

---

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

**FORM 20-F**

(Mark One)

**REGISTRATION STATEMENT PURSUANT TO SECTION 12(b) OR (g) OF THE SECURITIES EXCHANGE ACT OF 1934**

OR

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

For the fiscal year ended June 30, 2005

OR

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

For the transition period from \_\_\_\_\_ to \_\_\_\_\_

OR

**SHELL COMPANY REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934**

Date of event requiring this shell company report

Commission file number 000-51505

**Pharmaxis Ltd**

(Exact name of Registrant as specified in its charter)

---

(Translation of Registrant's name into English)

**Australia**

(Jurisdiction of incorporation or organization)

**Unit 2, 10 Rodborough Road, Frenchs Forest, NSW 2086, Australia**

(Address of principal executive offices)

Securities registered or to be registered pursuant to Section 12(b) of the Act.

Title of each class

Name of each exchange on which registered

None

Securities registered or to be registered pursuant to Section 12(g) of the Act.

**Ordinary Shares in the form of American Depositary Shares, evidenced by American Depositary Receipts**  
(Title of Class)

Securities for which there is a reporting obligation pursuant to Section 15(d) of the Act.

None

As of June 30, 2005 the number of outstanding shares of each of the issuer's classes of capital or common stock was as follows: 134,770,092 fully paid Ordinary Shares.

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 which financial statement item the registrant has elected to follow.

Item 17  Item 18

If this is an annual report, indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act).

Yes  No

(APPLICABLE ONLY TO ISSUERS INVOLVED IN BANKRUPTCY PROCEEDINGS DURING THE PAST FIVE YEARS)

Indicate by check mark whether the registrant has filed all documents and reports required to be filed by Sections 12, 13 or 15(d) of the Securities Exchange Act of 1934 subsequent to the distribution of the securities under a plan confirmed by a court.

Yes  No

---

## TABLE OF CONTENTS

<u>FORWARD LOOKING STATEMENTS</u>	1
ITEM 1. <u>IDENTITY OF DIRECTORS, SENIOR MANAGEMENT AND ADVISORS</u>	2
ITEM 2. <u>OFFER STATISTICS AND EXPECTED TIMETABLE</u>	2
ITEM 3. <u>KEY INFORMATION</u>	2
ITEM 4. <u>INFORMATION ON THE COMPANY</u>	26
ITEM 5. <u>OPERATING AND FINANCIAL REVIEW AND PROSPECTS</u>	43
ITEM 6. <u>DIRECTORS, SENIOR MANAGEMENT AND EMPLOYEES</u>	54
ITEM 7. <u>MAJOR SHAREHOLDERS AND RELATED PARTY TRANSACTIONS</u>	72
ITEM 8. <u>FINANCIAL INFORMATION</u>	76
ITEM 9. <u>THE OFFER AND LISTING</u>	77
ITEM 10. <u>ADDITIONAL INFORMATION</u>	78
ITEM 11. <u>QUANTITATIVE AND QUALITATIVE DISCLOSURES ABOUT MARKET RISK</u>	94
ITEM 12. <u>DESCRIPTION OF SECURITIES OTHER THAN EQUITY SECURITIES</u>	94
ITEM 13. <u>DEFAULTS, DIVIDEND ARREARAGES AND DELINQUENCIES</u>	95
ITEM 14. <u>MATERIAL MODIFICATIONS TO THE RIGHTS OF SECURITY HOLDERS AND USE OF PROCEEDS</u>	95
ITEM 15. <u>CONTROLS AND PROCEDURES</u>	95
ITEM 16A. <u>AUDIT COMMITTEE FINANCIAL EXPERT</u>	96
ITEM 16B. <u>CODE OF ETHICS</u>	96
ITEM 16C. <u>PRINCIPAL ACCOUNTANT FEES AND SERVICES</u>	96
ITEM 16D. <u>EXEMPTIONS FROM THE LISTING STANDARDS FOR AUDIT COMMITTEES</u>	97
ITEM 16E. <u>PURCHASES OF EQUITY SECURITIES BY THE ISSUER AND AFFILIATED PURCHASERS</u>	97
ITEM 17. <u>FINANCIAL STATEMENTS</u>	98
ITEM 18. <u>FINANCIAL STATEMENTS</u>	98
ITEM 19. <u>EXHIBITS</u>	98

In this registration statement on Form 20-F, the terms “we,” “our,” “us,” “Pharmaxis Ltd”, “Pharmaxis” and “the Company” refer to Pharmaxis Ltd. Unless otherwise noted herein, all references to “dollars” and dollar amounts are references to Australian dollars. Our Australian company number is ACN 082 811 630 and our Australian business number is ABN 75 082 811 630.

## **FORWARD LOOKING STATEMENTS**

Some of the statements made under “Item 3. Key Information – D. Risk Factors” “Item 4. Information on the Company” “Item 5. Operating and Financial Review and Prospects” and elsewhere in this report on Form 20-F constitute forward-looking statements. In some cases, you can identify forward-looking statements by terminology such as “may,” “will,” “should,” “expects,” “plans,” “anticipates,” “believes,” “estimates,” “predicts,” “potential,” or “continue,” or the negative of these terms or other comparable terminology. These statements are only current predictions and are subject to known and unknown risks, uncertainties and other factors that may cause our or our industry’s actual results, levels of activity, performance or achievements to be materially different from those anticipated by the forward-looking statements. These factors include, among other things, those listed under “Item 3. Key Information – D. Risk Factors” and elsewhere in this report on Form 20-F.

Although we believe that the expectations reflected in the forward-looking statements are reasonable, we cannot guarantee future results, levels of activity, performance or achievements. Except as required by law, we are under no duty to update or revise any of our forward-looking statements, whether as a result of new information, future events or otherwise, after the date of this report on Form 20-F.

## PART I

### ITEM 1. IDENTITY OF DIRECTORS, SENIOR MANAGEMENT AND ADVISORS

#### A. Directors and Senior Management

Not applicable

#### B. Advisors

Not applicable.

#### C. Auditors

Not applicable.

### ITEM 2. OFFER STATISTICS AND EXPECTED TIMETABLE

Not applicable.

### ITEM 3. KEY INFORMATION

#### A. Selected Financial Data

The following table presents our selected financial data for the dates and periods indicated. This data should be read together with Item 5 - "Operating and Financial Review and Prospects." The statement of operations data for the years ended June 30, 2003, 2004 and 2005, and for the period from inception (May 29, 1998) to June 30, 2005, and the balance sheet data as of June 30, 2004 and 2005, were derived from our audited financial statements and related notes thereto included elsewhere in this annual report, were prepared in accordance with U.S. GAAP and are presented in Australian dollars (except as otherwise noted). The statement of operations data for the year ended June 30, 2002, and the balance sheet data as of June 30, 2002 and 2003, are derived from our audited financial statements and related notes thereto which are not included in this report, were prepared in accordance with U.S. GAAP and are presented in Australian dollars (except as otherwise noted). Selected financial data for the year ended June 30, 2001 has been omitted because the underlying, discrete standalone financial information is unavailable, and therefore, such financial data cannot be provided without unreasonable expense or effort. Our fiscal year ends on June 30. We designate our fiscal year by the year in which that fiscal year ends; e.g., fiscal year 2005 refers to our fiscal year ended June 30, 2005.

	Year ended June 30,				Period from inception (May 29, 1998) to June 30, 2005	Year ended June 30, 2005(2)
	2002	2003	2004	2005	A\$	U.S.\$
	A\$	A\$	A\$	A\$		
(in thousands, except per share and footnote data)						
<b>Statement of Operations Data:</b>						
Revenue	\$ —	\$ —	\$ —	\$ —	\$ —	\$ —
Operating expenses:						
Research and development(1)	486	925	4,806	7,885	14,664	6,007
General and administrative	141	981	2,182	3,105	6,550	2,365
Commercial	—	—	—	807	807	615
Amortization of intangible assets	83	86	89	90	485	69
Fair value of stock options issued to employees related to:						
Research and development	32	261	253	115	753	88
Commercial	—	—	—	116	116	88
General and administrative	37	122	279	29	527	22
Total operating expenses	779	2,375	7,609	12,147	23,902	9,254
Loss from operations	(779)	(2,375)	(7,609)	(12,147)	(23,902)	(9,254)
Interest and other income	44	327	1,123	1,702	3,256	1,297
Amortization of preference share issue expenses	—	(65)	(161)	—	(226)	—
Net loss	\$ (735)	\$ (2,113)	\$ (6,647)	\$ (10,445)	\$ (20,872)	\$ (7,957)
Basic and diluted net loss per share	\$ (0.07)	\$ (0.19)	\$ (0.09)	\$ (0.08)	\$ (0.61)	\$ (0.06)

Weighted average number of ordinary shares used in calculating basic and diluted net loss per share(3)	11,200	11,200	75,744	123,933	34,068	123,933
--	--------	--------	--------	---------	--------	---------

(1) Research and development expenses have been reduced by government research grants of A\$663,000, A\$751,000, A\$1,105,000, A\$1,132,000 and A\$4,551,000 in fiscal 2002, 2003, 2004 and 2005, and the period from inception (May 29, 1998) to June 30, 2005, respectively.

