ 16,816        | \$ 4,344    | \$          | \$                |
| Research and development               | 33,048    | 24,139           | 8,909       |             |                   |
| Operating leases                       | 15,860    | 3,020            | 8,458       | 4,245       | 137               |
| Selling, general and administrative    | 2,199     | 1,917            | 282         |             |                   |
| Total contractual obligations          | \$ 72,267 | \$ 45,892        | \$ 21,993   | \$ 4,245    | \$ 137            |

Included above are inventory-related non-cancellable commitments due to Lonza Braine totaling \$14.8 million during 2007 and \$4.3 million during 2008 and \$2.0 million in remaining Angiomax-related filling, finishing and packaging non-cancellable commitments through 2007. Of total estimated contractual obligations for research and development activities, \$7.5 million is non-cancellable. Of estimated contractual obligations for consulting, employment and professional services agreements associated with selling, general and administrative activities, \$0.8 million is non-cancellable.

In addition to the contractual obligations above, we have agreed to make payments upon the achievement of sales and regulatory milestones, and agreed to pay royalties, to Biogen Idec under our product license agreement for Angiomax and to AstraZeneca under our product license agreements for Cleviprex and cangrelor. Under the Angiomax license, we have agreed to pay up to an additional \$8.0 million upon the first commercial sales of Angiomax for the treatment of AMI in the United States and Europe. Under the Cleviprex license, we have agreed to pay up to an additional \$5.0 million upon reaching certain regulatory milestones, including a payment of \$1.5 million within 30 days of the acceptance for review of the NDA we plan to submit to the FDA for Cleviprex. Under the cangrelor license, we agreed to make milestone payments upon regulatory approval in major markets. The foregoing amounts do not include royalties that we may also have to pay.

#### Item 7A. Quantitative and Qualitative Disclosure About Market Risk

Market risk is the risk of change in fair value of a financial instrument due to changes in interest rates, equity prices, creditworthiness, financing, exchange rates or other factors. Our primary market risk exposure relates to changes in interest rates in our cash, cash equivalents and available for sale securities. We place our investments in high-quality financial instruments, primarily money market funds, corporate debt and U.S. government agency securities with maturities of less than two years, which we believe are subject to limited interest rate and credit risk. We currently do not hedge interest rate exposure. At December 31, 2006, we held \$196.8 million in cash, cash equivalents and available for sale securities which had an average interest rate of approximately 4.4%. Of this amount, approximately 96% of the cash, cash equivalents and available for sale securities were due on demand or within one year and had an average interest rate of approximately of 4.4%. The remaining 4% were due within two years and had an average interest rate of approximately 5%.

Most of our transactions are conducted in U.S. dollars. We do have certain agreements with parties located outside the United States. Transactions under certain of these agreements are conducted in U.S. dollars, subject to adjustment based on significant fluctuations in currency exchange rates. Transactions under certain other of these agreements are conducted in the local foreign currency. If the applicable exchange rate undergoes a change of 10.0%, we do not believe that it would have a material impact on our results of operations or cash flows.

**Item 8.** Financial Statements and Supplementary Data

All financial statements and schedules required to be filed hereunder are filed as Appendix A to this annual report on Form 10-K and incorporated herein by this reference.

**Item 9.** Changes in and Disagreements with Accountants on Accounting and Financial Disclosure

None.

**Item 9A.** Controls and Procedures

*Disclosure Controls and Procedures*

Our management, with the participation of our chief executive officer and chief financial officer, evaluated the effectiveness of our disclosure controls and procedures as of December 31, 2006. The term "disclosure controls and procedures," as defined in Rules 13a-15(e) and 15d-15(e) under the Exchange Act, means controls and other procedures of a company that are designed to ensure that information required to be disclosed by us in the reports that we file or submit under the Exchange Act is recorded, processed, summarized and reported within the time periods specified in the SEC's rules and forms. Disclosure controls and procedures include, without limitation, controls and procedures designed to ensure that information required to be disclosed by a company in the reports that it files or submits under the Exchange Act is accumulated and communicated to the company's management, including its principal executive and principal financial officers, as appropriate to allow timely decisions regarding required disclosure. Management recognizes that any controls and procedures, no matter how well designed and operated, can provide only reasonable assurance of achieving their objectives and management necessarily applies its judgment in evaluating the cost-benefit relationship of possible controls and procedures. Based on the evaluation of our disclosure controls and procedures as of December 31, 2006, our chief executive officer and chief financial officer concluded that, as of such date, our disclosure controls and procedures were effective at the reasonable assurance level.

*Management's Annual Report on Internal Control Over Financial Reporting*

The report required to be filed hereunder is included in Appendix A to this annual report on Form 10-K and incorporated herein by this reference.

*Attestation Report of Our Registered Public Accounting Firm*

The report required to be filed hereunder is included in Appendix A to this annual report on Form 10-K and incorporated herein by this reference.

*Changes in Internal Control Over Financial Reporting*

No change in our internal control over financial reporting (as defined in Rules 13a-15(f) and 15d-15(f) under the Exchange Act) occurred during the quarter ended December 31, 2006 that has materially affected, or is reasonably likely to materially affect, our internal control over financial reporting.

**Item 9B.** Other Information

Not applicable.

**PART III**

Pursuant to Paragraph G(3) of the General Instructions to Form 10-K, the information required by Part III (Items 10, 11, 12, 13 and 14) is being incorporated by reference herein from our proxy statement to be filed with the Securities and Exchange Commission within 120 days of the end of the fiscal year



ended December 31, 2006 in connection with our 2007 Annual Meeting of Stockholders (our 2006 Proxy Statement ).

**Item 10. Directors, Executive Officers and Corporate Governance**

The information required by this item will be contained in our 2007 Proxy Statement under the captions Discussion of Proposals, Information About Corporate Governance, Information About Our Executive Officers and Other Information and is incorporated herein by this reference.

We have adopted a code of business conduct and ethics applicable to all of our directors and employees, including our principal executive officer, principal financial officer and our controller. The code of business conduct and ethics is available on the corporate governance section of Investor Relations of our website, *www.themedicinescompany.com*.

Any waiver of the code of business conduct and ethics for directors or executive officers, or any amendment to the code that applies to directors or executive officers, may only be made by the board of directors. We intend to satisfy the disclosure requirement under Item 5.05 of Form 8-K regarding an amendment to, or waiver from, a provision of this code of ethics by posting such information on its website, at the address and location specified above. To date, no such waivers have been requested or granted.

**Item 11. Executive Compensation**

The information required by this item will be contained in our 2007 Proxy Statement under the captions Corporate Governance, Information About Our Executive Officers and Other Information and is incorporated herein by this reference.

**Item 12. Security Ownership of Certain Beneficial Owners and Management and Related Stockholder Matters**

The information required by this item will be contained in our 2007 Proxy Statement under the captions Information About Our Executive Officers and Other Information and is incorporated herein by this reference.

**Item 13. Certain Relationships and Related Transactions, and Director Independence**

The information required by this item will be contained in our 2007 Proxy Statement under the caption Information About Our Executive Officers and is incorporated herein by this reference.

**Item 14. Principal Accountant Fees and Services**

The information required by this item will be contained in our 2007 Proxy Statement under the caption Discussion of Proposals and is incorporated herein by this reference.

**PART IV**

**Item 15. Exhibits, Financial Statement Schedules**

(a) Documents filed as part of this annual report:

(1) Financial Statements. The Consolidated Financial Statements are included as Appendix A hereto and are filed as part of this annual report. The Consolidated Financial Statements include:

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| <u>Report of Independent Registered Public Accounting Firm</u>  | F-3         |
| <u>Report of Independent Registered Public Accounting Firm on Internal Control over Financial Reporting</u>   | F-4         |
| <u>Consolidated Balance Sheets</u>  | F-5         |
| <u>Consolidated Statements of Operations</u>  | F-6         |
| <u>Consolidated Statements of Stockholders' Equity</u>  | F-7         |
| <u>Consolidated Statements of Cash Flows</u>  | F-8         |
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(2) Financial Statement Schedule. The financial statement schedule following the Notes to Consolidated Financial Statements is filed as part of this annual report. All other schedules are omitted because they are not applicable or are not required, or because the required information is included in the consolidated financial statements or notes filed as part of this annual report.

(3) Exhibits. The exhibits set forth on the Exhibit Index following the signature page to this annual report are filed as part of this annual report. This list of exhibits identifies each management contract or compensatory plan or arrangement required to be filed as an exhibit to this annual report.



**SIGNATURES**

Pursuant to the requirements of Section 13 or 15(d) of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned, thereunto duly authorized, on March 1, 2007.

THE MEDICINES COMPANY

By: */s/ CLIVE A. MEANWELL*  
Clive A. Meanwell  
*Chief Executive Officer*

Pursuant to the requirements of the Securities Exchange Act of 1934, this report has been signed below by the following persons on behalf of the registrant and in the capacities indicated on March 1, 2007:

| <b>Signature</b>  | <b>Title(s)</b>  |
|---|--|
| <i>/s/ CLIVE A. MEANWELL</i><br>Clive A. Meanwell       | Chief Executive Officer and Chairman of the Board of Directors (Principal Executive Officer)                 |
| <i>/s/ GLENN P. SBLENDORIO</i><br>Glenn P. Sblendorio   | Executive Vice President, Chief Financial Officer and Treasurer (Principal Financial and Accounting Officer) |
| <i>/s/ JOHN P. KELLEY</i><br>John P. Kelley             | President, Chief Operating Officer and Director  |
| <i>/s/ WILLIAM W. CROUSE</i><br>William W. Crouse       | Director   |
| <i>/s/ ROBERT J. HUGIN</i><br>Robert J. Hugin           | Director   |
| <i>/s/ T. SCOTT JOHNSON</i><br>T. Scott Johnson         | Director   |
| <i>/s/ ARMIN M. KESSLER</i><br>Armin M. Kessler         | Director   |
| <i>/s/ ROBERT G. SAVAGE</i><br>Robert G. Savage         | Director   |
| <i>/s/ MELVIN K. SPIGELMAN</i><br>Melvin K. Spigelman   | Director   |
| <i>/s/ ELIZABETH H.S. WYATT</i><br>Elizabeth H.S. Wyatt | Director   |

APPENDIX A

INDEX TO THE  
CONSOLIDATED FINANCIAL STATEMENTS OF  
THE MEDICINES COMPANY

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**Management's Report on Consolidated Financial Statements and Internal Control over Financial Reporting**

The management of The Medicines Company has prepared, and is responsible for, The Medicines Company's consolidated financial statements and related footnotes. These consolidated financial statements have been prepared in conformity with U.S. generally accepted accounting principles.

The Medicines Company's management is responsible for establishing and maintaining adequate internal control over financial reporting. Internal control over financial reporting is defined in Rule 13a-15(f) or 15d-15(f) promulgated under the Securities Exchange Act of 1934 as a process designed by, or under the supervision of the Company's principal executive and principal financial officers and effected by the Company's board of directors, management, and other personnel, to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles and includes those policies and procedures that:

- pertain to the maintenance of records that in reasonable detail accurately and fairly reflect the transactions and dispositions of the assets of The Medicines Company;
- provide reasonable assurance that transactions are recorded as necessary to permit preparation of financial statements in accordance with generally accepted accounting principles, and that receipts and expenditures of The Medicines Company are being made only in accordance with authorizations of management and directors of The Medicines Company; and
- provide reasonable assurance regarding prevention or timely detection of unauthorized acquisition, use or disposition of The Medicines Company's assets that could have a material effect on the financial statements.

Because of its inherent limitations, internal control over financial reporting may not prevent or detect misstatements. Also, projections of any evaluation of effectiveness to future periods are subject to the risk that controls may become inadequate because of changes in conditions, or that the degree of compliance with the policies or procedures may deteriorate.

The Medicines Company's management assessed the Company's internal control over financial reporting as of December 31, 2006. Management's assessment was based upon the criteria established in *Internal Control - Integrated Framework* issued by the Committee of Sponsoring Organizations of the Treadway Commission. Based on its assessment, management concluded that, as of December 31, 2006, The Medicines Company's internal control over financial reporting is effective based on those criteria. The Company's assessment of the effectiveness over its financial reporting, as of December 31, 2006, has been audited by Ernst & Young LLP, an independent registered public accounting firm, that has audited The Medicines Company's financial statements, and their attestation report is included herein.