- (2) The amounts have been translated into U.S. dollars from Australian dollars based upon the noon buying rates in New York City as determined by the Federal Reserve Bank of New York on June 30, 2005. These translations are merely for the convenience of the reader and should not be construed as representations that the Australian dollar amounts actually represent such U.S. dollar amounts or could be converted into U.S. dollars at the rate indicated.
- (3) Fiscal 2002, 2003 and 2004, and the period from inception (May 29, 1998) to June 30, 2005, have been retroactively adjusted for an eight-for-one share split effected immediately prior to the closing of our initial public offering in Australia. The increase in ordinary shares in fiscal 2004 is attributable to our Australian initial public offering in which a total of 50,000,000 new ordinary shares were issued, and immediately before which 46,816,000 convertible redeemable preference shares converted to ordinary shares on a one-for-one basis. The increase in ordinary shares in fiscal 2005 is primarily attributable to a share placement and share purchase plan in which a total of 26,362,092 new ordinary shares were issued.

	As of June 30,				
	2002	2003	2004	2005	2005
	A\$	A\$	A\$	A\$	U.S.\$(2)
	(in thousands)				
<b>Balance Sheet Data:</b>					
Cash and cash equivalents	\$ 751	\$ 7,384	\$25,101	\$33,268	\$25,344
Total assets	2,144	10,459	28,111	37,836	28,824
Convertible redeemable preference shares(1)	2,000	11,630	—	—	—
Total shareholders' (deficit) equity	(46)	(1,776)	26,631	35,467	27,019

- (1) Convertible redeemable preference shares were converted to ordinary shares immediately prior to the quotation of our ordinary shares on the Australian Stock Exchange after our Australian initial public offering in November 2003.
- (2) The amounts have been translated into U.S. dollars from Australian dollars based upon the noon buying rates in New York City as determined by the Federal Reserve Bank of New York on June 30, 2005. These translations are merely for the convenience of the reader and should not be construed as representations that the Australian dollar amounts actually represent such U.S. dollar amounts or could be converted into U.S. dollars at the rate indicated.

## Exchange Rate Information:

The following tables present exchange rates of the Australian dollar into the U.S. dollars for the periods indicated. Annual averages are calculated using the average of month-end rates of the relevant year. Monthly averages are calculated using the average of the daily rates during the relevant period.

Period	Average	High	Low
	U.S.\$	U.S.\$	U.S.\$
Five most recent fiscal years			
2001	\$0.5320	\$0.5996	\$0.4828
2002	0.5240	0.5748	0.4841
2003	0.5884	0.6729	0.5280
2004	0.7155	0.7979	0.6390
2005	0.7568	0.7974	0.6880
Six most recent months			
June 2005	0.7667	0.7792	0.7498
July 2005	0.7524	0.7661	0.7403
August 2005	0.7614	0.7739	0.7469
September 2005	0.7651	0.7731	0.7537
October 2005	0.7535	0.7630	0.7468
November 2005	0.7353	0.7451	0.7267
As of December 8, 2005, the exchange rate was \$0.7507			

## B. Capitalization and Indebtedness

Not applicable

## C. Reasons for the Offer and Use of Proceeds

Not applicable.

## D. Risk Factors

*The following risks relate specifically to the Company's business and should be considered carefully. Our business, financial condition and results of operations could be materially and adversely affected by any of the following risks. As a result, the trading price of our ordinary shares and our American Depositary Shares, or ADSs, could decline and the holders could lose part or all of their investment.*

### **Risks Related to Our Business**

**We are at an early stage of our development as an integrated pharmaceutical company and we do not have, and may never have, any products that generate revenues. Unless we are able to generate sufficient product revenue, we will continue to incur losses from operations and may not achieve or maintain profitability.**

We are at an early stage of our development as an integrated pharmaceutical company. We were incorporated in May 1998 and we have a limited operating history on which to evaluate our business and prospects. To date, we do not have, and may never have, any products that generate revenues. We have funded our operations through the private sales of our securities, the initial public offering of our securities in Australia and a share purchase plan. We have also received funding from government research grants and earned interest income on invested funds.

We have incurred losses in each year since our inception and expect to continue to incur substantial losses. We incurred losses of approximately A\$2.1 million, A\$6.6 million and A\$10.4 million in the fiscal years ended June 30, 2003, 2004 and 2005, respectively. Our accumulated losses from inception to June 30, 2005 are A\$20.9 million. These losses, among other things, have had and will continue to have an adverse effect on our shareholders' equity



and working capital. Unless we are able to generate sufficient product revenue, we will continue to incur losses from operations and may not achieve or maintain profitability.

We expect our expenses to increase significantly in the short term in connection with:

- ongoing clinical trials of Aridol. Although we have completed a number of clinical trials in respect of Aridol, further Phase III clinical trials of Aridol are required to be completed in the U.S. before an application for marketing authorization can be filed with the U.S. Food and Drug Administration, or the FDA. These U.S. clinical trials are expensive;
- the regulatory marketing authorization process to approve the sale of Aridol. Aridol is the first of our product candidates to complete Phase III trials in any jurisdiction and the first of our product candidates for which we have sought marketing authorization. The work involved in seeking regulatory marketing authorization for Aridol is extensive, time consuming and expensive;
- the further expansion of our existing manufacturing facilities and an increase in the number of manufacturing personnel to enable the commercial manufacture of Aridol. The costs associated with the further expansion of our manufacturing facilities represent a significant short-term increase in our expenses. The increased costs relating to additional manufacturing personnel also represents a significant increase in our expenses which are likely to continue over time;
- the development of our sales and marketing capability. Our existing sales and marketing capability is currently limited and, to date, has not represented a significant cost to us. In the short term, our sales and marketing capability must be increased to enable the sales and marketing of Aridol in Europe and Australia and thereafter in the U.S.;
- the continuation of simultaneous Phase II clinical trials of Bronchitol and the commencement of Phase III clinical trials of Bronchitol. These clinical trials are carried out in a number of jurisdictions and are expensive. Our expenses will increase as we commence new clinical trials or we progress existing trials to more advanced phases. The more advanced clinical trials typically require more clinical trial participants, clinical trial sites and research investigators than earlier stage clinical trials and are consequently more expensive;
- the commencement of Phase I clinical trials of PXS64, which will represent a significant new expense for us; and
- although we already have existing expenses associated with the preclinical testing of PXS2076 and other products candidates, in the short term, we will continue to incur expenses as a result of preclinical testing of PXS2076 and any other clinical trials and preclinical testing that we may initiate.

We also expect to incur increased general and administrative expenses in support of our increased operations as well as the increased costs to operate as a company listed on the Australian Stock Exchange and on the Nasdaq National Market. Over the longer term, the costs referred to above will fluctuate, primarily dependant on the number and type of clinical trials being undertaken by us at any one time. Costs will also increase if we are able to progress any further clinical trial candidates from preclinical testing to clinical trials.

**We may not become profitable if our initial product candidates are unsuccessful in clinical trials or we are unable to obtain regulatory approvals. Even if we receive regulatory approval for any product candidates, profitability will depend on our ability to generate revenues from the sale of our products or the licensing of our technology.**

We cannot be certain that the clinical development of Aridol or Bronchitol or any of our other product candidates in preclinical testing or clinical development will be successful, that they will receive the regulatory approvals required to commercialize them, or that any of our research and development programs will yield additional product candidates suitable for investigation through clinical trials.

If Aridol or Bronchitol is unsuccessful in clinical trials or we are unable to obtain marketing authorization, we may not be profitable. We have recently completed the Phase III clinical trials of Aridol necessary for Australian and European registration of Aridol. However, we cannot be certain that marketing approval will be granted in

Australia and/or Europe. Further Phase III clinical trials are required to be completed in the U.S. before an application for marketing authorization can be filed with the FDA. There is a risk that these Phase III clinical trials in the U.S. may not be successful and that marketing authorization may not be granted in the U.S. We are undertaking simultaneous Phase II clinical trials of Bronchitol and Phase III clinical trials of Bronchitol. Clinical trials of Bronchitol will continue for several years, but may take significantly longer to complete. There is a risk that these clinical trials of Bronchitol may not be successful or that marketing approval may not be granted in the future. If we are not able to successfully complete clinical trials of Aridol (in the U.S.) and Bronchitol, and if we are unable to subsequently obtain marketing authorization of Aridol and Bronchitol, we may not be profitable.

The process to develop, obtain regulatory approval for, and commercialize potential product candidates is long, complex and costly. Even if we receive regulatory approval for any product candidates, profitability will depend on our ability to generate revenues from the sale of our products or the licensing of our technology that will offset the significant and continuing expenditures required for us to advance our research, protect and extend our intellectual property rights and develop, manufacture, license, market, distribute and sell our technology and products successfully. Our ability to generate revenue depends on a number of factors, including our ability to:

- successfully conduct and complete clinical trials for Aridol (in the U.S.) and Bronchitol and our other product candidates;
- develop and obtain regulatory marketing authorization, as well as approvals concerning pricing and reimbursement, which may be necessary in some of the E.U. member states, for Aridol and Bronchitol in our target markets and, in the future, to develop and obtain regulatory marketing authorization for our other product candidates;
- manufacture or obtain commercial quantities of Aridol and Bronchitol or our other product candidates at acceptable cost levels; and
- successfully market and sell Aridol, Bronchitol and our other product candidates. In circumstances where we have licensed our technology to third parties, our ability to generate revenue will depend on the success of the licensor of the technology to successfully market and sell the licensed technology.

Although we have a pipeline of potential product candidates and our own internal and contract research and development team, our business is currently substantially dependent on our ability to complete development, obtain regulatory approval for, and successfully commercialize Aridol and Bronchitol in a timely manner. If we are unable to successfully commercialize Aridol and/or Bronchitol, we may not be able to earn sufficient revenues to continue our business. If we fail to become and remain profitable, or if we are unable to fund our continuing losses, there would be a material adverse effect on our business and the holders of our ordinary shares and ADSs could lose all or part of their investment.

**Unsuccessful or delayed marketing authorization could increase our future development costs or impair our future revenue. Approvals that may be given may not cover all the indications for which we seek approval or may contain significant limitations.**

To receive regulatory authorization for the commercial sale of our product candidates, we must conduct preclinical studies and clinical trials to demonstrate safety and efficacy in humans. This process of attempting to gain regulatory approval is expensive and can take many years, and failure can occur at any stage of the testing. Our failure to adequately demonstrate the safety and efficacy of Aridol and/or Bronchitol and/or any of our other product candidates will prevent regulatory approval and commercialization of such product candidates. Our inability to successfully and effectively complete clinical trials for our product candidates, in particular Aridol and Bronchitol, will severely harm our business and we may not be profitable.

Significant delays in clinical development could materially increase our product development costs, delay our receipt of revenue or allow our competitors to bring product candidates to market before we do, impairing our ability to effectively commercialize Aridol and Bronchitol or our other product candidates.

In addition, any authorization we may obtain may not cover all of the clinical indications for which we seek approval. Also, an authorization might contain significant limitations in the form of narrow indications, warnings, precautions or contraindications with respect to conditions of use.

**We will continue to need significant amounts of additional capital that may not be available to us on favorable terms or at all or which may be dilutive.**

To date, we have funded our operations and capital expenditures with proceeds from the sale of our securities, government grants and interest on investments.

In order to achieve our goal of being a fully integrated pharmaceutical company and to conduct the lengthy and expensive research, preclinical studies, clinical trials, regulatory approval process, manufacture, sales and marketing necessary to complete the full development of our product candidates, we will require substantial additional funds in addition to the funds received in connection with the U.S. public offering and the Australian placement completed in November 2005. In particular, we will require substantial additional funding as a result of our strategy of not always entering into strategic alliances or partnerships in order to commercialize our product candidates.

To meet these financing requirements, we may raise funds through public or private equity offerings, debt financings, and through other means, including collaborations and license agreements. Raising additional funds by issuing equity or convertible debt securities may cause our shareholders to experience significant additional dilution in their ownership interests. Raising additional funds through debt financing, if available, may involve covenants that restrict our business activities. Additional funding may not be available to us on favorable terms, or at all. If we are unable to obtain additional funds, we may be forced to delay, reduce the scope or terminate our clinical trials and the development, manufacturing and marketing of our products. To the extent that we raise additional funds through collaborations and licensing arrangements, we may have to relinquish valuable rights and control over our technologies, research programs or product candidates, or grant licenses on terms that may not be favorable to us.

**If we fail to obtain additional financing, we may be unable to fund our operations and commercialize our product candidates.**

We expect that our cash expenditure will increase for the next several years, and that we will spend substantial amounts to complete the clinical development and commercialization of Aridol, Bronchitol, PXS64 and our other product candidates, and to license or acquire other product candidates. We believe that our existing cash and cash equivalents will be sufficient to meet our projected operating requirements for at least 12 months.

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

- the scope, results, rate of progress, timing and costs of preclinical studies and clinical trials and other development activities. The funding requirements for clinical trials of Aridol, Bronchitol and PXS64 are significant. The funding requirements for the preclinical testing and potential future clinical testing of PXS2076 and any other testing that we may initiate is also significant;
- the costs and timing of seeking and obtaining regulatory approvals;
- the costs of filing, prosecuting, defending and enforcing any patent claims and other intellectual property rights;
- the costs of developing our sales and marketing capabilities and establishing distribution capabilities. We will also need to establish a distribution capability;
- the cost of expanding our manufacturing capabilities. Our existing facilities are capable of limited commercial manufacture and we have recently expanded the commercial manufacturing capacity of our existing facilities to coincide with the proposed commercial launch of Aridol in Europe and Australia;
- the cost of additional management and scientific, manufacturing and sales and marketing personnel. We will be required to increase the number of our personnel over time;
- the terms, timing and cash requirements of any future acquisitions, collaborative arrangements, licensing of product candidates or investing in businesses, product candidates and technologies;
- the costs of securing coverage, payment and reimbursement of our product candidates, if any of our product candidates receive regulatory approval; and
- the effects of competing clinical, technological and market developments.

If we are not able to secure additional funding when needed, we may have to delay, reduce the scope of, or eliminate one or more of our clinical trials or research and development programs or future commercialization efforts.

**The suspension or termination of our government research grants may result in lost revenue. We may also be required to repay previously received grant revenue in certain circumstances which would have an adverse effect on our cash position.**

We currently receive substantial grant funding under two separate grant agreements with the Commonwealth of Australia. There is a risk that we will not be entitled to the grant payments for failing to incur eligible expenditure or failing to undertake activities associated with the applicable grant or otherwise for failing to satisfy the relevant conditions in the applicable grant agreement. Furthermore, there is a risk we will not be entitled to the funds under the grant agreements, including, if the Commonwealth of Australia has insufficient funding for the relevant grant program, if we fail to submit reports when required, if we have not otherwise complied with our obligations under the relevant funding agreement, or if the Commonwealth of Australia is entitled to or does terminate the relevant agreements. The Commonwealth of Australia may terminate the grant agreements on different bases under different grant agreements, including by giving us written notice of termination if we are in breach of the relevant agreement and if in the opinion of the Commonwealth of Australia the breach is not capable of being remedied, or if capable of being remedied it is not remedied after receipt of written notice, if we fail to submit reports, if our research and development activities or the quality of those activities do not satisfy the grant eligibility criteria, if there is a change of control of us or if we become insolvent.

In certain circumstances where we fail to use our best endeavors to commercialize the project within a reasonable time of completion of the project or upon termination of a grant due to our breach of agreement or our insolvency, the Commonwealth of Australia may require us to repay some or all of the grant. If required to repay the grant amounts, we may be required to reallocate funds needed to continue the commercialization of our products and such repayment may have a material adverse effect on our cash position and us.

**Currency fluctuations may expose us to increased costs and revenue decreases.**

Our business may in the future be affected by fluctuations in foreign exchange rates. Currency fluctuations could, therefore, cause our costs to increase and revenues to decline. The majority of our expenses will continue to be denominated in Australian dollars although we will also be expending cash in other denominations, including U.S. dollars and the European euro. In the last two years, the Australian dollar has as a general trend appreciated against the U.S. dollar. If this trend continues, this may have a positive effect on any costs which we incur in the U.S. but may have an adverse effect on our revenues sourced from the U.S. We cannot anticipate whether this trend will continue in respect of the U.S. dollar. The exchange rates of the Australian dollar to the European euro have fluctuated over the same period. In circumstances where the Australian dollar devalues against either or both of the U.S. dollar or the European euro, this may have an adverse effect on our costs incurred in either the U.S. or the European Union (as applicable) but may have a positive effect on any revenues which we source from the U.S. or the European Union (as applicable). The same principles apply in respect of our costs and revenues in other jurisdictions. In addition, we conduct clinical trials in many different countries and we have manufacturing of some of our product candidates undertaken outside of Australia, which exposes us to potential cost increases resulting from fluctuations in exchange rates. We do not currently have any plans to hedge the effect of currency fluctuations on our overseas expenditures. We manage our currency risks by settling foreign currency payables immediately upon recognition of a foreign currency liability.