Dated February 23, 2007

/s/ Clive A. Meanwell  
Chairman and  
Chief Executive Officer

/s/ Glenn P. Sblendorio  
Executive Vice President-  
Chief Financial Officer

**Report of Independent Registered Public Accounting Firm**

**The Board of Directors and Stockholders of The Medicines Company**

We have audited the accompanying consolidated balance sheets of The Medicines Company as of December 31, 2006 and 2005, and the related consolidated statements of operations, consolidated statements of stockholders' equity and cash flows for each of the three years in the period ended December 31, 2006. Our audits also included the financial statement schedule listed in item 15 (a). These financial statements and schedule are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements and schedule based on our audits.

We conducted our audits 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 audits provide a reasonable basis for our opinion.

In our opinion, the financial statements referred to above present fairly, in all material respects, the consolidated financial position of The Medicines Company at December 31, 2006 and 2005, and the consolidated results of its operations and its consolidated cash flows for each of the three years in the period ended December 31, 2006 in conformity with U.S. generally accepted accounting principles. Also in our opinion, the related financial statement schedule when considered in relation to the basic financial statements taken as a whole, presents fairly in all material respects the information set forth therein.

As discussed in Note 2 to the Financial Statements, the Company changed its method of accounting for stock-based compensation on January 1, 2006.

We also have audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States), the effectiveness of The Medicines Company's internal control over financial reporting as of December 31, 2006, based on criteria established in Internal Control - Integrated Framework issued by the Committee of Sponsoring Organizations of the Treadway Commission and our report dated February 23, 2007 expressed an unqualified opinion thereon.

/s/ Ernst and Young LLP

MetroPark, NJ  
February 23, 2007

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**Report of Independent Registered Public Accounting Firm on Internal Control over Financial Reporting**

**The Board of Directors and Stockholders of The Medicines Company**

We have audited management's assessment, included in the accompanying Management's Report on Consolidated Financial Statements and Internal Control over Financial Reporting, that The Medicines Company maintained effective internal control over financial reporting as of December 31, 2006, based on criteria established in Internal Control - Integrated Framework issued by the Committee of Sponsoring Organizations of the Treadway Commission (the COSO criteria). The Medicines Company's management is responsible for maintaining effective internal control over financial reporting and for its assessment of the effectiveness of internal control over financial reporting. Our responsibility is to express an opinion on management's assessment and an opinion on the effectiveness of the company's internal control over financial reporting 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 effective internal control over financial reporting was maintained in all material respects. Our audit included obtaining an understanding of internal control over financial reporting, evaluating management's assessment, testing and evaluating the design and operating effectiveness of internal control, and performing such other procedures as we considered necessary in the circumstances. We believe that our audit provides a reasonable basis for our opinion.

A company's internal control over financial reporting is a process designed to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles. A company's internal control over financial reporting includes those policies and procedures that (1) pertain to the maintenance of records that, in reasonable detail, accurately and fairly reflect the transactions and dispositions of the assets of the company; (2) provide reasonable assurance that transactions are recorded as necessary to permit preparation of financial statements in accordance with generally accepted accounting principles, and that receipts and expenditures of the company are being made only in accordance with authorizations of management and directors of the company; and (3) provide reasonable assurance regarding prevention or timely detection of unauthorized acquisition, use, or disposition of the company's assets that could have a material effect on the financial statements.

Because of its inherent limitations, internal control over financial reporting may not prevent or detect misstatements. Also, projections of any evaluation of effectiveness to future periods are subject to the risk that controls may become inadequate because of changes in conditions, or that the degree of compliance with the policies or procedures may deteriorate.

In our opinion, management's assessment that The Medicines Company maintained effective internal control over financial reporting as of December 31, 2006, is fairly stated, in all material respects, based on the COSO criteria. Also, in our opinion, The Medicines Company maintained, in all material respects, effective internal control over financial reporting as of December 31, 2006, based on the COSO criteria.

We also have audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States), the 2006 consolidated financial statements of The Medicines Company and our report dated February 23, 2007 expressed an unqualified opinion thereon.

/s/ Ernst and Young LLP

MetroPark, NJ  
February 23, 2007

**THE MEDICINES COMPANY**  
**CONSOLIDATED BALANCE SHEETS**  
(in thousands, except per share amounts)

|   | <b>December 31,</b> |                   |
|---|---------------------|-------------------|
|   | <b>2006</b>         | <b>2005</b>       |
| <b>ASSETS</b>   |                     |                   |
| Current assets:   |                     |                   |
| Cash and cash equivalents   | \$ 75,530           | \$ 25,706         |
| Available for sale securities   | 121,287             | 114,384           |
| Accrued interest receivable   | 1,414               | 922               |
| Accounts receivable, net of allowance of approximately \$0.80 million and \$0.85 million at December 31, 2006 and 2005  | 21,504              | 14,611            |
| Inventory   | 41,628              | 47,985            |
| Prepaid expenses and other current assets   | 12,963              | 970               |
| <b>Total current assets</b>   | <b>274,326</b>      | <b>204,578</b>    |
| Fixed assets, net   | 3,071               | 3,990             |
| Deferred tax assets   | 41,032              |                   |
| Other assets  | 139                 | 139               |
| <b>Total assets</b>   | <b>\$ 318,568</b>   | <b>\$ 208,707</b> |
| <b>LIABILITIES AND STOCKHOLDERS EQUITY</b>  |                     |                   |
| Current liabilities:  |                     |                   |
| Accounts payable  | \$ 8,885            | \$ 5,989          |
| Accrued expenses  | 36,918              | 28,677            |
| <b>Total current liabilities</b>  | <b>45,803</b>       | <b>34,666</b>     |
| Commitments and contingencies   |                     |                   |
| Deferred revenue  | 2,814               | 3,142             |
| Stockholders' equity:   |                     |                   |
| Preferred stock, \$1.00 par value per share, 5,000,000 shares authorized; no shares issued and outstanding  |                     |                   |
| Common stock, \$.001 par value per share, 125,000,000 shares authorized; 51,227,313 and 49,723,756 issued and outstanding at December 31, 2006 and 2005, respectively | 51                  | 50                |
| Additional paid-in capital  | 511,076             | 476,012           |
| Accumulated deficit   | (241,172)           | (304,898)         |
| Accumulated other comprehensive (loss)  | (4)                 | (265)             |
| <b>Total stockholders' equity</b>   | <b>269,951</b>      | <b>170,899</b>    |
| <b>Total liabilities and stockholders' equity</b>   | <b>\$ 318,568</b>   | <b>\$ 208,707</b> |

*See accompanying notes*

**THE MEDICINES COMPANY**  
**CONSOLIDATED STATEMENTS OF OPERATIONS**  
(in thousands, except per share amounts)

|  | Year Ended December 31, |             |            |
|--|-------------------------|-------------|------------|
|  | 2006                    | 2005        | 2004       |
| Net revenue  | \$ 213,952              | \$ 150,207  | \$ 144,251 |
| Operating expenses:  |                         |             |            |
| Cost of revenue  | 51,812                  | 34,762      | 29,123     |
| Research and development   | 63,536                  | 64,389      | 49,290     |
| Selling, general and administrative                                | 88,265                  | 63,053      | 50,275     |
| Total operating expenses   | 203,613                 | 162,204     | 128,688    |
| Income/(loss) from operations                                      | 10,339                  | (11,997 )   | 15,563     |
| Other income/(expense):  |                         |             |            |
| Interest income  | 7,319                   | 4,344       | 2,126      |
| Income/(loss) before income taxes                                  | 17,658                  | (7,653 )    | 17,689     |
| Benefit from/(provision for) income taxes                          | 46,068                  | (100 )      | (690 )     |
| Net income/(loss)  | \$ 63,726               | \$ (7,753 ) | \$ 16,999  |
| Basic earnings/(loss) per common share                             | \$ 1.27                 | \$ (0.16 )  | \$ 0.36    |
| Shares used in computing basic earnings/(loss) per common share:   | 50,300                  | 49,443      | 47,855     |
| Diluted earnings/(loss) per common share                           | \$ 1.25                 | \$ (0.16 )  | \$ 0.34    |
| Shares used in computing diluted earnings/(loss) per common share: | 51,034                  | 49,443      | 49,772     |

*See accompanying notes*

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**THE MEDICINES COMPANY**  
**CONSOLIDATED STATEMENTS OF STOCKHOLDERS' EQUITY**  
**For The Years Ended December 31, 2004, 2005 and 2006**

(in thousands)

|   | Common Stock<br>Shares | Common Stock<br>Amount | Additional<br>Paid-in<br>Capital | Deferred Stock<br>Compensation | Accumulated<br>Deficit | Accumulated<br>Comprehensive<br>Income<br>(Loss) | Total<br>Stockholders<br>Equity |
|---|------------------------|------------------------|----------------------------------|--------------------------------|------------------------|--|---------------------------------|
| Balance at December 31, 2003                          | 47,444                 | \$ 47                  | \$ 454,804                       | \$ (744 )                      | \$ (314,144 )          | \$ 202   | \$ 140,165                      |
| Employee stock purchases                              | 1,097                  | 1                      | 13,643                           |                                |                        |  | 13,644                          |
| Issuance of common stock Warrant purchases            | 104                    |                        | 81                               |                                |                        |  | 81                              |
| Adjustments to deferred compensation for terminations |                        |                        | (19 )                            | 19                             |                        |  |                                 |
| Amortization of deferred stock compensation           |                        |                        |                                  | 725                            |                        |  | 725                             |
| Non-cash stock compensation Consultants               |                        |                        | 143                              |                                |                        |  | 143                             |
| Tax benefit from option exercises                     |                        |                        | 449                              |                                |                        |  | 449                             |
| Net income  |                        |                        |                                  |                                | 16,999                 |  | 16,999                          |
| Currency translation adjustment                       |                        |                        |                                  |                                |                        | 12   | 12                              |
| Unrealized loss on available for sale securities      |                        |                        |                                  |                                |                        | (547 )   | (547 )                          |
| Comprehensive income                                  |                        |                        |                                  |                                |                        |  | 16,464                          |
| Balance at December 31, 2004                          | 48,645                 | 48                     | 469,101                          |                                | (297,145 )             | (333 )   | 171,671                         |
| Employee stock purchases                              | 579                    | 1                      | 6,825                            |                                |                        |  | 6,826                           |
| Issuance of common stock Warrant purchases            | 500                    | 1                      | (1 )                             |                                |                        |  |                                 |
| Non-cash stock compensation Consultants               |                        |                        | 35                               |                                |                        |  | 35                              |
| Tax benefit from option exercises                     |                        |                        | 52                               |                                |                        |  | 52                              |
| Net loss  |                        |                        |                                  |                                | (7,753 )               |  | (7,753 )                        |
| Currency translation adjustment                       |                        |                        |                                  |                                |                        | (22 )  | (22 )                           |
| Unrealized gain on available for sale securities      |                        |                        |                                  |                                |                        | 90   | 90                              |
| Comprehensive loss                                    |                        |                        |                                  |                                |                        |  | (7,685 )                        |
| Balance at December 31, 2005                          | 49,724                 | 50                     | 476,012                          |                                | (304,898 )             | (265 )   | 170,899                         |
| Employee stock purchases                              | 1,503                  | 1                      | 23,964                           |                                |                        |  | 23,965                          |
| Non-cash stock compensation                           |                        |                        | 8,459                            |                                |                        |  | 8,459                           |
| Tax benefit from option exercises                     |                        |                        | 2,641                            |                                |                        |  | 2,641                           |
| Net income  |                        |                        |                                  |                                | 63,726                 |  | 63,726                          |
| Currency translation adjustment                       |                        |                        |                                  |                                |                        | 23   | 23                              |
| Unrealized gain on available for sale securities      |                        |                        |                                  |                                |                        | 238  | 238                             |
| Comprehensive income                                  |                        |                        |                                  |                                |                        |  | 63,987                          |
| Balance at December 31, 2006                          | 51,227                 | \$ 51                  | \$ 511,076                       | \$                             | \$ (241,172 )          | \$ (4 )  | \$ 269,951                      |

See accompanying notes.