***Risks Related to Research and Development of Our Products***

**Clinical trials are expensive, time consuming, subject to delay and their outcome is uncertain and may not be completed at all.**

Before we can obtain regulatory authorization for the commercial sale of any product or product candidate, we are required to complete preclinical development and extensive clinical trials in humans to demonstrate its safety and efficacy. Preclinical development and clinical trials are subject to extensive regulation by the FDA and other

regulatory authorities in the United States, the European Union, Australia and elsewhere. In addition, clinical trials must be conducted with product candidates produced under applicable current Good Manufacturing Practices. Clinical trials are expensive and complex, can take many years, are often subject to delay and have uncertain outcomes. We have recently completed the Phase III clinical trials of Aridol that we believe are necessary for Australian and European marketing authorization of Aridol. Further Phase III clinical trials are required to be completed in the U.S. before an application for marketing authorization can be filed with the FDA. The FDA has accepted an Investigational New Drug Application, or IND, for Aridol and Bronchitol. We have yet to reach an agreement with the FDA for U.S. trials of Bronchitol. Clinical trials of our product candidates, including Aridol, Bronchitol and PXS64, will continue for several years, but may take significantly longer to complete.

There are numerous factors that could affect the timing of the commencement, continuation and completion of clinical trials which may delay the clinical trials or prevent us from completing these trials successfully, including but not limited to:

- delays in securing clinical investigators or trial sites for our clinical trials, scheduling conflicts with participating clinicians and clinical institutions, and delays in obtaining institutional review board, or IRB, and other regulatory approvals to commence a clinical trial. There are a limited number of clinical investigators and clinical trials sites worldwide able to conduct the clinical trials required by us. Clinical investigators and trial sites may have demands from a number of companies competing to use their resources;
- slower than anticipated recruitment and enrollment of patients who meet the trial eligibility criteria or the loss of patients during the course of the clinical trials. By way of example, we experienced delays on our Bronchitol clinical trials for the treatment of cystic fibrosis because recruitment was slower than anticipated. We managed the enrollment problem by modifying the protocol to widen the eligibility of clinical trial participants, adding additional clinical trial sites and assisting with resourcing at clinical trial sites. We may not be able to successfully manage such enrollment problems in the future;
- the requirement to repeat or undertake large clinical trials. Our Phase II and Phase III clinical trials involve a large number of patients and are typically carried out in different jurisdictions and may also need to be repeated if required by regulatory authorities. Although we have completed Phase III clinical trials for Aridol to enable us to seek marketing approval in Europe and Australia, we are required to undertake additional Phase III clinical trials in the U.S. prior to seeking marketing authorization of Aridol in the U.S.;
- negative or inconclusive results from clinical trials, or deficiencies in the conduct of the clinical trials may require us to repeat clinical trials;
- unforeseen safety issues or unforeseen adverse side effects or fatalities or other adverse events arising during a clinical trial due to medical problems that may or may not be related to clinical trial treatments;
- the product candidate may not be as good as current therapies. For example, Aridol must prove to be convenient and effective as a lung capacity test which assists in the identification of and determines the severity of asthma. In addition, Bronchitol must improve the quality of life and health of people with chronic obstructive lung diseases such as cystic fibrosis and bronchiectasis;
- quality or stability of the product candidate may fall below acceptable standards;
- shortages of available product supply. We may be required to simultaneously provide product to patients in a range of jurisdictions and there may be shortages or delays in supplying the product in those jurisdictions;
- uncertain dosing issues; and
- inability to monitor patients adequately during or after treatment or problems with investigator or patient compliance with the trial protocols.

Due to the foregoing and other factors, the regulatory approval of Aridol in the U.S. or in other jurisdictions, Bronchitol, PXS64 and any of our other future product candidates could take a significantly longer

time to gain regulatory approval than we expect or these products may never gain approval, which could reduce or eliminate our revenue by delaying or terminating the potential commercialization of our products or product candidates. If we suffer any significant delays, setbacks or negative results in, or termination of, our clinical trials, we may be unable to continue the development of our products or product candidates or generate revenue and our business may be materially adversely affected.

**Ongoing and future clinical trials of our product candidates may not show sufficient safety or efficacy to obtain requisite regulatory approvals.**

Ongoing and future clinical trials of our product candidates may not show sufficient safety or efficacy to obtain regulatory approval for marketing. Phase I and Phase II clinical trials are not primarily designed to test the efficacy of a product candidate but rather to test safety, to study pharmacokinetics and pharmacodynamics and to understand the product candidate's side effects at various doses and schedules. Furthermore, success in preclinical and early clinical trials does not ensure that later large-scale trials will be successful nor does it predict final results. Acceptable results in early trials may not be repeated in later trials. Further, Phase III clinical trials of Aridol are required to be completed in the U.S. before an application for marketing authority can be filed with the FDA and there is a risk that these may not show sufficient safety or efficacy to obtain regulatory approval for marketing Aridol in the U.S. despite the completion of the Phase III trials of Aridol in other jurisdictions. Likewise, clinical trials of Bronchitol and our other product candidates may not show sufficient safety or efficacy to obtain regulatory approval for marketing.

We may conduct lengthy and expensive clinical trials of our product candidates, only to learn that the product candidate is not an effective treatment. A number of companies in the biotechnology and pharmaceutical industries have suffered significant setbacks in advanced clinical trials, even after promising results in earlier trials. In addition, clinical results are frequently susceptible to varying interpretations that may delay, limit or prevent regulatory approvals. Negative or inconclusive results or adverse medical events during a clinical trial could cause the clinical trial to be delayed, redone or terminated. In addition, failure to construct appropriate clinical trial protocols or other factors could result in the test or control group experiencing a disproportionate number of adverse events and could cause a clinical trial to be redone or terminated. The length of time necessary to complete clinical trials and to submit an application for marketing approval for a final decision by applicable regulatory authorities may also vary significantly based on the type, complexity and novelty of the product candidate involved, as well as other factors.

**Due to our reliance on contract research organizations, hospitals and investigators to conduct clinical trials, we are unable to directly control the timing, conduct and expense of our clinical trials. We also use third parties to provide research and development services and do not have direct control of the timing, conduct and expense of certain of our research programs.**

We rely on third parties such as contract research organizations, hospitals and research investigators to provide services in connection with our clinical trials. Our clinical trials are conducted by a number of third parties at a number of different sites in different jurisdictions. With respect to the clinical trials we have completed to date, we have been involved with approximately 25 investigators, and a small number of contract research organizations, consultants and other professionals. We are in the process of preparing for clinical trials of Bronchitol for the treatment of cystic fibrosis in the United Kingdom which will involve two investigators and clinical trials of Bronchitol for the treatment of cystic fibrosis in Canada which will involve six investigators and one contract research organization. We are also preparing for our Phase III clinical trials of Aridol as a lung capacity test in the U.S. which will involve approximately 25 investigators and one contract research organization.

We believe that the agreements that we enter into with these third parties are customary for agreements relating to the provision of clinical trial services. The agreements set out the parameters and protocols for the relevant clinical trials, set out the amount payable by us, as well as setting out the rights and obligations of the third parties and us.

To date, we have been able to manage the use of these third parties in order to effectively carry out our clinical trials. If these third parties do not successfully carry out their contractual duties or regulatory obligations or meet expected deadlines, if the third parties need to be replaced or if the quality or accuracy of the data they obtain is compromised due to the failure to adhere to our clinical protocols or regulatory requirements or for other reasons, our clinical trials may be extended, delayed, suspended or terminated, and we may not be able to obtain regulatory

approval for or successfully commercialize our product candidates. Although there are a range of suitable institutions and investigators that would be able to conduct the clinical trials on our behalf, there is no guarantee that we will be able to enter into any such arrangement on acceptable terms, if at all.

Although we have our own internal research and development team, we also contract with the ANU Enterprises Pty Ltd (part of the Australian National University) to provide us with certain human resources and services to identify product candidates and to undertake preclinical research and development of product candidates. If ANU Enterprises Pty Ltd does not successfully carry out their contractual duties and provide suitable human resources and other services, or if ANU Enterprises Pty Ltd needs to be replaced or if the quality of the human resources and services they provide are inadequate, our research and development programs may be extended, delayed, suspended or terminated. In addition to our arrangements with ANU Enterprises Pty Ltd, we currently utilize the services of approximately twenty other organizations that conduct research and development services on our behalf from time to time.

#### ***Risks Related to the Manufacture of Our Products***

##### **The failure to secure an adequate supply of the inhalers to be used in the administration of Aridol and Bronchitol could compromise the commercialization of Aridol and Bronchitol.**

Both Aridol and Bronchitol are administered through a dry powder inhaler. If we are not able to enter into a supply agreement or the number of inhalers supplied are inadequate to meet patient demand, we would be subject to costly delays which may compromise the commercialization of Aridol and/or Bronchitol. The supply of inhalers may be delayed or we may need to change our supplier of inhalers which could delay the commercialization of Aridol and/or Bronchitol.