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**THE MEDICINES COMPANY**  
**CONSOLIDATED STATEMENTS OF CASH FLOWS**  
(in thousands)

|  | Year Ended December 31, |             |            |
|--|-------------------------|-------------|------------|
|  | 2006                    | 2005        | 2004       |
| Cash flows from operating activities:  |                         |             |            |
| Net income/(loss)  | \$ 63,726               | \$ (7,753 ) | \$ 16,999  |
| Adjustments to reconcile net income/(loss) to net cash (used in)/provided by operating activities: |                         |             |            |
| Depreciation   | 1,465                   | 998         | 591        |
| Amortization of net premiums and discounts on available for sale securities                        | (1,160 )                | (18 )       | 1,281      |
| Non-cash stock compensation expense  | 8,459                   | 35          | 868        |
| Loss on disposal of fixed assets   | 244                     |             | 49         |
| Deferred tax provision   | (49,200)                |             |            |
| Tax benefit from option exercises  | 2,641                   | 52          | 449        |
| Changes in operating assets and liabilities:   |                         |             |            |
| Accrued interest receivable  | (492 )                  | (10 )       | 79         |
| Accounts receivable  | (6,893 )                | 3,776       | (2,727 )   |
| Inventory  | 6,357                   | (20,644 )   | (15,882 )  |
| Prepaid expenses and other current assets  | (3,825 )                | 280         | (275 )     |
| Other assets   |                         | 21          | 39         |
| Accounts payable   | 2,896                   | (5,524 )    | 5,166      |
| Accrued expenses   | 8,231                   | 5,356       | 4,446      |
| Deferred revenue   | (328 )                  | (374 )      | 2,246      |
| Net cash provided by (used in) operating activities  | 32,121                  | (23,805 )   | 13,329     |
| Cash flows from investing activities:  |                         |             |            |
| Purchases of available for sale securities   | (149,852 )              | (134,638 )  | (112,838 ) |
| Maturities and sales of available for sale securities  | 144,347                 | 144,171     | 79,666     |
| Purchases of fixed assets  | (790 )                  | (3,313 )    | (804 )     |
| Net cash (used in) provided by investing activities  | (6,295 )                | 6,220       | (33,976 )  |
| Cash flows from financing activities:  |                         |             |            |
| Proceeds from issuances of common stock, net   | 23,965                  | 6,825       | 13,725     |
| Net cash provided by financing activities  | 23,965                  | 6,825       | 13,725     |
| Effect of exchange rate changes on cash  | 33                      | (39 )       | 25         |
| Increase (decrease) in cash and cash equivalents   | 49,824                  | (10,799 )   | (6,897 )   |
| Cash and cash equivalents at beginning of period   | 25,706                  | 36,505      | 43,402     |
| Cash and cash equivalents at end of period   | \$ 75,530               | \$ 25,706   | \$ 36,505  |
| Supplemental disclosure of cash flow information:  |                         |             |            |
| Interest paid  | \$                      | \$          | \$         |
| Taxes paid   | \$ 395                  | \$ 316      | \$ 69      |

See accompanying notes.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS**  
**December 31, 2006**

**1. Nature of Business**

The Medicines Company (the Company) was incorporated in Delaware on July 31, 1996. The Company is a pharmaceutical company providing innovative, cost effective acute care hospital products to the worldwide hospital marketplace. In December 2000, the U.S. Food and Drug Administration (the FDA) approved the Company's product, Angiomax® (bivalirudin), a direct thrombin inhibitor, for use as an anticoagulant in combination with aspirin in patients with unstable angina undergoing coronary angioplasty. In 2005, the Company received approvals from the FDA for new prescribing information for Angiomax. The Company is currently developing Angiomax for use in additional patient populations. The Company markets Angiomax to interventional cardiology customers for its approved uses in PCI, including in patients with or at risk of HIT/HITTS. The Company has concentrated its commercial sales and marketing resources on the United States hospital market, and revenues to date have been generated principally from sales of Angiomax in the United States. In September 2004, the Company received authorization from the European Commission to market Angiomax® (bivalirudin) as Angiox® (bivalirudin) in the member states of the European Union for use as an anticoagulant in combination with aspirin in patients undergoing percutaneous coronary interventions. In addition to Angiomax, the Company is currently developing two other pharmaceutical products as potential acute care hospital products. The first of these, Cleviprex (clevidipine), is an intravenous drug intended for the control of blood pressure in intensive care patients who require rapid and precise control of blood pressure. The second potential product, cangrelor, is an intravenous antiplatelet agent that is intended to prevent platelet activation and aggregation, which the Company believes has potential advantages in the treatment of vascular disease.

The Company's net revenue of \$214.0 million in 2006, \$150.2 million in 2005 and \$144.3 million in 2004 was generated principally from sales of Angiomax in the United States. International sales and revenue resulting from the amortization of milestone payments included in total net revenue were \$11.3 million in 2006, \$9.5 million in 2005 and \$8.6 million in 2004. Additionally, during 2006, in collaboration with a third party, the Company received reimbursements for certain research and development expenses it had incurred. The reimbursements of \$1.9 million are reported as part of net revenue and the fees for the services are included in research and development expenses. The Company has invested, and plans to continue investing, in Angiomax development programs to expand the indications for which Angiomax is approved. Additionally, the Company plans to continue investing in the development of Cleviprex and cangrelor.

**2. Significant Accounting Policies**

*Basis of Presentation*

The consolidated financial statements include the accounts of the Company and its wholly-owned subsidiaries. All significant intercompany balances and transactions have been eliminated in consolidation. The Company has no unconsolidated subsidiaries or investments accounted for under the equity method.

*Use of Estimates*

The preparation of financial statements in conformity with U.S. generally accepted accounting principles requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and the reported amounts of revenues and expenses during the reporting period. Actual results could differ from those estimates.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

*Risks and Uncertainties*

The Company is subject to risks common to companies in the pharmaceutical industry including, but not limited to, uncertainties related to commercialization of products, regulatory approvals, dependence on key products, dependence on key customers and suppliers, and protection of intellectual property rights.

*Concentrations of Credit Risk*

Financial instruments that potentially subject the Company to concentration of credit risk include cash, cash equivalents, available for sale securities and accounts receivable. The Company believes it minimizes its exposure to potential concentrations of credit risk by placing investments in high-quality financial instruments with high quality institutions. At December 31, 2006 and 2005, approximately \$49 million and \$12.6 million, respectively, of the cash and cash equivalents balance was invested in a single fund, the Evergreen Institutional Money Market Fund, a no-load money market fund, with the Capital Advisors Group.

The Company sells Angiomax primarily to a limited number of domestic wholesalers with distribution centers located throughout the United States and to several international distributors. These wholesalers and distributors then sell Angiomax to hospitals. Three domestic wholesalers, AmerisourceBergen Drug Corporation, McKesson Corporation and Cardinal Health, Inc. accounted for 31%, 24% and 33%, respectively, of the Company's net revenue for the year ended December 31, 2006. Revenue from each of these customers accounted for similar percentages of net revenue in 2005 and 2004. During 2006, 2005 and 2004, the Company's net revenue from these three customers totaled approximately 88%, 90% and 77%, respectively, of net revenue. At December 31, 2006 and 2005, amounts due from these three wholesaler customers represented approximately \$20.8 million, or 89%, and \$15.2 million, or 98%, respectively, of gross accounts receivable. The Company's trade accounts receivable are reported net of allowances for chargebacks, cash discounts and doubtful accounts. The Company performs ongoing credit evaluations of its customers and generally does not require collateral. The Company maintains reserves for potential credit losses and, during 2006, such losses were within the expectations of management.

*Cash, Cash Equivalents and Available for Sale Securities*

The Company considers all highly liquid investments purchased with an original maturity at date of purchase of three months or less to be cash equivalents. Cash equivalents at December 31, 2006 included investments of \$75.5 million in money market funds and commercial paper with original maturities of less than three months. Cash equivalents at December 31, 2005 included investments of \$12.6 million in money market funds and \$2.0 million of corporate bonds with original maturities of less than three months. These investments are carried at cost, which approximates fair value. The Company measures all original maturities from the date the investment was originally purchased by the Company.

The Company considers securities with original maturities of greater than three months to be available for sale securities. Securities under this classification are recorded at fair market value and unrealized gains and losses are recorded as a separate component of stockholders' equity. The estimated fair value of the available for sale securities is determined based on quoted market prices or rates for similar instruments. In addition, the cost of debt securities in this category is adjusted for amortization of

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

premium and accretion of discount to maturity. The Company evaluates securities with unrealized losses to determine whether such losses are other than temporary.

At December 31, 2006 and December 31, 2005, the Company held available for sale securities with fair value totaling \$121.3 million and \$114.4 million, respectively. These available for sale securities included various corporate debt securities and United States government agency notes. At December 31, 2006, \$113.3 million of the Company's available for sale securities had maturities within one year and \$8.0 million had maturities which were more than one year but less than two years. At December 31, 2005 all of the Company's available for sale securities had maturities within one year. Available for sale securities, including estimated fair values, are summarized as follows:

| (in thousands)               | Cost              | Unrealized Loss        | Fair Value        |
|------------------------------|-------------------|------------------------|-------------------|
| <b>2006</b>                  |                   |                        |                   |
| Corporate debt securities    | \$ 62,023         | \$ (15 )               | \$ 62,008         |
| U.S. government agency notes | 59,301            | (22 )                  | 59,279            |
| <b>Total</b>                 | <b>\$ 121,324</b> | <b>\$ (37 )</b>        | <b>\$ 121,287</b> |
|                              | <b>Cost</b>       | <b>Unrealized Loss</b> | <b>Fair Value</b> |
| <b>2005</b>                  |                   |                        |                   |
| Corporate debt securities    | \$ 23,873         | \$ (69 )               | \$ 23,804         |
| U.S. government agency notes | 90,786            | (206 )                 | 90,580            |
| <b>Total</b>                 | <b>\$ 114,659</b> | <b>\$ (275 )</b>       | <b>\$ 114,384</b> |

**Revenue Recognition**

*Product Sales.* The Company sells its products primarily to domestic wholesalers and international distributors, who, in turn, sell to hospitals. The Company does not recognize revenue from product sales until there is persuasive evidence of an arrangement, delivery has occurred, the price is fixed and determinable, the buyer is obligated to pay the Company, the obligation to pay is not contingent on resale of the product, the buyer has economic substance apart from the Company, the Company has no obligation to bring about sale of the product, the amount of returns can be reasonably estimated and collectibility is reasonably assured.

*Domestic Sales.* The Company records allowances for chargebacks and other discounts, and accruals for product returns, rebates and fee-for-service charges at the time of sale, and reports revenue net of such amounts. In determining the amounts of certain allowances and accruals, the Company must make significant judgments and estimates. For example, in determining these amounts, the Company estimates hospital demand, buying patterns by hospitals and group purchasing organizations from wholesalers and the levels of inventory held by wholesalers. Making these determinations involves estimating whether trends in past wholesaler and hospital buying patterns will predict future product sales. In 2005, the Company agreed with its largest wholesalers to enter into fee-for-service arrangements under which these wholesalers agreed to provide the Company with more frequent data on wholesaler inventory levels and hospital purchases. As these arrangements are implemented, the Company expects to apply this data in determining the amounts of certain of these allowances and accruals.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

The nature of the Company's allowances and accruals requiring critical estimates, and the specific considerations it uses in estimating their amounts, are as follows:

- *Product returns.* The Company's customers have the right to return any unopened product during the 18-month period beginning six months prior to the labeled expiration date and ending 12 months after the labeled expiration date. As a result, in calculating the accrual for product returns, the Company must estimate the likelihood that product sold to wholesalers might remain in their inventory to within six months of expiration and analyze the likelihood that such product will be returned within 12 months after expiration.

In estimating the likelihood of product remaining in wholesalers' inventory, the Company relies on information from wholesalers regarding their inventory levels, measured hospital demand as reported by third party sources and on internal sales data. The Company also considers its wholesalers' past buying patterns, estimated remaining shelf life of product previously shipped and the expiration dates of product currently being shipped.

In estimating the likelihood of product return, the Company relies primarily on historic patterns of returns and estimated remaining shelf life of product previously shipped.

At December 31, 2006 and 2005, the Company's accrual for product returns was \$0.4 million and \$0.2 million, respectively.

- *Chargebacks and rebates.* Although the Company sells Angiomax primarily to wholesalers and distributors, the Company typically enters into agreements with hospitals, either directly or through group purchasing organizations acting on behalf of their hospital members, in connection with the hospitals' purchases of Angiomax from the Company's wholesalers. Based on the terms of these agreements, most of the Company's hospital customers have the right to receive a discounted price and volume based rebate on product purchases. The Company provides a credit to the wholesaler, or a chargeback, representing the difference between the wholesaler's acquisition list price and the discounted price.

As a result of these contracts, at the time of product shipment, the Company must estimate the likelihood that Angiomax sold to wholesalers might be ultimately sold to a contracting hospital or group purchasing organization. The Company must also estimate the contracting hospital's or group purchasing organization's volume of purchases.

The Company bases its estimates on the historic chargeback data it receives from wholesalers, which detail historic buying patterns and sales mix for particular hospitals and group purchasing organizations, and the applicable customer chargeback rates and rebate thresholds.

At December 31, 2006 and 2005, the Company's allowance for chargebacks was \$0.3 million and \$0.5 million, respectively, and its accrual for rebates was \$0.8 million and \$1.5 million, respectively.

The Company has adjusted its allowances for chargebacks and accruals for product returns and rebates in the past based on actual sales experience, and the Company will likely be required to make adjustments to these allowances and accruals in the future. The Company continually monitors its allowances and accruals and makes adjustments when the Company believes actual experience may differ from its estimates. The allowances included in the table below reflect these adjustments.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

The following table provides a summary of activity with respect to the Company's allowances and accruals during 2006, 2005 and 2004 (amounts in thousands):

|   | Returns | Chargebacks | Rebates  |
|---|---------|-------------|----------|
| <b>Balance at December 31, 2003</b>         | 1,102   | 1,772       | 1,828    |
| Allowances for sales during 2004            | 121     | 5,978       | 2,663    |
| Actual credits issued for prior years sales | (617 )  | (1,852 )    | (1,913 ) |
| Actual credits issued for sales during 2004 | (3 )    | (2,795 )    | (954 )   |
| <b>Balance at December 31, 2004</b>         | 603     | 3,103       | 1,624    |
| Allowances for sales during 2005            | (240 )  | 1,776       | 2,334    |
| Actual credits issued for prior years sales | (146 )  | (2,895 )    | (1,317 ) |
| Actual credits issued for sales during 2005 |         | (1,478 )    | (1,187 ) |
| <b>Balance at December 31, 2005</b>         | 217     | 506         | 1,454    |
| Allowances for sales during 2006            | 404     | 4,240       | 2,247    |
| Actual credits issued for prior years sales | (212 )  | (737 )      | (1,318 ) |
| Actual credits issued for sales during 2006 | (8 )    | (3,681 )    | (1,549 ) |
| <b>Balance at December 31, 2006</b>         | \$ 401  | \$ 328      | \$ 834   |

*International Distributors.* Under our agreements with our primary international distributors, we sell our product to these distributors at a fixed transfer price. The established transfer price is typically determined twice in each year, prior to the first and last shipment of Angiomax to the distributor each year. If not agreed upon by the parties prior to such shipments, the price is determined by taking an average of the transfer price for the preceding three shipments. The minimum selling price used in determining the transfer price of 50% of the average net unit selling price, subject to mutually agreed adjustments. The transfer price is denominated in Euros, which are then converted to U.S. Dollars, payable by our distributors, at the exchange rate between the two currencies, as quoted by the European Central Bank, just prior to shipment.

Revenue from the sale of distribution rights includes the amortization of milestone payments. These milestone payments are recorded as deferred revenue until contractual performance obligations have been satisfied, and they are typically recognized ratably over the term of these agreements. When the period of deferral cannot be specifically identified from the contract, the Company must estimate the period based upon other critical factors contained within the contract. The Company reviews these estimates at least annually, which could result in a change in the deferral period.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

***Reimbursement Revenue***

In collaboration with a third party, the Company has paid fees for services rendered by a research organization and other out-of-pocket costs for which it was reimbursed at cost, without mark-up or profits. The Company accounts for these arrangements using FASB EITF 01-14 Income Statement Characterization of Reimbursements Received for Out-of-Pocket Expenses Incurred ( EITF 01-14 ) and FASB EITF 99-19 Reporting Revenue Gross as a principal versus Net as an Agent ( EITF 99-19 ). The reimbursements received have been reported as part of net revenue in our consolidated statements of operations. The fees for the services rendered and the out-of-pocket costs have been included in research and development expenses.

***Cost of Revenue***

Cost of revenue consists of expenses in connection with the manufacture of the Angiomax sold, royalty expenses under the Company's agreement with Biogen Idec, Inc., and the logistics costs of selling Angiomax, such as distribution, storage and handling.

***Advertising Costs***

The Company expenses advertising costs as incurred. Advertising costs were approximately \$2.7 million, \$1.3 million and \$0.7 million, for the years ended December 31, 2006, 2005, and 2004, respectively.

***Inventory***

Inventory is recorded upon the transfer of title from the Company's vendors. Inventory is stated at the lower of cost or market value and valued using first-in, first-out methodology. Angiomax bulk substance is classified as raw materials and its costs are determined using acquisition costs from the Company's contract manufacturer. Work-in-progress costs of filling, finishing and packaging are recorded against specific product batches. The Company obtains all of its Angiomax bulk substance from the manufacturing division of UCB Bioproducts S.A., which was recently acquired by Lonza Ltd. and is now known as Lonza Braine, S.A. Under the terms of the Company's agreement with Lonza Braine, the Company provides forecasts of Angiomax bulk substance needs eighteen months in advance of the year of delivery. The Company also has a separate agreement with Ben Venue Laboratories, Inc. for the fill-finish of Angiomax drug product.

The major classes of inventory were as follows:

| <b>Inventory</b> | <b>2006</b>           | <b>2005</b>      |
|------------------|-----------------------|------------------|
|                  | <b>(in thousands)</b> |                  |
| Raw materials    | \$ 25,456             | \$ 21,048        |
| Work-in-progress | 12,506                | 23,630           |
| Finished goods   | 3,666                 | 3,307            |
| <b>Total</b>     | <b>\$ 41,628</b>      | <b>\$ 47,985</b> |

The Company reviews inventory for slow moving or obsolete amounts based on expected revenues. If annual revenues are less than expected, the Company may be required to make allowances for excess or obsolete inventory in the future.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

*Fixed Assets*

Fixed assets are stated at cost. Depreciation is provided using the straight-line method based on estimated useful lives or, in the case of leasehold improvements, over the lesser of the useful lives or the lease terms.

*Research and Development*

Research and development costs are expensed as incurred.

*Stock-Based Compensation*

Prior to January 1, 2006, the Company elected to account for stock-based compensation using the intrinsic value method prescribed in Accounting Principles Board Opinion No. 25, Accounting for Stock Issued to Employees as permitted by Statement of Financial Accounting Standards ( SFAS ) No. 123, Accounting for Stock-Based Compensation ( SFAS No.123 ).

Effective January 1, 2006, the Company adopted the fair value recognition provisions of Financial Accounting Standards Board Statement No. 123 (revised 2004), Share-Based Payment ( SFAS 123(R) ), and the Company has elected the modified prospective transition method and, therefore, adjustments to prior periods are not required as a result of adopting SFAS 123(R). Under this method, the provisions of SFAS 123(R) apply to all awards granted after January 1, 2006, and to any unrecognized expense of awards unvested at the date of adoption based on the grant date fair value. The Company is recognizing expense using the accelerated expense attribution method specified in FASB Interpretation No. 28, Accounting for Stock Appreciation Rights and Other Variable Stock Option or Award Plans (FIN 28).

The following table illustrates the effect on net (loss)/ income and (loss)/earnings per share if the Company had applied the fair value recognition provisions of SFAS No. 123 to stock-based employee compensation:

|  | <b>Years Ended December 31,</b>                 |             |
|--|---|-------------|
|  | <b>2005</b>                                     | <b>2004</b> |
|  | <b>(in thousands, except per share amounts)</b> |             |
| Net (loss)/income As reported  | \$ (7,753 )                                     | \$ 16,999   |
| Deduct: Total stock-based compensation expense determined under fair value based method for all stock option awards and discounts under the employee stock purchase plan, net of tax | (42,670 )                                       | (15,002 )   |
| Add: Amortization of deferred stock compensation reported pursuant to APB 25, net of tax   |   | 725         |
| Net (loss)/income Pro forma  | \$ (50,423 )                                    | \$ 2,722    |
| Net (loss)/earnings per common share, basic As reported  | \$ (0.16 )                                      | \$ 0.36     |
| Net (loss)/earnings per common share, basic Pro forma  | \$ (1.02 )                                      | \$ 0.06     |
| Net (loss)/earnings per common share, diluted As reported  | \$ (0.16 )                                      | \$ 0.34     |
| Net (loss)/earnings per common share, diluted Pro forma  | \$ (1.02 )                                      | \$ 0.05     |



**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

Expected volatilities are based on historic volatility of the Company's common stock as well as peer companies in the life science industry over a range of periods from 12 to 60 months and other factors. The Company uses historical data to estimate forfeiture rate. The expected term of options represents the period of time that options granted are expected to be outstanding. The Company has made a determination of expected term by analyzing employees' historical exercise experience and has made estimates of future exercises of unexercised options based on the midpoint between the vesting date and end of the contractual term. The risk-free interest rate is based on the U.S. Treasury yield in effect at the time of grant corresponding with the expected life of the options.

For purposes of applying SFAS 123(R) during the year ended December 31, 2006, the Company estimates the fair value of each option on the date of grant using the Black-Scholes closed-form option-pricing model applying the weighted average assumptions in the following table. The Company allocated this fair value to compensation expense using the accelerated expense attribution method specified in FIN 28.

For purposes of performing the valuation, employees were separated into two groups according to patterns of historical exercise behavior; the weighted average assumptions below result from the two groups of employees exhibiting different behavior.

|                                 | Years Ended December 31, |   |      |   |      |   |
|---------------------------------|--------------------------|---|------|---|------|---|
|                                 | 2006                     |   | 2005 |   | 2004 |   |
| Expected dividend yield         | 0                        | % | 0    | % | 0    | % |
| Expected stock price volatility | 45%, 47%                 |   | 55   | % | 79   | % |
| Risk-free interest rate         | 4.77%, 4.78%             |   | 4.05 | % | 2.84 | % |
| Expected option term (years)    | 3.38, 3.60               |   | 2.94 |   | 2.84 |   |

On December 23, 2005, upon the recommendation of its Compensation Committee, the Board of Directors of the Company approved full acceleration of the vesting of each otherwise unvested stock option:

- with an exercise price per share equal to or greater than \$20.50,
- granted under the 1998 Stock Incentive Plan, 2000 Outside Director Stock Option Plan, 2001 Non-Officer, Non-Director Employee Stock Incentive Plan, 2004 Stock Incentive Plan, and
- held by employees, officers and non-employee directors of the Company.

The acceleration of vesting on December 23, 2005 affected options to purchase approximately 3,894,350 shares of the Company's common stock, par value \$0.001 per share. These options would have otherwise vested between December 23, 2005 and October 1, 2009. The Company accelerated the vesting of these options to eliminate future compensation expense that otherwise would have been recognized under SFAS 123(R). The Company estimated that the aggregate future expense that it eliminated as a result of the acceleration of the vesting of these options was approximately \$22.2 million, which would otherwise have been recognized over the respective vesting periods of the individual options. The above pro forma information for the year ended December 31, 2005 includes the effect of accelerating these options.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**2. Significant Accounting Policies (Continued)**

*Translation of Foreign Currencies*

The functional currencies of the Company's foreign subsidiaries are the local currencies: British pound sterling and New Zealand dollar. In accordance with SFAS No. 52 Foreign Currency Translation, the Company's assets and liabilities are translated using the current exchange rate as of the balance sheet date. Stockholders' equity is translated using historical rates at the balance sheet date. Expenses and items of income are translated using a weighted average exchange rate over the period ended on the balance sheet date. Adjustments resulting from the translation of the financial statements of the Company's foreign subsidiaries into U.S. dollars are excluded from the determination of net earnings/(loss) and are accumulated in a separate component of stockholders' equity. Foreign exchange transaction gains and losses are included in the results of operations and are not material to the Company's consolidated financial statements.

*Income Taxes*

Deferred tax assets and liabilities are determined based on differences between financial reporting and income tax bases of assets and liabilities, as well as net operating loss carryforwards, and are measured using the enacted tax rates and laws in effect when the differences are expected to reverse. Deferred tax assets are reduced by a valuation allowance to reflect the uncertainty associated with ultimate realization.