##### **Delays in the supply of the necessary quantity or quality of mannitol could compromise the commercialization of our products.**

Any delays in the supply of the necessary quantity or quality of mannitol for the manufacture of Aridol and Bronchitol could compromise the commercialization of our products. The supply of mannitol may be delayed, or we may need to change our supplier of mannitol, which could potentially result in delays to the supply of the product to clinical trials and for sale while we sourced alternative suppliers of mannitol.

##### **We currently have limited manufacturing capacity and outsource some manufacturing for the clinical development and commercial production of our products, all of which puts us at risk of lengthy and costly delays of bringing our products to market.**

We currently operate manufacturing facilities in Sydney, Australia. Our manufacturing facilities are licensed by the Australian Therapeutic Goods Administration, or TGA, to manufacture Good Manufacturing Practice grade material for clinical trials. We have outsourced the manufacturing of Good Manufacturing Practice grade PXS64 for preclinical trials and clinical trials as our manufacturing facilities are not suitable for the production of PXS64.

We are required to obtain a license to commercially manufacture Good Manufacturing Practice grade Aridol for commercial sale. If we are not able to obtain a license to commercially manufacture Good Manufacturing Practice grade Aridol or if the license is subject to conditions or is revoked, the commercialization of Aridol and consequentially our other product candidates may be delayed or severely compromised and our results of operations may be harmed. We expect that it will be necessary to build or establish suitable additional or alternative manufacturing facilities as the commercial demand for our products increase. If we establish new manufacturing facilities, we will incur significant costs and undetermined risks associated with the funding and building of such facilities which may delay or severely compromise the commercialization of our products and our results and operations may be harmed. There is also a risk of delays to our research and clinical trial activities if we needed to change our existing outsourced manufacturers of PXS64.

**In circumstances where we seek to outsource the manufacture of certain products, there is no guarantee that we will be able to enter into any such arrangement on acceptable terms, if at all, and as a result we are at risk of lengthy and costly delays of bringing our products to market.**

In circumstances where we seek to outsource the manufacture of certain product candidates, such as PXS64, there is no guarantee that we will be able to enter into any such arrangement on acceptable terms, if at all. We may be required to enter into long-term manufacturing agreements that contain exclusivity provisions and/or substantial termination penalties. To date, the manufacturing agreements for the manufacture of PXS64 do not contain any such exclusivity provisions or termination penalties. In addition, contract manufacturers may have a limited number of facilities in which our products can be produced and any interruption of the operation of those facilities could result in the cancellation of shipments and loss of product, resulting in delays and additional costs.

We and our contract manufacturers are required to produce our clinical product and commercial product under FDA and E.U. current Good Manufacturing Practices in order to meet acceptable standards. If such standards change, our ability and the ability of contract manufacturers to produce our products when we require may be affected.

We have outsourced the manufacturing of Good Manufacturing Practice grade PXS64 for Phase I clinical trials as our manufacturing facilities are not suitable for the production of PXS64. Our existing manufacturers of PXS64 and any future contract manufacturers for PXS64 or any of our other product candidates which we seek to contract manufacture may not perform as agreed or may not remain in the contract manufacturing business for the time required to successfully produce, store and distribute our products. We, or our contract manufacturers, may also fail to achieve and maintain required production yields or manufacturing standards which could result in patient injury or death, product recalls or withdrawals, product shortages, delays or failures in product testing or delivery or other problems that could seriously harm our business. In addition, we are, and our contract manufacturers are, subject to ongoing inspections and regulation of regulatory authorities, including by the TGA and the FDA.

The ability to find an acceptable manufacturer or to change manufacturers may be difficult for a number of reasons, including that the number of potential manufacturers is limited and we may not be able to negotiate agreements with manufacturers on commercially reasonable terms, the complex nature of the manufacturing process of certain of our product candidates, such as PXS64, which may require a significant learning curve for the manufacturer, and the FDA must approve any replacement manufacturer prior to manufacturing, which requires new testing and compliance inspections.

If we were required and able to change manufacturers, the FDA would also require that we demonstrate structural and functional comparability between the same product manufactured by different organizations and may require comparability studies.

#### ***Risks Related to Marketing, Distribution and Sales***

**If we are unable to establish a sales and marketing force our business may be harmed.**

We currently have a limited number of sales and marketing staff and no distribution capabilities. Our goal is to build an integrated pharmaceutical business undertaking research and development, clinical trials, sales and marketing for certain of our product candidates. We are proposing to develop our sales and marketing capability for products which address highly concentrated markets served by specialist physicians. We intend to contract or partner with third parties in respect of sales and marketing of products where the markets are larger, more diverse or less accessible. For our early stage products or any new products, we may also form strategic alliances with third parties, which have established distribution systems and sales forces, in order to commercialize our products. Assuming Aridol is approved for commercial sale, we intend to market it directly in Australia and to use a combination of direct marketing to pulmonary specialists and third parties in Europe and later in the U.S.

We will need to incur significant additional expenses and commit significant additional management resources to establish a sales and marketing force. Although we have already begun to develop our sales and marketing capability, we may not be able to successfully establish these capabilities despite these additional expenditures. Even if we are successful in establishing a sales and marketing force, it may not be as effective as a third-party sales and marketing force. In circumstances where we elect to rely on third parties, we may receive less



revenue than if we sold such products directly. In addition, we may have little or no control over the sales efforts of those third parties and they may not perform as agreed. In the event we are unable to sell Aridol, Bronchitol and other product candidates, either directly or through third parties, our business may be significantly harmed and we may be forced to delay, reduce the scope or terminate our clinical trials and the development, manufacturing and marketing of our products.

**Our failure to establish and manage a distribution network for our products could delay or compromise the commercialization of Aridol and Bronchitol and other products.**

We have not yet established systems and processes necessary for distributing products to customers. Failure to secure contracts with distributors could negatively impact the distribution of our products. If we are unable to effectively establish and manage the distribution process, the commercialization of Aridol and Bronchitol and other products may be delayed or severely compromised and our results of operations may be harmed.

**To the extent we are able to enter into collaborative arrangements or strategic alliances, we will be exposed to risks related to those collaborations and alliances.**

Although our goal is to be a fully integrated pharmaceutical company, an important element of our strategy for developing, manufacturing and commercializing our product candidates is entering into partnerships and strategic alliances with other pharmaceutical companies or other industry participants to advance our programs and enable us to maintain our financial and operational capacity. We may not be able to negotiate alliances on acceptable terms, if at all. Although we are not currently party to any collaborative arrangement or strategic alliance that is material to our business, in the future we may rely on collaborative arrangements or strategic alliances to complete the development and commercialization of some of our product candidates. These arrangements may result in us receiving less revenue than if we sold such products directly, may place the development, sales and marketing of our products outside our control, may require us to relinquish important rights or may otherwise be on terms unfavorable to us. Collaborative arrangements or strategic alliances will subject us to a number of risks, including the risk that:

- we may not be able to control the amount and timing of resources that our strategic partner/collaborators may devote to the product candidates;
- our strategic partner/collaborators may experience financial difficulties;
- we may be required to relinquish important rights such as marketing and distribution rights;
- business combinations or significant changes in a collaborator's business strategy may also adversely affect a collaborator's willingness or ability to complete its obligations under any arrangement;
- a collaborator could independently move forward with a competing product developed either independently or in collaboration with others, including our competitors; and
- collaborative arrangements are often terminated or allowed to expire, which would delay the development and may increase the cost of developing our product candidates.

**We face costs associated with importing our products into markets outside of Australia.**

As much of our product is likely to be manufactured in Australia, we may face difficulties in importing our products into various jurisdictions as a result of, among other things, import inspections, incomplete or inaccurate import documentation or defective packaging. There will be increased costs associated with importing/exporting our product.

***Risks Relating to Competition***

**If our competitors are able to develop and market products that are preferred over Aridol, Bronchitol or our other product candidates our commercial opportunity may be significantly reduced or eliminated.**

We face intense competition from established pharmaceutical and biotechnology companies, as well as from academic institutions, government agencies and private and public research institutions. We are seeking to develop and market products that will compete with other products and drugs that currently exist or are being developed or may be developed in the future. For Aridol, various products and treatments are currently marketed for

monitoring lung capacity and the identification and assessment of asthma, including methacholine (Provocholine<sup>®</sup>) by Methapharm Inc as a direct bronchiol provocation agent and nitric oxide (Niox<sup>®</sup>) by Aerocrine AB of Stockholm, Sweden. We believe Aridol is the only lung capacity test that is being developed using dry powder inhalation technology. This test may not be well accepted in the market place or the medical community. Similarly, for Bronchitol, various products and treatments are currently marketed, including inhaled antibiotics, mucolytic agents and bronchodilators. Bronchitol may not work well in conjunction with existing marketed therapies. In addition, a number of companies are developing new approaches for the treatment of cystic fibrosis, including new antibiotic preparations by Corus Inc. and Chiron Inc. and new agents to restore salt balance from Inspire Pharmaceuticals Inc. In addition, many companies are interested in gene therapy. New antibiotic preparations are being tested in patients with bronchiectasis. For patients with chronic bronchitis, new anti-inflammatory agents and new bronchodilating agents are under development.

Our commercial opportunity will be reduced or eliminated if our competitors develop and commercialize products that are safer, more effective, have fewer side effects, are more convenient, are less expensive, or that reach the market sooner than our products. Scientific, clinical or technical developments by our competitors may render Aridol and/or Bronchitol or our other product candidates obsolete or noncompetitive. Further, public announcements regarding the development of any such competing products could adversely affect the market price of our ordinary shares or ADSs. We anticipate that we will face increased competition in the future as new companies enter the markets and as scientific developments progress. If our products obtain regulatory approvals, but do not compete effectively in the marketplace, our business will suffer.

Many of our competitors currently have significantly greater financial resources and expertise in conducting clinical trials, obtaining regulatory approvals, undertaking and managing manufacturing and sales and marketing of products than we do. Other early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements they may have with large and established companies. In addition, these third parties compete with us in recruiting and retaining qualified scientific and management personnel, establishing clinical trial sites and patient registration for clinical trials, as well as in acquiring therapies and therapy licenses complementary to our programs or advantageous to our business. We expect that our ability to compete effectively will depend upon our ability to:

- successfully and rapidly complete clinical trials and obtain all requisite regulatory approvals in a cost-effective manner;
- attract and retain key personnel;
- demonstrate the competitive advantages of our product candidates. For example, Aridol must prove to be convenient and effective as a test for airway inflammation which identifies and determines the severity of asthma. In addition, Bronchitol must improve the quality of life and health of people with chronic obstructive lung diseases such as cystic fibrosis and bronchiectasis;
- build an adequate manufacturing, sales and marketing infrastructure. We face the risk that our future manufacturing, sales and marketing infrastructure may be inadequate for the commercialization of our products or that competitors have faster and more effective manufacturing, sales and marketing capacities;
- secure the support of key clinicians and physicians. The success of our products is dependant on the acceptance of our products by key clinicians and physicians. We face the risk that our products may not be well received or that a product will be released by a competitor which is preferred by key clinicians and physicians; and
- identify and obtain other product candidates on commercially reasonable terms which will provide us with a pipeline of potential product candidates which may reduce the risk if any of our existing product candidates or are adversely effected.

**Future sales of our products may suffer if they are not accepted in the marketplace by physicians, patients and the medical community.**

There is a risk that Aridol, Bronchitol or our other product candidates may not gain market acceptance among physicians, patients and the medical community. The degree of market acceptance of Aridol and Bronchitol or our other product candidates will depend on a number of factors. For example, Aridol must prove to be convenient and effective as a test for airway inflammation which identifies and determines the severity of asthma. In addition, Bronchitol must improve the quality of life for people with chronic obstructive lung diseases such as cystic fibrosis and bronchiectasis. The prevalence and severity of any side effects to Aridol or Bronchitol could negatively affect market acceptance of both Aridol and Bronchitol. Failure to achieve market acceptance of Aridol and Bronchitol would significantly harm our business.

The degree of market acceptance of any of our approved products will depend on a variety of factors, including:

- timing of market introduction and number and clinical profile of competitive products. There are currently a range of existing alternative products to each of our products and we are aware that new products are being developed;
- our ability to provide acceptable evidence of safety and efficacy and our ability to secure the support of key clinicians and physicians for our products;
- relative convenience and ease of administration. In the case of Aridol and Bronchitol, there is a risk that using dry powder inhalation technology may not be well accepted in the market place;
- cost-effectiveness compared to existing and new treatments;
- availability of coverage, reimbursement and adequate payment from health maintenance organizations and other third-party payors;
- prevalence and severity of adverse side effects; and
- other advantages over other treatment methods.

**If we are unable to obtain acceptable prices or adequate reimbursement from third-party payors for Aridol and Bronchitol, or any other product candidates that we may seek to commercialize, our revenues and prospects for profitability will suffer.**

The commercial success of our product candidates is substantially dependent on whether third-party coverage and reimbursement is available from governmental payors such as Medicare and Medicaid, private health insurers, including managed care organizations, and other third-party payors.

Many patients will not be capable of paying for our products themselves and will rely on third-party payors to pay for their medical needs. The U.S. Centers for Medicare and Medicaid Services, health maintenance organizations and other third-party payors in the United States, the European Union, Australia and other jurisdictions are increasingly attempting to contain healthcare costs by limiting both coverage and the level of reimbursement of new drugs and, as a result, they may not cover or provide adequate payment for our products. Our products may not be considered cost-effective and reimbursement may not be available to consumers or may not be sufficient to allow our products to be marketed on a competitive basis. Large private payors, managed care organizations, group purchasing organizations and similar organizations are exerting increasing influence on decisions regarding the use of, and reimbursement levels for, particular treatments. Such third-party payors, including Medicare, are challenging the prices charged for medical products and services, and many third-party payors limit or delay reimbursement for newly approved health care products. In particular, third-party payors may limit the reimbursed indications. Cost-control initiatives could decrease the price we might establish for products, which could result in product revenues lower than anticipated. If the prices for our product candidates decrease or if governmental and other third-party payors do not provide adequate coverage and reimbursement levels, our prospects for revenue and for profitability will suffer.

**If there are fewer individuals in our target markets than we estimate, we may not generate sufficient revenues to continue development of our product candidates or continue operations.**

Our estimate of the patient population of our target markets is based on published studies as well as internal analyses and studies we have commissioned. If the results of these studies or our analysis of them do not accurately reflect the number of patients in our target markets, our assessment of the market may be wrong, making it difficult or impossible for us to meet our revenue goals. In addition, it is difficult to determine the portion of the patient population that might use Aridol and/or Bronchitol, or our other product candidates.

**Our orphan drug exclusivity for Bronchitol may not provide us with a competitive advantage.**

The FDA has granted Orphan Drug designation to Bronchitol for the treatment of both bronchiectasis and CF for patients at risk for developing bronchiectasis. Orphan drug exclusivity for Bronchitol for the treatment of both bronchiectasis and cystic fibrosis for patients at risk of developing bronchiectasis is an important element of our competitive strategy. Any company that obtains the first FDA approval for a designated orphan drug for a rare disease generally receives marketing exclusivity for use of that drug for the designated condition for a period of seven years from approval. However, the FDA may permit other companies to market a form of mannitol, the active ingredient in Bronchitol, not covered by our patent, to treat bronchiectasis and cystic fibrosis for patients at risk of developing bronchiectasis if any such product demonstrates clinical superiority, or if we are unable to provide sufficient drug supply to meet medical needs. More than one product may also be approved by the FDA for the same orphan indication or disease as long as the products are different drugs. Any of these FDA actions could create a more competitive market for us. Our orphan drug exclusivity for Bronchitol does not apply to drugs to treat bronchiectasis and cystic fibrosis for patients at risk of developing bronchiectasis that do not contain mannitol, or to drugs containing mannitol that seek approval for uses other than bronchiectasis and cystic fibrosis for patients at risk of developing bronchiectasis. Our orphan drug exclusivity may thus not ultimately provide us a true competitive advantage, and our business could suffer as a result.

***Risks Relating to Regulatory Issues***

**Our products are subject to extensive regulation, which can be costly and time-consuming, and we may not obtain approvals for the commercialization of some or all of our products.**

The clinical development, manufacturing, sales and marketing of our products are subject to extensive regulation by regulatory authorities in the United States, the European Union, Australia and elsewhere. These regulations vary in important, meaningful ways from country to country.

We are not permitted to market a potential drug in the United States until we receive approval of a New Drug Application, or NDA, from the FDA. We have not yet received an NDA approval from the FDA for any of our products. Obtaining an NDA approval is expensive and is a complex, lengthy and uncertain process. The FDA approval process for a new drug involves completion of preclinical studies and the submission of the results of these studies to the FDA, together with proposed clinical protocols, manufacturing information, analytical data and other information in an IND, which must become effective before human clinical trials may begin. Clinical development typically involves three phases of study: Phase I, II and III. The most significant costs associated with clinical development are the Phase III clinical trials as they tend to be the longest and largest studies conducted during the drug development process. After completion of clinical trials, an NDA may be submitted to the FDA. In responding to an NDA, the FDA may refuse to file the application, or if accepted for filing, the FDA may grant marketing approval, request additional information or deny the application if it determines that the application does not provide an adequate basis for approval. In addition, failure to comply with FDA and other applicable foreign and U.S. regulatory requirements may subject us to administrative or judicially imposed sanctions. We have recently completed the Phase III clinical trials of Aridol necessary for Australian and European registration of Aridol. Further Phase III clinical trials are required to be completed in the U.S. before an application for marketing authority can be filed with the FDA. The FDA has accepted an IND for Aridol and for Bronchitol. Clinical trials of our other product candidates, including Bronchitol and PXS64, will continue for several years, but may take significantly longer to complete. We have completed a Phase II clinical trial of Bronchitol for the treatment of bronchiectasis and expect to commence additional clinical trials during the fourth quarter of 2005 or the first quarter of 2006. We have also completed a Phase II clinical trial of Bronchitol for the treatment of cystic fibrosis and will prepare additional

clinical trials. Our other product candidates, PXS64 and PXS2076, are currently in varying stages of the research or preclinical phase of development.