*Comprehensive Income/(Loss)*

The Company reports comprehensive income/(loss) and its components in accordance with the provisions of SFAS No. 130, Reporting Comprehensive Income. Comprehensive income/(loss) includes net income/(loss), all changes in equity for cumulative translations adjustments resulting from the consolidation of foreign subsidiaries' financial statements and unrealized gain/(loss) on available for sale securities.

|   | <b>Years Ended December 31,</b> |             |             |
|---|---------------------------------|-------------|-------------|
|   | <b>2006</b>                     | <b>2005</b> | <b>2004</b> |
|   | <b>(in thousands)</b>           |             |             |
| Net income/(loss) As reported                           | \$ 63,726                       | \$ (7,753 ) | \$ 16,999   |
| Unrealized gain/(loss) on available for sale securities | 238                             | 90          | (547 )      |
| Foreign currency translation adjustment                 | 23                              | (22 )       | 12          |
| Comprehensive income/(loss)                             | \$ 63,987                       | \$ (7,685 ) | \$ 16,464   |

*Segments*

The Company manages its business and operations as one segment and is focused on the acquisition, development and commercialization of late-stage development drugs and drugs approved for marketing. The Company has licensed rights to Angiomax®, Cleviprex and cangrelor. Revenues reported to date are derived primarily from the sales of Angiomax in the United States.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**3. Recent Accounting Pronouncements**

In June 2006, the Financial Accounting Standards Board (FASB) issued FASB Interpretation No. (FIN) 48 Accounting for Uncertainty in Income Taxes an interpretation of FASB Statement No. 109 ( FIN 48 ). FIN 48 prescribes a threshold for the financial statement recognition and measurement of a tax position taken or expected to be taken in a tax return. Only tax positions meeting the more-likely-than-not recognition threshold at the effective date may be recognized or continue to be recognized upon adoption of this interpretation. FIN 48 also provides guidance on accounting for derecognition, interest and penalties, and classification and disclosure of matters related to uncertainty in income taxes. FIN 48 will be effective for the Company beginning January 1, 2007. The Company currently expects that adoption of this interpretation will not have a material effect on the Company's consolidated financial position and results of operations.

In September 2006, the FASB issued Statement of Financial Accounting Standards (SFAS) No. 157 Fair Value Measurements, ( SFAS No. 157 ) which defines fair value, establishes a framework for consistently measuring fair value under GAAP, and expands disclosures about fair value measurements. SFAS No. 157 is effective for the Company beginning January 1, 2008, and the provisions of SFAS No. 157 will be applied prospectively as of that date. The Company is currently evaluating the effect that adoption of this statement will have on the Company's consolidated financial position and results of operations when it becomes effective in 2008.

**4. The Company's Plans and Financing**

Except for the years ended December 31, 2006 and December 31, 2004, the Company has incurred net losses on an annual basis since inception. To date, the Company has primarily funded its operations through the issuance of debt and equity, and, in 2006 and 2004, from cash flow from operations. The Company expects to continue to expend substantial amounts for continued product research, development and commercialization activities for the foreseeable future, and the Company plans to fund these expenditures from revenue or through debt or equity financing, if possible. Should revenue or additional debt or equity financing be unavailable to the Company, it will restrict certain of its planned activities and operations, as necessary, to sustain operations and conserve cash resources.

In January 2007, the Company announced its intention to commence an offering of 6,000,000 shares of its Common Stock pursuant to an effective shelf registration in an underwritten public offering. However, the Company subsequently determined not to proceed with the public offering of common stock at that time.

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**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**5. Fixed Assets**

Fixed assets consist of the following:

|                                   | <b>Estimated<br/>Life (Years)<br/>(in thousands)</b> | <b>December 31,<br/>2006</b> | <b>2005</b> |
|-----------------------------------|--|------------------------------|-------------|
| Furniture, fixtures and equipment | 3  | \$ 2,386                     | \$ 2,109    |
| Computer software                 | 3  | 1,337                        | 1,398       |
| Computer hardware                 | 3  | 1,651                        | 1,426       |
| Leasehold improvements            | 5-10   | 1,269                        | 1,265       |
|                                   |  | 6,643                        | 6,198       |
| Less: Accumulated depreciation    |  | (3,572 )                     | (2,208 )    |
|                                   |  | \$ 3,071                     | \$ 3,990    |

Depreciation expense was approximately \$1.5 million, \$1.0 million, and \$0.6 million for the years ended December 31, 2006, 2005 and 2004, respectively.

**6. Accrued Expenses**

Accrued expenses consisted of the following at December 31:

|   | <b>2006<br/>(in thousands)</b> | <b>2005</b> |
|---|--------------------------------|-------------|
| Research and development services         | \$ 8,848                       | \$ 10,688   |
| Royalties                                 | 11,722                         | 6,356       |
| Compensation related                      | 8,073                          | 5,058       |
| Product returns and rebates               | 1,234                          | 1,671       |
| Manufacturing, logistics and related fees | 1,723                          | 1,058       |
| Legal, accounting and other               | 3,076                          | 2,503       |
| Sales and marketing                       | 2,242                          | 1,343       |
|   | \$ 36,918                      | \$ 28,677   |

**7. Common Stock Purchase Warrants**

In March 2000, the Company issued \$13.4 million of 8% convertible notes ( March Notes ) and warrants (the March Warrants ) to purchase 2,255,687 shares of Common Stock to then existing investors, raising proceeds of \$13.4 million. The March Notes were ultimately converted into shares of Common Stock of the Company. Each March Warrant provided the holder with the right to purchase one share of Common Stock at a price of \$5.92 per share at any time prior to March 2, 2005. At December 31, 2004 there were March Warrants outstanding to purchase 661,561 shares of Common Stock. All of these warrants were exercised on or before March 2, 2005.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders' Equity**

*Preferred Stock*

The Company has 5,000,000 shares of preferred stock (the Preferred Stock) authorized, none of which are issued.

*Common Stock*

Common stockholders are entitled to one vote per share and dividends when declared by the Company's Board of Directors, subject to the preferential rights of any outstanding shares of Preferred Stock.

Employees, directors and consultants of the Company purchased 1,503,557; 578,763 and 1,097,041 shares of Common Stock during the years ended December 31, 2006, 2005 and 2004, respectively, pursuant to option exercises and the Company's employee stock purchase plan. The aggregate net proceeds to the Company resulting from these purchases were approximately \$24.0 million, \$6.8 million, and \$13.7 million during the years ended December 31, 2006, 2005 and 2004, respectively, and are included within the financing activities section of the consolidated statements of cash flows.

Pursuant to provisions of the March Warrants, 500,179 shares of Common Stock were issued on a cashless exercise basis to holders of the underlying warrants during the year ended December 31, 2005, resulting in no proceeds to the Company.

*Stock Plans*

The Company has adopted the following stock incentive plans:

- the 2004 Stock Incentive Plan (the 2004 Plan),
- the 2001 Non-Officer, Non-Director Stock Incentive Plan (the 2001 Plan),
- the 2000 Outside Director Stock Option Plan (the 2000 Director Plan), and
- the 1998 Stock Incentive Plan (the 1998 Plan).

Each of these plans provides for the grant of stock options and other stock-based awards to employees, officers, directors, consultants and advisors of the Company and its subsidiaries. Stock option grants have an exercise price equal to the fair market value of the Company's Common Stock on the date of grant and generally have a 10-year term. The fair value of stock option grants is recognized, net of an estimated forfeiture rate of approximately 17%, using an accelerated method over the vesting period of the options, which is generally four years.

*2004 Plan*

In April 2004, the Board of Directors adopted, subject to stockholder approval, the 2004 Plan, which provides for the grant of stock options, restricted stock awards, stock appreciation rights and other stock-based awards to the Company's employees, officers, directors, consultants and advisors, including any individuals who have accepted an offer of employment. The Company's stockholders approved the 2004

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders Equity (Continued)**

Plan in May 2004. The number of shares the Company may issue is restated to reflect an amendment approved by the Board of Directors on April 11, 2006 and by stockholders at the 2006 annual meeting.

The Company may issue up to 8,800,000 shares of Common Stock, subject to adjustment in the event of stock splits and other similar events, pursuant to awards granted under the 2004 Plan. The Board of Directors has delegated its authority under the 2004 Plan to the Compensation Committee, consisting of independent directors, which administers the 2004 Plan, including granting options and other awards under the 2004 Plan. In addition, pursuant to the terms of the 2004 Plan, the Board of Directors has delegated to the Company's executive officers limited authority to grant stock options to employees without further action by the Board of Directors or the Compensation Committee. Options granted under the 2004 Plan generally vest in increments over 4 years and have a 10-year term.

The Board of Directors has adopted a program under the 2004 Plan providing for automatic options grants to the Company's non-employee directors. Each non-employee director is granted non-statutory stock options under the 2004 Plan to purchase:

- 20,000 shares of Common Stock on the date of his or her initial election to the Board of Directors (the Initial Options ); and
- 15,000 shares of the Common Stock on the date of each annual meeting of the Company's stockholders (the Annual Options ), except if such non-employee director was initially elected to the Board of Directors at such annual meeting. The lead director will be granted an additional option to purchase 5,000 shares of the Common Stock on the date of each annual meeting of the Company's stockholders.

These options have an exercise price equal to the closing price of the Common Stock on the NASDAQ Global Select Market on the date of grant and have a 10-year term. The Initial Options vest in 36 equal monthly installments beginning on the date one month after the grant date. The Annual Options vest in 12 equal monthly installments beginning on the date one month after the date of grant. All vested options are exercisable at any time prior to the first anniversary of the date the director ceases to be a director.

*2001 Plan*

In May 2001, the Board of Directors approved the 2001 Plan, which provides for the grant of non-statutory stock options to employees, consultants and advisors of the Company and its subsidiaries, including individuals who have accepted an offer of employment, other than those employees who are officers or directors of the Company. The 2001 Plan provides for the issuance of up to 1,250,000 shares of Common Stock. The Board of Directors has delegated its authority under the 2001 Plan to the Compensation Committee, which administers the 2001 Plan, including granting options under the 2001 Plan. In addition, pursuant to the terms of the 2001 Plan, the Board of Directors has delegated to the Company's chief executive officer limited authority to grant stock options to employees without further action by the Board of Directors or the Compensation Committee. The Company ceased making grants under the 2001 Plan following adoption of an amendment to the 2004 Plan at its annual stockholders' meeting on May 25, 2006. Unexercised options under the 2001 Plan remain outstanding.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders Equity (Continued)**

*2000 Director Plan*

Prior to the adoption of the 2004 Plan, the Company granted non-statutory stock options to the Company's non-employee directors pursuant to the 2000 Director Plan. The Company ceased making grants under the 2000

Director Plan following adoption of the 2004 Plan. Unexercised options under the 2000 Director Plan remain outstanding.

*1998 Plan*

In April 1998, the Company adopted the 1998 Plan, which provides for the grant of stock options, restricted stock and other stock-based awards to employees, officers, directors, consultants, and advisors of the Company and its subsidiaries, including any individuals who have accepted an offer of employment. The Board of Directors has authority to determine the term of each option, the option price, the number of shares for which each option is granted and the rate at which each option becomes exercisable. As a result of subsequent amendments, the 1998 Plan currently provides that 6,118,259 shares of Common Stock may be issued pursuant to awards under the 1998 Plan. During 1999, the Board of Directors amended all then-outstanding options to allow holders to exercise the options prior to vesting, provided that the shares of Common Stock issued upon exercise of the option would be subject to transfer restrictions and vesting provisions that allowed the Company to repurchase unvested shares at the exercise price. There were no outstanding unvested shares of Common Stock at December 31, 2005 and 2006. Pursuant to the terms of the 1998 Plan, the Board of Directors has delegated its authority under the 1998 Plan to the Compensation Committee. Accordingly, the Compensation Committee administers the 1998 Plan, including granting options and other awards under the 1998 Plan. In addition, pursuant to the terms of the 1998 Plan, the Board of Directors has delegated to the Company's chief executive officer limited authority to grant stock options to employees without further action by the Board of Directors or the Compensation Committee. Options granted under the 1998 Plan generally vest in increments over four years and have a ten-year term. The Company ceased making grants under the 1998 Plan following adoption of an amendment to the 2004 Plan at its annual stockholders' meeting on May 25, 2006. Unexercised options under the 1998 Plan remain outstanding.

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**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders Equity (Continued)**

*Stock Option Activity*

The following table presents a summary of option activity and data under the Company's stock incentive plans as of December 31, 2006:

|   | Number of Shares | Weighted-Average Exercise Price Per Share | Weighted-Average Remaining Contractual Term | Aggregate Intrinsic Value |
|---|------------------|---|---|---------------------------|
| Outstanding, December 31, 2003                  | 5,315,993        | \$ 15.98                                  |   |                           |
| Granted   | 2,262,500        | 27.64                                     |   |                           |
| Exercised                                       | (1,060,174 )     | 12.06                                     |   |                           |
| Forfeited and expired                           | (409,285 )       | 21.61                                     |   |                           |
| Outstanding, December 31, 2004                  | 6,109,034        | 20.60                                     |   |                           |
| Granted   | 2,884,750        | 20.61                                     |   |                           |
| Exercised                                       | (526,557 )       | 11.05                                     |   |                           |
| Forfeited and expired                           | (788,091 )       | 24.57                                     |   |                           |
| Outstanding, December 31, 2005                  | 7,679,136        | 20.85                                     |   |                           |
| Granted   | 1,496,789        | 20.60                                     |   |                           |
| Exercised                                       | (1,415,605 )     | 16.15                                     |   |                           |
| Forfeited and expired                           | (1,006,913 )     | 24.66                                     |   |                           |
| Outstanding, December 31, 2006                  | 6,753,407        | \$ 21.21                                  | 7.80  | \$ 71,173,236             |
| Exercisable, December 31, 2006                  | 4,747,702        | \$ 21.75                                  | 7.22  | \$ 47,540,322             |
| Available for future grant at December 31, 2006 | 4,139,305        |   |   |                           |

Aggregate intrinsic value is the sum of the amounts by which the quoted market price of the Company's common stock exceeded the exercise price of the options at December 31, 2006, for those options for which the quoted market price was in excess of the exercise price. The weighted-average grant date fair value of options granted during the years ended December 31, 2006, 2005 and 2004 was \$7.95, \$8.06 and \$13.79, respectively. The total intrinsic value of options exercised during the years ended December 31, 2006, 2005 and 2004 was \$10.3 million, \$6.9 million and \$17.1 million, respectively.

In accordance with SFAS 123(R), the Company recorded approximately \$8.5 million of stock-based compensation expense for the year ended December 31, 2006. As of December 31, 2006, there was approximately \$10.6 million of total unrecognized compensation costs related to non-vested share-based employee compensation arrangements granted under the Company's equity compensation plans. This cost is expected to be recognized over a weighted average period of 1.52 years.



**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders Equity (Continued)**

The following table summarizes information regarding options outstanding as of December 31, 2006:

| Range of Exercise Prices Per Share | Options Outstanding            |   | Weighted Average Remaining Contractual Life (Years) | Weighted Average Exercise Price Per Share | Options Vested Number Outstanding at 12/31/06 | Weighted Average Exercise Price Per Share |
|------------------------------------|--------------------------------|---|---|---|---|---|
|                                    | Number Outstanding at 12/31/06 | Weighted Average Remaining Contractual Life (Years) |   |   |   |   |
| \$1.23-\$10.04                     | 405,252                        | 3.78  | \$ 5.70   | 405,252                                   | \$ 5.70                                       |   |
| \$10.11-\$13.80                    | 149,357                        | 5.12  | 11.41   | 149,357                                   | 11.41   |   |
| \$15.32-\$17.38                    | 446,878                        | 6.37  | 16.31   | 379,607                                   | 16.16   |   |
| \$17.45-\$19.09                    | 1,827,019                      | 8.90  | 18.43   | 702,223                                   | 18.31   |   |
| \$19.11-\$22.51                    | 1,145,253                      | 8.68  | 21.16   | 589,768                                   | 21.60   |   |
| \$22.56-\$24.60                    | 791,157                        | 7.92  | 23.32   | 652,504                                   | 23.43   |   |
| \$24.92-\$27.81                    | 919,948                        | 7.66  | 26.39   | 842,448                                   | 26.33   |   |
| \$27.87-\$30.27                    | 780,043                        | 7.62  | 28.22   | 754,043                                   | 28.17   |   |
| \$30.69-\$34.95                    | 288,500                        | 7.27  | 32.25   | 272,500                                   | 32.32   |   |
|                                    | 6,753,407                      | 7.80  | 21.21   | 4,747,702                                 | \$ 21.75                                      |   |

The following table presents a summary of the Company's non-vested shares of restricted stock granted as of December 31, 2006:

|                               | Number of Shares |        |  | Weighted Average Grant-Date Fair Value |          |  |
|-------------------------------|------------------|--------|--|--|----------|--|
| Non-vested, December 31, 2005 |                  |        |  |  |          |  |
| Awarded                       |                  | 25,000 |  |  | \$ 20.11 |  |
| Vested                        |                  |        |  |  |          |  |
| Forfeited                     |                  |        |  |  |          |  |
| Non-vested, December 31, 2006 |                  | 25,000 |  |  | \$ 20.11 |  |

The Company granted a restricted stock award under the 2004 stock incentive plan during the first quarter of 2006. The restricted stock grant vests in equal increments of 25% per year on an annual basis commencing twelve months after grant date. Expense of approximately \$0.2 million was recognized in the year ended December 31, 2006. The remaining expense of approximately \$0.3 million will be recognized over a period of 3.17 years.

*2000 ESPP*

In May 2000, the Board of Directors and the Company's stockholders approved the 2000 Employee Stock Purchase Plan (the 2000 ESPP), which provides for the issuance of up to 505,500 shares of Common Stock. The number of shares the Company may issue under the 2000 ESPP reflects an amendment approved by the Board of Directors on April 11, 2006 and by stockholders at the 2006 annual meeting. The 2000 ESPP permits eligible employees to purchase shares of Common Stock at the lower of 85% of the fair market value of the Common Stock at the beginning or at the end of each offering period. Employees who own 5% or more of the Common Stock are not eligible to participate in the 2000 ESPP. Participation is voluntary.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**8. Stockholders Equity (Continued)**

As of December 31, 2006, the Company had issued 240,442 shares over the life of the 2000 ESPP. The Company issued 62,952 shares and 52,206 shares under the 2000 ESPP during the years ended 2006 and 2005, respectively, and currently has 265,058 shares in reserve for future issuance under the 2000 ESPP. The Company recorded approximately \$0.4 million in compensation expense related to the 2000 ESPP in the year ended December 31, 2006.

The fair value of each option element of the 2000 ESPP is estimated on the date of grant using the Black-Scholes closed-form option valuation model that applies the assumptions noted in the following table. Expected volatilities are based on historical volatility of the Company's Common Stock. Expected term represents the six-month offering period for the 2000 ESPP. The risk-free interest rate is based on the U.S. Treasury yield curve in effect at the time of grant.

|                                 | Years Ended December 31, |   |      |   |      |   |
|---------------------------------|--------------------------|---|------|---|------|---|
|                                 | 2006                     |   | 2005 |   | 2004 |   |
| Expected dividend yield         | 0                        | % | 0    | % | 0    | % |
| Expected stock price volatility | 31%, 41                  | % | 31   | % | 79   | % |
| Risk-free interest rate         | 4.63, 5.06               | % | 3.57 | % | 2.84 | % |
| Expected option term (years)    | 0.5                      |   | 0.5  |   | 0.5  |   |

During 2005 and 2004, under the provisions of APB 25, the Company did not record any expense for the 2000 ESPP Plan.

*Common Stock Reserved for Future Issuance*

At December 31, 2006, there were 11,157,770 shares of Common Stock reserved for future issuance under the 2000 ESPP and grants made under the 1998 Plan, the 2000 Director Plan, the 2001 Plan, and the 2004 Plan.

**9. Net Earnings/(Loss) per Share**

The following table sets forth the computation of basic and diluted net earnings/(loss) per share for the years ended December 31, 2006, 2005 and 2004.

|   | Years Ended December 31,                 |             |           |
|---|--|-------------|-----------|
|   | 2006                                     | 2005        | 2004      |
|   | (in thousands, except per share amounts) |             |           |
| <b><u>Basic and diluted</u></b>                     |  |             |           |
| Net income/(loss) As reported                       | \$ 63,726                                | \$ (7,753 ) | \$ 16,999 |
| Weighted average common shares outstanding, basic   | 50,321                                   | 49,443      | 47,855    |
| Less: unvested restricted common shares outstanding | 21                                       |             |           |
| Weighted average common shares outstanding, basic   | 50,300                                   | 49,443      | 47,855    |
| Net effect of dilutive stock options and warrants   | 734                                      |             | 1,917     |
| Weighted average common shares outstanding, diluted | 51,034                                   | 49,443      | 49,772    |
| Net earnings/(loss) per common share, basic         | \$ 1.27                                  | \$ (0.16 )  | \$ 0.36   |
| Net earnings/(loss) per common share, diluted       | \$ 1.25                                  | \$ (0.16 )  | \$ 0.34   |

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**9. Net Earnings/(Loss) per Share (Continued)**

Basic net earnings/(loss) per share is computed using the weighted average number of shares of Common Stock outstanding during the period reduced, where applicable, for outstanding, yet unvested shares. As of December 31, 2006, there were options to purchase 6,753,407 shares of Common Stock outstanding and at December 31, 2004 there were options to purchase 6,109,034 shares of Common Stock outstanding and warrants to purchase 662,000 shares of Common Stock outstanding. Except for 207,500 options in 2006 and 565,833 options in 2004, which were considered anti-dilutive, these options and warrants were included in the computation of diluted net earnings per share for the years ended December 31, 2006 and 2004, respectively. The number of dilutive common stock equivalents for the years 2006 and 2004 was calculated using the treasury stock method. As of December 31, 2005, there were options to purchase 7,679,136 shares of Common Stock outstanding. The Company has not included these options in the computation of diluted net loss per share for the year ended December 31, 2005, as their effect would have been anti-dilutive.

**10. Income Taxes**

The benefit from/(provision for) income taxes in 2006, 2005 and 2004 consists of current and deferred federal, state and foreign taxes paid based on net income and state taxes based on net worth as follows:

|  | 2006<br>(in thousands) | 2005             | 2004             |
|--|------------------------|------------------|------------------|
| <b>Current:</b>  |                        |                  |                  |
| Federal  | \$ (348 )              | \$ (100 )        | \$ (411 )        |
| State  | (143 )                 | (100 )           | (240 )           |
| Foreign  |                        |                  | (39 )            |
|  | (491 )                 | (100 )           | (690 )           |
| <b>Deferred:</b>                                       |                        |                  |                  |
| Federal  | 43,300                 |                  |                  |
| State  | 3,259                  |                  |                  |
| Foreign  |                        |                  |                  |
|  | 46,559                 |                  |                  |
| <b>Total benefit from/(provision for) income taxes</b> | <b>\$ 46,068</b>       | <b>\$ (100 )</b> | <b>\$ (690 )</b> |

The components of Income Before Income taxes consisted of:

|               | 2006<br>(in thousands) | 2005               | 2004             |
|---------------|------------------------|--------------------|------------------|
| Domestic      | \$ 17,689              | \$ (7,682 )        | \$ 17,643        |
| International | (31 )                  | 29                 | 46               |
| <b>Total</b>  | <b>\$ 17,658</b>       | <b>\$ (7,653 )</b> | <b>\$ 17,689</b> |

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**10. Income Taxes (Continued)**

The difference between tax expense and the amount computed by applying the statutory federal income tax rate (34%) to income before income taxes is as follows:

|  | <b>Year Ended December 31,</b> |             |             |
|--|--------------------------------|-------------|-------------|
|  | <b>2006</b>                    | <b>2005</b> | <b>2004</b> |
|  | <b>(in thousands)</b>          |             |             |
| Statutory rate applied to pre-tax income/(loss)          | \$ 6,004                       | \$ (2,643 ) | \$ 6,007    |
| Add (deduct):  |                                |             |             |
| State income taxes, net of federal benefit               | (2,057 )                       | 65          | 321         |
| Foreign  | 4                              | (10 )       | 23          |
| Tax credits  | (2,326 )                       | (2,389 )    | (1,949 )    |
| Other  | 100                            | 342         | 283         |
| Increase/(decrease) to federal valuation allowance (net) | (47,793)                       | 4,735       | (3,995 )    |
| Income taxes   | \$ (46,068)                    | \$ 100      | \$ 690      |

The significant components of the Company's deferred tax assets are as follows:

|                                  | <b>December 31,</b>   | <b>2005</b> |
|----------------------------------|-----------------------|-------------|
|                                  | <b>(in thousands)</b> |             |
| Deferred tax assets:             |                       |             |
| Net operating loss carryforwards | \$ 80,541             | \$ 86,765   |
| Research and development credit  | 16,185                | 13,879      |
| Intangible assets                | 545                   | 635         |
| Stock based compensation         | 3,008                 |             |
| Other                            | 3,645                 | 3,741       |
| Total deferred tax assets        | 103,924               | 105,020     |
| Valuation allowance              | (54,724 )             | (105,020 )  |
| Net deferred tax assets          | \$ 49,200             | \$          |

The net change in deferred tax assets is primarily due to current year net operating income, offset partially by increases in income tax credits and stock compensation benefits. Additionally, the Company recorded a portion of its deferred tax assets by reducing the valuation allowances that had been recorded in prior years since the realization of these future benefits is now considered more likely than not. The amount of the deferred tax asset considered realizable is subject to change based on estimates of future taxable income during the carryforward period. If the Company maintains profitability, these deferred tax assets are available to offset future income taxes. The Company had increased its valuation allowance against these assets by \$7,171 in 2005 and had restated certain deferred tax assets relating to net operating loss carryforwards as of December 31, 2004 to reflect the expiration of certain state net operating loss carryforwards.