Despite the substantial time and expense invested in preparation and submission of an NDA or equivalents in other jurisdictions, regulatory approval is never guaranteed. The FDA and other regulatory authorities in the United States, the European Union, Australia and elsewhere, exercise substantial discretion in the drug approval process. The number, size and design of preclinical studies and clinical trials that will be required will vary depending on the product, the disease or condition for which the product is intended to be used and the regulations and guidance documents applicable to any particular product. The FDA or other regulators can delay, limit or deny approval of a product for many reasons, including, but not limited to, the fact that regulators may not approve our or our third-party manufacturer's processes or facilities or that new laws may be enacted or regulators may change their approval policies or adopt new regulations requiring new or different evidence of safety and efficacy for the intended use of a product.

**Even if our product candidates receive regulatory approval, we may still face development and regulatory difficulties that may delay or impair future sales of our products and we would be subject to ongoing regulatory obligations and restrictions, which may result in significant expense and limit our ability to commercialize our product candidates.**

If we receive regulatory approval to sell Aridol or any other product candidate, the relevant regulatory authorities may, nevertheless, impose significant restrictions on the indicated uses, manufacturing, labeling, packaging, storage, advertising, promotion and record keeping or impose ongoing requirements for post-approval studies and adverse event reporting. In addition, regulatory agencies subject a marketed product, its manufacturer and the manufacturer's facilities to continual review and periodic inspections. Potentially costly follow-up or post-marketing clinical studies may be required as a condition of approval to further substantiate safety or efficacy, or to investigate specific issues of interest to the regulatory authority. Previously unknown problems with the product candidate, including adverse events of unanticipated severity or frequency, may result in restrictions on the marketing of the product, and could include withdrawal of the product from the market. If we discover previously unknown problems with a product or our manufacturing facilities or the manufacturing facilities of a contract manufacturer, a regulatory agency may impose restrictions on that product, on us or on our third-party contract manufacturers, including requiring us to withdraw the product from the market. If we fail to comply with applicable regulatory requirements, a regulatory agency may:

- issue warning letters;
- impose civil or criminal penalties;
- suspend our regulatory approval;
- suspend any of our ongoing clinical trials;
- refuse to approve pending applications or supplements to approved applications filed by us;
- impose restrictions on our operations, including closing our contract manufacturers' facilities or terminating licenses to manufacture Good Manufacturing Practice grade material; or
- seize or detain products or require a product recall.

Any of the foregoing could seriously harm the commercialization of our products and our results and operations may be seriously harmed. Likewise, any failure to comply with ongoing regulatory requirements may significantly and adversely affect our ability to commercialize our product.

In addition, the law or regulatory policies governing pharmaceuticals may change. New statutory requirements may be enacted or additional regulations may be enacted that could prevent or delay regulatory approval of our products. We cannot predict the likelihood, nature or extent of adverse government regulation that may arise from future legislation or administrative action. If we are not able to maintain regulatory compliance, we might not be permitted to market our products and our business could suffer.

### ***Risks Relating to Product Liability Claims***

**If product liability lawsuits are successfully brought against us, we will incur substantial liabilities and damage to our reputation and may be required to limit commercialization of Aridol and Bronchitol or other product candidates.**

We face product liability exposure related to the testing of our product candidates in human clinical trials. We are currently conducting clinical trials in respect of Aridol and Bronchitol and we expect to commence clinical trials of PXS25. If any of our products are approved for sale, we may face exposure to claims by an even greater number of persons than were involved in the clinical trials once we begin marketing, distribution and sales our products commercially. We expect to commence commercial sales of Aridol in Australia and Europe in the course of 2006.

Regardless of merit or eventual outcome, liability claims may result in:

- decreased demand for our products and product candidates;
- injury to our reputation;
- withdrawal of clinical trial participants;
- costs of related litigation;
- substantial monetary awards to patients and others;
- loss of revenues; and
- the inability to commercialize our products and product candidates.

We enter into indemnity agreements with the hospitals, institutions and authorities in which the clinical trials are conducted and with the clinicians and investigators who carry out the clinical trials on our behalf. The majority of the indemnities are in a substantially similar form and are based on an Australian industry standard agreement. Certain of the agreements have been negotiated on a case by case basis and vary from the standard. The standard indemnities typically provide that we will indemnify in respect of all claims and proceedings made by any of the patients or non-patient volunteers participating in the relevant clinical trials for personal injury arising from the administration of the product under investigation or any clinical intervention or procedure required.

We have liability insurance that covers our clinical trials in North America and in other countries where we conduct clinical trials. Having regard to the insurance requirements of the ethics committees who oversee the clinical trials, the good safety profile of Aridol and Bronchitol, the varied use of mannitol in humans, the number of clinical trials undertaken to date without a material claim being made against us, the use of informed consents by trial participants, and the number of clinical trials being undertaken at any one time, we consider that our clinical trial insurance is reasonable for our current activities. We intend to expand our insurance coverage to include the sale of commercial products if marketing approval is obtained for Aridol and in the future for other product candidates. However, insurance coverage is increasingly expensive. We may not be able to maintain insurance coverage at a reasonable cost and we may not be able to obtain insurance coverage that we consider reasonable or that will be adequate to satisfy any liability that may arise and the claim for damages could be substantial. If we are not able to obtain adequate coverage at a reasonable cost, the commercialization of our products may be delayed or severely compromised.

If there is a claim made against us or some other problem that is attributable to our products or product candidates, our ordinary share and ADS prices may be negatively affected. Even if we were ultimately successful in product liability litigation, the litigation would consume substantial amounts of our financial and managerial resources and may create adverse publicity, all of which would impair our ability to generate sales of the product the subject of the litigation as well as our other potential products.

### ***Risks Relating to Intellectual Property and License Arrangements***

**The development programs for our two lead product candidates are based in part on intellectual property rights we license from others, and any termination of those licenses could seriously harm our business as the loss of any rights to market key products would seriously harm our operating results.**

We have an exclusive worldwide license from Central Sydney Area Health Service to develop and commercialize certain intellectual property relating to the use of mannitol, the component part of both Aridol and Bronchitol, to induce sputum and promote airway clearance and also in the use as a test of airway function and susceptibility to asthma. We also have an exclusive license from ANU Enterprises Pty Ltd to develop and commercialize intellectual property relating to the treatment of inflammatory or immune-mediated conditions in patients by administering a phosphosugar. Both of these license agreements impose payment and other material obligations on us. If we were to breach any such obligations and our counterparties were entitled to, and did, terminate the licenses, this would restrict or delay or eliminate our ability to develop and commercialize our key product candidates, which could seriously harm our business. In particular, if our agreement with Central Sydney Area Health Service were terminated, then we would have no further rights to develop and commercialize Aridol and Bronchitol which would seriously harm our business as this license is critical to both Aridol and Bronchitol.

**Third parties may own or control patents or patent applications that we may be required to license to commercialize our product candidates, that we may infringe, or that could result in litigation that would be costly and time consuming.**

Our ability to commercialize Aridol and Bronchitol and our other product candidates depends upon our ability to develop, manufacture, market and sell these products without infringing the proprietary rights of third parties. A number of pharmaceutical and biotechnology companies, universities and research institutions have or may be granted patents that cover technologies similar to the technologies owned by or licensed to us. We may choose to seek, or be required to seek, licenses under third-party patents, which would likely require the payment of license fees or royalties or both. A license may not be available to us on commercially reasonable terms, or at all. We may also be unaware of existing patents or other proprietary rights of third parties that may be infringed by Aridol and Bronchitol or our other product candidates. As patent applications can take many years to issue, there may be other currently pending applications which may later result in issued patents that are infringed by Aridol and Bronchitol or our other product candidates.

There has been substantial litigation and other proceedings regarding patent and other intellectual property rights in the pharmaceutical and biotechnology industries. Defending ourselves against third-party claims, including litigation in particular, would be costly and time consuming and would divert management's attention from our business, which could lead to delays in our development or commercialization efforts. If third parties are successful in their claims, we might have to pay substantial damages or take other actions that are adverse to our business. As a result of intellectual property infringement claims, or to avoid potential claims, we might be:

- prohibited from selling or licensing any product candidate that we may develop unless the patent holder licenses the patent to us, which it is not required to do;
- required to expend considerable amounts of money in defending the claim;
- required to pay substantial royalties or grant a cross license to our patents to another patent holder;
- required to pay substantial monetary damages; or
- required to redesign the formulation of a product so it does not infringe, which may not be possible or could require substantial funds and time.