The Company considers the level of past and future taxable income as well as the utilization of carryforwards and other factors when considering the recognition of deferred tax assets. Based upon these considerations, the Company reduced its valuation allowance and recognized a \$49.2 million deferred tax asset in the fourth quarter of 2006, which resulted in a \$46.6 million benefit to deferred income taxes and a \$2.6 million credit to additional paid-in capital, representing the current year's portion of the deferred tax

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**10. Income Taxes (Continued)**

asset related to stock option tax benefits. This benefit was primarily related to the portion of deferred tax assets that management believes it is more likely than not will be realized in future periods.

Factors that could significantly impact the Company's valuation allowance include the regulatory approval of products currently under development, extension of the patent rights relating to Angiomax or failure to achieve future anticipated revenues. Should we further reduce or increase the valuation allowance on deferred tax assets, a current year tax benefit or expense would be recognized and future periods would then include income taxes at a higher or lower rate than the effective rate in the period that the adjustment is made.

The valuation allowance was decreased in total by \$50.3 million during 2006 as a result of the reduction in valuation allowance partially offset by changes in deferred tax assets.

In 1998 and 2002 the Company experienced a change in ownership as defined in Section 382 of the Internal Revenue Code. Section 382 can potentially limit a company's ability to use net operating losses, tax credits and other tax attributes in periods subsequent to a change in ownership. However, based on the market value of the Company at such dates, the Company believes that these ownership changes will not significantly impact its ability to use net operating losses or tax credits in the future to offset taxable income. At December 31, 2006, the Company has federal net operating loss carryforwards available to reduce taxable income, and federal research and development tax credit carryforwards available to reduce future tax liabilities, which expire approximately as follows:

| <b>Year of Expiration</b> | <b>Federal Net<br/>Operating Loss<br/>Carryforwards<br/>(in thousands)</b> | <b>Federal Research<br/>and Development<br/>Tax Credit<br/>Carryforwards</b> |
|---------------------------|--|--|
| 2011                      | \$   | \$ 22  |
| 2012                      |  | 527  |
| 2018                      | 21,250   | 425  |
| 2019                      | 33,803   | 1,002  |
| 2020                      | 45,270   | 1,176  |
| 2021                      | 51,100   | 477  |
| 2022                      | 41,403   | 1,876  |
| 2023                      | 19,693   | 2,282  |
| 2024                      | 11   | 1,949  |
| 2025                      | 12,506   | 3,620  |
| 2026                      |  | 2,326  |
|                           | \$ 225,036   | \$ 15,682  |

At December 31, 2006 a total of \$10.8 million of the deferred tax asset valuation allowance related to net operating loss carryforwards is associated with the exercise of non-qualified stock options. Such benefits, when realized, will be credited to additional paid-in capital.

For state tax purposes, net operating loss carryforwards of approximately \$64.4 million expire in the years 2007 through 2025. State research and development tax credit carryforwards are approximately \$0.5 million.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**11. License Agreements**

*Angiomax® (bivalirudin)*

In March 1997, the Company entered into an agreement with Biogen, Inc., a predecessor of Biogen Idec Inc., for the license of the anticoagulant pharmaceutical bivalirudin, which the Company has developed as Angiomax. Under the terms of the agreement, the Company acquired exclusive worldwide rights to the technology, patents, trademarks, inventories and know-how related to Angiomax. In exchange for the license, the Company paid \$2.0 million on the closing date and is obligated to pay up to an additional \$8.0 million upon the first commercial sales of Angiomax for the treatment of acute myocardial infarction in the United States and Europe. In addition, the Company is obligated to pay royalties on sales of Angiomax and on any sublicense royalties on a country-by-country basis earned until the later of (1) 12 years after the date of the first commercial sales of the product in a country or (2) the date on which the product or its manufacture, use or sale is no longer covered by a valid claim of the licensed patent rights in such country. Under the terms of the agreement, the royalty rate due to Biogen Idec on sales increases with growth in annual sales of Angiomax. The agreement also stipulates that the Company use commercially reasonable efforts to meet certain milestones related to the development and commercialization of Angiomax, including expending at least \$20 million for certain development and commercialization activities, which the Company met in 1998. The license and rights under the agreement remain in force until the Company's obligation to pay royalties ceases. Either party may terminate the agreement for material breach by the other party, if the material breach is not cured within 90 days after written notice. In addition, the Company may terminate the agreement for any reason upon 90 days prior written notice. The Company recognized royalty expense under the agreement of \$27.2 million in 2006, \$16.1 million in 2005 and \$10.8 million in 2004 for Angiomax sales.

*Cleviprex*

The Company exclusively licensed Cleviprex in March 2003 from AstraZeneca AB for all countries other than Japan. In May 2006, the Company amended its license agreement with AstraZeneca to provide exclusive license rights in Japan in exchange for an upfront payment. The Company acquired this license after having studied Cleviprex under a study and exclusive option agreement with AstraZeneca that the Company entered into in March 2002. Under the terms of the agreement, the Company has the rights to the patents, trademarks, inventories and know-how related to Cleviprex. In exchange for the license, the Company paid \$1.0 million in 2003 upon entering into the license and agreed to pay up to an additional \$5.0 million upon reaching certain regulatory milestones, including a payment of \$1.5 million within 30 days of the acceptance for review of the NDA the Company plans to submit to the FDA for Cleviprex. In addition, the Company will be obligated to pay royalties on a country-by-country basis on future annual sales of Cleviprex, and on any sublicense royalties earned, until the later of (1) the duration of the licensed patent rights which are necessary to manufacture, use or sell Cleviprex in a country or (2) ten years from the Company's first commercial sale of Cleviprex in such country. The licenses and rights under the agreement remain in force on a country-by-country basis until the Company ceases selling Cleviprex in such country or the agreement is otherwise terminated. The Company may terminate the agreement upon 30 days written notice, unless AstraZeneca, within 20 days of having received the Company's notice, requests that the Company enter into good faith discussions to redress its concerns. If the Company cannot reach a mutually agreeable solution with AstraZeneca within three months of the commencement of such discussions, the Company may then terminate the agreement upon 90 days written notice. Either party may

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**11. License Agreements (Continued)**

terminate the agreement for material breach upon 60 days prior written notice, if the breach is not cured within such 60 days.

*Cangrelor*

In December 2003, the Company acquired from AstraZeneca AB exclusive license rights to cangrelor for all countries other than Japan, China, Korea, Taiwan and Thailand. Under the terms of the agreement, the Company has the rights to the patents, trademarks, inventories and know-how related to cangrelor. In exchange for the license, the Company paid in January 2004 an upfront payment upon entering into the license and agreed to make additional payments upon reaching certain regulatory milestones. Under the terms of the license agreement, the Company will be obligated to pay royalties on a country-by-country basis on future annual sales of cangrelor, and on any sublicense royalties earned, until the later of (1) the duration of the licensed patent rights which are necessary to manufacture, use or sell cangrelor in a country or (2) ten years from the Company's first commercial sale of cangrelor in such country. The licenses and rights under the agreement remain in force on a country-by-country basis until the Company ceases selling cangrelor in such country or the agreement is otherwise terminated. The Company may terminate the agreement upon 30 days written notice, unless AstraZeneca, within 20 days of having received the Company's notice, requests that the Company enter into good faith discussions to redress its concerns. If the Company cannot reach a mutually agreeable solution with AstraZeneca within three months of the commencement of such discussions, the Company may then terminate the agreement upon 90 days written notice. Either party may terminate the agreement for material breach upon 60 days prior written notice, if the breach is not cured within such 60 days.

**12. Related Party Transactions and Strategic Alliances**

*UCB Bioproducts*

In December 1999, the Company entered into a commercial supply agreement with UCB Bioproducts S.A., now Lonza Braine, for the development and supply of the Angiomax bulk drug substance. Under the terms of the commercial supply agreement, UCB completed development of a modified production process known as the Chemilog process and filed an amendment in 2001 to its drug master file for regulatory approval of the Chemilog process by the FDA. The Chemilog process was approved by the FDA in May 2003. The Company has agreed to purchase a substantial portion of its Angiomax bulk drug product manufactured using the Chemilog process from UCB at agreed upon prices for a period ending in September 2010. Following the expiration of the agreement, which automatically renews for consecutive three-year periods unless either party provides notice of non-renewal within one year prior to the expiration of the initial term or any renewal term, or if the Company terminates the agreement prior to its expiration, Lonza Braine has agreed to transfer the development technology to the Company. If the Company engages a third party to manufacture Angiomax using this technology prior to bivalirudin becoming a generic drug in the United States, the Company will be obligated to pay UCB a royalty based on the amount paid by the Company to the third party manufacturer. The Company may only terminate the agreement prior to its expiration in the event of a material breach by UCB.

During 2006, 2005 and 2004 the Company recorded \$10.8 million, \$32.4 million and \$25.9 million, respectively, in costs related to UCB's production of Angiomax bulk drug substance. In 2003, the Company recorded \$1.1 million in costs due to Angiomax related development activities. These development costs

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**12. Related Party Transactions and Strategic Alliances (Continued)**

were expensed as research and development in 2003, as FDA approval of the Chemilog processes had not yet been received.

*Nycomed*

In March 2002, the Company entered into an agreement with Nycomed to market and distribute Angiomax in Europe. Nycomed sources, manufactures and markets pharmaceuticals and consumer health products. In September 2004, the Company received authorization from the European Commission to market Angiomax® (bivalirudin) as Angiox (bivalirudin) in the member states of the European Union for use as an anticoagulant in patients undergoing percutaneous coronary interventions.

Nycomed is the Company's exclusive distributor of Angiox in all countries of the European Union excluding Greece, Portugal and Spain. Nycomed paid an initial distributor fee to the Company of \$1.5 million in 2002 and paid to the Company an additional \$2.5 million in 2004 in additional milestones based on regulatory approval in Europe. These payments were recorded as deferred revenue when received and are being amortized over the expected life of this agreement. In addition, in 2002 Nycomed also made a \$1 million equity investment in the Company.

*Strategic Imagery, LLC*

In December 2004, the Company entered into a consulting agreement with Strategic Imagery LLC, a consulting company owned by Mr. Robert Savage, a director of the Company. Under the terms of the consulting agreement, Mr. Savage has agreed to provide consulting services to the Company from time to time on organizational development and senior management coaching. Either party may terminate the consulting agreement at any time upon thirty days written notice. The Company incurred \$49,300 of expenses in 2005 pursuant to the consulting agreement. This agreement expired in December 2005.

**13. Commitments and Contingencies**

The Company's long-term contractual obligations include commitments and estimated purchase obligations entered into in the normal course of business. These include commitments related to purchases of inventory of the Company's products, research and development service agreements, operating leases and selling, general and administrative obligations.

Future estimated contractual obligations as of December 31, 2006 are:

| <b>Contractual Obligations</b>       | <b>2007</b>           | <b>2008</b>      | <b>2009</b>     | <b>2010</b>     | <b>2011</b>     | <b>Later Years</b> | <b>Total</b>     |
|--------------------------------------|-----------------------|------------------|-----------------|-----------------|-----------------|--------------------|------------------|
|                                      | <b>(in thousands)</b> |                  |                 |                 |                 |                    |                  |
| Inventory related commitments        | \$ 16,816             | \$ 4,344         | \$              | \$              | \$              | \$                 | \$ 21,160        |
| Research and development             | 24,139                | 8,153            | 756             |                 |                 |                    | 33,048           |
| Operating Leases                     | 3,020                 | 3,005            | 2,770           | 2,683           | 2,575           | 1,807              | 15,860           |
| Selling, general and administrative  | 1,917                 | 282              |                 |                 |                 |                    | 2,199            |
| <b>Total contractual obligations</b> | <b>\$ 45,892</b>      | <b>\$ 15,784</b> | <b>\$ 3,526</b> | <b>\$ 2,683</b> | <b>\$ 2,575</b> | <b>\$ 1,807</b>    | <b>\$ 72,267</b> |

Included above are inventory-related non-cancellable commitments due to Lonza Braine totaling \$14.8 million during 2007 and \$4.3 million during 2008 and \$2.0 million in remaining Angiomax-related filling, finishing and packaging non-cancellable commitments through 2007. The Company has estimated contractual obligations for research and development activities, of which \$7.5 million is non-cancellable. The Company also has \$2.2 million of estimated contractual obligations for consulting, employment and professional services agreements associated with selling, general and administrative activities, of which \$0.8 million is non-cancellable.



**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**13. Commitments and Contingencies (Continued)**

In addition to the contractual obligations above, the Company has agreed to make payments upon the achievement of sales and regulatory milestones, and agreed to pay royalties, to Biogen Idec under its product license agreement for Angiomax and to AstraZeneca under the Company's product license agreements for Cleviprex and cangrelor. Under the Angiomax license, the Company has agreed to pay up to an additional \$8.0 million upon the first commercial sales of Angiomax for the treatment of acute myocardial infarction in the United States and Europe. Under the Cleviprex license, the Company has agreed to pay up to an additional \$5.0 million upon reaching certain regulatory milestones, including a payment of \$1.5 million within 30 days of the acceptance for review of the NDA the Company plans to submit to the FDA for Cleviprex. Under the cangrelor license, the Company agreed to make milestone payments upon regulatory approval in major markets. The foregoing amounts do not include royalties that the Company may also have to pay.

The Company leases its facilities in Parsippany, New Jersey and Waltham, Massachusetts. The leases for Parsippany and Waltham expire in January 2013 and December 2008, respectively. Rent expense was approximately \$1.6 million, \$1.5 million and \$1.1 million in 2006, 2005 and 2004, respectively.

During 2006 the Company entered into an agreement to lease fleet vehicles for its domestic sales force. The lease agreement provides for management services for the fleet over a five year period and is renewable as vehicles are returned or needed. Expenses associated with these leases in 2006 were less than \$0.1 million. Future minimum annual lease payments will be \$1.2 million for 2007 and 2008 with expenses for the following years included in operating leases in the above table.

*Litigation*

The Company is involved in ordinary and routine matters and litigation incidental to its business. In the opinion of management, there are no matters outstanding that would have a material adverse effect on the consolidated financial position or results of operations of the Company.

**14. Employee Benefit Plan**

The Company has an employee savings and retirement plan which is qualified under Section 401(k) of the Internal Revenue Code. The Company's employees may elect to reduce their current compensation up to the statutorily prescribed limit and have the amount of such reduction contributed to the 401(k) plan. The Company may make matching or additional contributions to the 401(k) plan in amounts to be determined annually by the Board of Directors. The Company has not made any matching or additional contributions to date.

**THE MEDICINES COMPANY**  
**NOTES TO CONSOLIDATED FINANCIAL STATEMENTS (Continued)**  
**December 31, 2006**

**15. Selected Quarterly Financial Data (Unaudited)**

The following table presents selected quarterly financial data for the years ended December 31, 2006 and 2005.

|  | Three Months Ended                           |                  |                   |                  |                  |                  |                   |                  |
|--|--|------------------|-------------------|------------------|------------------|------------------|-------------------|------------------|
|  | Mar. 31,<br>2006                             | June 30,<br>2006 | Sept. 30,<br>2006 | Dec. 31,<br>2006 | Mar. 31,<br>2005 | June 30,<br>2005 | Sept. 30,<br>2005 | Dec. 31,<br>2005 |
|  | <b>(in thousands, except per share data)</b> |                  |                   |                  |                  |                  |                   |                  |
| Net revenue                                | \$ 34,642                                    | \$ 59,372        | \$ 59,580         | \$ 60,357        | \$ 43,572        | \$ 42,595        | \$ 31,920         | \$ 32,120        |
| Cost of sales                              | 8,498  | 15,450           | 14,342            | 13,521           | 10,597           | 10,997           | 6,106             | 7,061            |
| Total operating expenses                   | 48,081                                       | 50,046           | 50,521            | 54,964           | 42,020           | 42,271           | 39,364            | 38,548           |
| Net income/(loss)                          | (12,114 )                                    | 10,914           | 10,673            | 54,253           | 2,338            | 1,251            | (6,232 )          | (5,110 )         |
| Basic net income/(loss) per common share   | \$ (0.24 )                                   | \$ 0.22          | \$ 0.21           | \$ 1.07          | \$ 0.05          | \$ 0.03          | \$ (0.13 )        | \$ (0.10 )       |
| Diluted net income/(loss) per common share | \$ (0.24 )                                   | \$ 0.22          | \$ 0.21           | \$ 1.04          | \$ 0.05          | \$ 0.02          | \$ (0.13 )        | \$ (0.10 )       |
| Market Price High                          | \$ 22.00                                     | \$ 21.34         | \$ 23.25          | \$ 36.18         | \$ 29.95         | \$ 24.95         | \$ 24.55          | \$ 23.70         |
| Low  | \$ 16.54                                     | \$ 16.81         | \$ 18.28          | \$ 22.05         | \$ 20.70         | \$ 20.83         | \$ 20.13          | \$ 15.50         |

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**Schedule II**  
**Valuation and Qualifying Accounts**  
**Year ended December 31, 2006, 2005 and 2004**

|  | <b>Balance at<br/>Beginning<br/>of Period</b> | <b>(Credit) Charged<br/>to Costs and<br/>Expenses(1)</b> | <b>Other<br/>Charges<br/>(Deductions)(2)</b> | <b>Balance<br/>at End<br/>of Period</b> |
|--|---|--|--|---|
| <b>2006</b>  |   |  |  |   |
| Allowances for chargebacks, cash discounts and doubtful accounts | \$ 851  | \$ 8,592   | \$ 8,643                                     | \$ 800                                  |
| <b>2005</b>  |   |  |  |   |
| Allowances for chargebacks, cash discounts and doubtful accounts | \$ 3,574                                      | \$ 4,842   | \$ 7,565                                     | \$ 851                                  |
| <b>2004</b>  |   |  |  |   |
| Allowances for chargebacks, cash discounts and doubtful accounts | \$ 2,226                                      | \$ 9,076   | \$ 7,728                                     | \$ 3,574                                |

- (1) amounts presented herein were charged to and reduced revenues
- (2) represents actual cash discounts, chargeback credits and other deductions

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**INDEX TO EXHIBITS**

| <b>Number</b> | <b>Description</b>   |
|---------------|--|
| 3.1(1)        | Third Amended and Restated Certificate of Incorporation of the registrant, as amended  |
| 3.2(2)        | Amended and Restated By-laws of the registrant, as amended   |
| 10.1(3)*      | 1998 Stock Incentive Plan, as amended  |
| 10.2(4)*      | 2000 Employee Stock Purchase Plan, as amended  |
| 10.3(5)*      | 2000 Outside Director Stock Option Plan, as amended  |
| 10.4(6)       | 2001 Non-Officer, Non-Director Employee Stock Incentive Plan   |
| 10.5(4)*      | 2004 Stock Incentive Plan, as amended  |
| 10.6(7)*      | Form of stock option agreement under 1998 Stock Incentive Plan   |
| 10.7(8)*      | Form of stock option agreement under 2004 Stock Incentive Plan   |
| 10.8(9)       | Form of restricted stock agreement under 2004 Stock Incentive Plan   |
| 10.9(10)      | Amended and Restated Registration Rights Agreement, dated as of August 12, 1998, as amended, by and among the registrant and the other parties thereto |
| 10.10(3)      | License Agreement, dated as of June 6, 1990, by and between Biogen, Inc. and Health Research, Inc., as assigned to the registrant                      |
| 10.11(3)      | License Agreement dated March 21, 1997, by and between the registrant and Biogen, Inc.   |
| 10.12(11)     | License Agreement effective as of March 28, 2003 by and between AstraZeneca AB and the registrant  |
| 10.13(4)      | Amendment No. 1 to License Agreement by and between AstraZeneca AB   |
| 10.14(11)     | License Agreement dated as of December 18, 2003 by and between AstraZeneca AB and the registrant   |
| 10.15(3)      | Chemilog Development and Supply Agreement, dated as of December 20, 1999, by and between the registrant and UCB Bioproducts S.A.                       |
| 10.16(12)     | Sales, Marketing and Distribution Agreement dated March 25, 2002 by and between Nycomed Danmark A/S and the registrant                                 |
| 10.17(11)     | Lease for 8 Campus Drive dated September 30, 2002 by and between Sylvan/Campus Realty L.L.C. and the registrant, as amended                            |
| 10.18(8)      | Third Amendment to Lease for 8 Campus Drive dated December 30, 2004 by and between Sylvan/Campus Realty L.L.C. and the registrant                      |
| 10.19(13)     | Lease for 200 Fifth Avenue, Waltham, MA dated June 19, 2003 by and between Prospect Hill Acquisition Trust and the registrant                          |
| 10.20(3)*     | Employment agreement dated September 5, 1996 by and between the registrant and Clive Meanwell  |
| 10.21(8)*     | Letter Agreement dated December 1, 2004 by and between the registrant and John Kelley  |
| 10.22(14)*    | Letter Agreement dated February 1, 2006 by and between the registrant and Catharine S. Newberry  |

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|            |  |
|------------|--|
| 10.23(14)* | Letter Agreement dated March 2, 2006 by and between the registrant and Glenn P. Sblendorio   |
| 10.24(8)*  | Summary of Board of Director Compensation  |
| 10.25*     | Form of Amended and Restated Management Severance Agreement dated as of August 17, 2006 by and between the registrant and each of Clive Meanwell and John Kelley                         |
| 10.26*     | Form of Amended and Restated Management Severance Agreement dated as of August 17, 2006 by and between the registrant and each of Glenn Sblendorio, Paul Antinori and Catharine Newberry |
| 10.27(14)* | Form of Lock-Up Agreement dated as of December 23, 2005 by and between the registrant and each of its executive officers and directors   |
| 21         | Subsidiaries of the registrant   |
| 23         | Consent of Ernst & Young LLP, Independent Registered Accounting Firm   |
| 31.1       | Chief Executive Officer Certification pursuant to Rule 13a-14(a) of the Securities Exchange Act of 1934, as adopted pursuant to Section 302 of the Sarbanes-Oxley Act of 2002            |
| 31.2       | Chief Financial Officer Certification pursuant to Rule 13a-14(a) of the Securities Exchange Act of 1934, as adopted pursuant to Section 302 of the Sarbanes-Oxley Act of 2002            |
| 32.1       | Chief Executive Officer Certification pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002   |
| 32.2       | Chief Financial Officer Certification pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002   |

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\* Management contract or compensatory plan or arrangement filed as an exhibit to this form pursuant to Items 15(a) and 15(c) of Form 10-K

Confidential treatment requested as to certain portions, which portions have been omitted and filed separately with the Securities and Exchange Commission

- (1) Incorporated by reference to the exhibits to amendment no. 1 to the registrant's registration statement on Form 8-A/A (registration no. 000-31191)
- (2) Incorporated by reference to exhibit 10.3 to the registrant's quarterly report on Form 10-Q for the quarter ended June 30, 2006
- (3) Incorporated by reference to the exhibits to the registration statement on Form S-1 (registration no. 333-37404)
- (4) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended June 30, 2006
- (5) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended March 31, 2003
- (6) Incorporated by reference to the exhibits to the registration statement on Form S-8 (registration no. 333-74612)

(7) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended June 30, 2006

(8) Incorporated by reference to the exhibits to the registrant's annual report on Form 10-K for the year ended December 31, 2004

(9) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended March 31, 2006

(10) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended June 30, 2002

(11) Incorporated by reference to the exhibits to the registrant's annual report on Form 10-K for the year ended December 31, 2003

(12) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended March 31, 2002

(13) Incorporated by reference to the exhibits to the registrant's quarterly report on Form 10-Q for the quarter ended June 30, 2003

(14) Incorporated by reference to the exhibits to the registrant's annual report on Form 10-K for the year ended December 31, 2005

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