We may also be forced to bring an infringement action if we believe that a third party is infringing our protected intellectual property. Any such litigation will be costly, time consuming and divert management's attention, and the outcome of any such litigation may not be favorable to us.

**Our intellectual property rights may not preclude competitors from developing competing products and our business may suffer.**

If we are not able to protect our proprietary technology, trade secrets and know-how, our competitors may use our intellectual property to develop competing products. Our patents, including our licensed patents relating to the use and manufacture of Aridol and Bronchitol, may not be sufficient to prevent others from competing with us. Most of our patents covering Aridol and Bronchitol expire in 2015. Therefore, we will not be able to depend on these patents past these relevant dates to exclude competitors from developing generic versions of Aridol and Bronchitol. Our issued patents and those that we may issue in the future, or those licensed to us, may be challenged, invalidated or circumvented, which could limit our ability to stop competitors from marketing related products or the term of patent protection that we may have for our product candidates. The occurrence of any of the foregoing events could harm our competitive position and seriously harm our business.

Our trade secrets relating to our product candidates and the manufacture of our product candidates may become known or independently discovered or competitors may develop alternatives. We disclose confidential information and trade secrets from time to time provided that the recipient executes a non-disclosure agreement or otherwise owes us obligations of confidentiality. Confidentiality agreements may be breached and we may have no effective remedy for such a breach. Enforcing a claim that a third party illegally obtained and is using our trade secrets is expensive and time consuming, and the outcome is unpredictable. Failure to obtain or maintain confidential information and trade secret protection could adversely affect our competitive business position.

**If we fail to enforce adequately or defend our intellectual property rights our business may be harmed.**

Our commercial success depends, to a large extent, on obtaining and maintaining patent and trade secret protection for our products, the methods used to manufacture those products and the methods for treating patients using those products. A key tool in protecting our products and our technologies from unauthorized use by third parties is the extent that valid and enforceable patents or trade secrets cover them. Our ability to obtain patents is uncertain and there is a risk that we may not be able to secure and maintain patents which we require to defend our intellectual property position. Patents provide only limited protections and may not adequately protect our rights or permit us to gain or keep any competitive advantage. Some countries in which we may sell our product candidates or license our intellectual property may fail to protect our intellectual property rights to the same extent as the protection that may be afforded in the United States or Australia. Some legal principles remain unresolved and there has not been a consistent policy regarding the breadth or interpretation of claims allowed in patents in the United States, the European Union, Australia or elsewhere. In addition, the specific content of patents and patent applications that are necessary to support and interpret patent claims is highly uncertain due to the complex nature of the relevant legal, scientific and factual issues. Changes in either patent laws or in interpretations of patent laws in the United States, the European Union or elsewhere may diminish the value of our intellectual property or narrow the scope of our patent protection.

Even if patents are issued, those patents can be challenged by our competitors who can argue such patents are invalid. Patents also will not protect our products if competitors devise ways of making these product candidates without legally infringing our patents. The U.S. Federal Food, Drug and Cosmetic Act and FDA regulations and policies and equivalents in other jurisdictions provide incentives to manufacturers to challenge patent validity or create modified, noninfringing versions of a drug in order to facilitate the approval of abbreviated new drug applications for generic substitutes.

Proprietary trade secrets and unpatented know-how are also very important to our business. We rely on trade secrets to protect our technology, especially where we do not believe that patent protection is appropriate or obtainable. However, trade secrets are difficult to protect. Our employees, consultants, contractors, outside scientific collaborators and other advisors may unintentionally or willfully disclose our confidential information to competitors, and confidentiality agreements may not provide an adequate remedy in the event of unauthorized disclosure of confidential information. Enforcing a claim that a third party illegally obtained and is using our trade secrets is expensive and time consuming, and the outcome is unpredictable. Failure to obtain or maintain trade secret protection could adversely affect our competitive business position. To date, we are not aware of any unintentional or willful disclosure of any of our material confidential information or any unauthorized use of our confidential information and we have not been required to seek remedy for any such unauthorized disclosure or use.



### ***Risks Relating to Resources***

#### **If we fail to attract and keep senior management and key scientific personnel, we may be unable to successfully develop and commercialize our product candidates.**

Our success depends on our continued ability to attract, retain and motivate highly qualified management, clinical and scientific personnel, manufacturing personnel, sales and marketing personnel and on our ability to develop and maintain important relationships with clinicians, scientists and leading academic and health institutions, including in particular ANU Enterprises Pty Ltd, the Australian National University and the Central Sydney Area Health Service.

The loss of services of one or more of our members of key management could delay or compromise the successful completion of our clinical trials or the commercialization of Aridol and Bronchitol and our other product candidates. We enter into employment agreements with each of our employees, including each member of our key management. Each of our employees agree to a specific period of notice that they or we must give in order to terminate their employment. Employees can terminate their employment by giving between one to three months notice (as set out in the relevant employee's employment agreement).

In the near term we will need to attract and retain manufacturing personnel and sales and marketing personnel and effectively integrate them into our organization to coincide with the expected commencement of commercial sales of Aridol in Europe and Australia. If we fail to attract or effectively integrate new personnel and consultants into our organization and create effective working relationships among them and other members of management, the future development and commercialization of Aridol and our other product candidates may suffer, harming future regulatory approvals, sales of our products and our results of operations.

There is significant competition from other companies and research and academic institutions for qualified personnel in the areas of our activities. If we fail to identify, attract, retain and motivate these highly skilled personnel, we may be unable to continue our development and commercialization activities.

The addition of new employees and the loss of key employees, particularly in key positions, can be disruptive and may also cause the future development and commercialization of our product candidates to suffer, harming future regulatory approvals, sales of our products and our results of operations.

We do not currently carry "key person" insurance on the lives of members of senior management. We consider that at this stage of our development it is reasonable not to carry any key person insurance.

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

We are currently a small company with 43 employees and full time contractors as of September 30, 2005. In order to continue our clinical trials and commercialize our product candidates, manufacture commercial quantities of Aridol and market and sell Aridol, we will need to increase our operations, including expanding our employee base. Our future financial performance and our ability to commercialize our products and to compete effectively will depend, in part, on our ability to manage any future growth effectively. To that end, we must be able to:

- manage our preclinical studies and clinical trials effectively;
- undertake and manage the manufacturing of product effectively;
- undertake and manage sales and marketing effectively;
- integrate current and additional management, administrative, financial and sales and marketing personnel;
- develop our administrative, accounting and management information systems and controls; and
- hire and train additional qualified personnel.

**The acquisition or licensing of other products or product candidates may put a strain on our operations and will likely require us to seek additional financing.**

One of our strategies is to develop and license or acquire complementary products or product candidates. We have no present agreement regarding any new material licensing or acquisitions. However, if we do undertake any licensing or acquisitions, the process of undertaking the licensing or acquisitions and integrating a licensed or acquired product or product candidate into our business may put a strain on our operations, including diversion of personnel and financial resources and diversion of management's attention. In addition, any acquisition would give rise to potentially significant additional operating costs which would likely require us to seek additional financing. Future acquisitions could result in additional issuances of equity securities that would dilute the ownership of existing shareholders and holders of our ADSs. Future acquisitions could also result in us incurring debt, contingent liabilities or the amortization of expenses related to other intangible assets, any of which could adversely affect our operating results.

***Risks Relating to Takeovers***

**Our constitution may discourage attempts by shareholders to make a proportional takeover for us and could restrict the ability for shareholders to obtain a premium from such a transaction.**

Provisions of our constitution contain a proportional takeover provision which provides that if a person makes a proportional takeover offer for less than all of the share capital in us, shareholders are entitled to vote to determine whether the proportional takeover offer may proceed. A person may wish to make a proportional takeover offer for a number of reasons, including, if they wish to increase their control of us and/or influence the composition of the board of directors. Arguably, the proportional takeover provisions in our constitution make it more difficult to achieve a proportional takeover and therefore may discourage proportional takeover offers and make it more difficult for a person to gain proportional control of us and could restrict the ability for shareholders to obtain a premium from such a transaction.

**Australian takeovers laws may discourage takeover offers being made for us or may discourage the acquisition of large numbers of our shares.**

We are incorporated in Australia and are subject to the takeovers laws of Australia. Among other things, we are subject to the Corporations Act 2001 (Commonwealth of Australia). Subject to a range of exceptions, the Corporations Act prohibits the acquisition of a direct or indirect interest in our issued voting shares (including through the acquisition of ADSs) if the acquisition of that interest will lead to a person's or someone else's voting power in us increasing from 20% or below to more than 20%, or increasing from a starting point that is above 20% and below 90%. Exceptions to the general prohibition include circumstances where the person makes a formal takeover bid for us, if the person obtains shareholder approval for the acquisition or if the person acquires less than 3% of the voting power of us in any rolling six month period. Australian takeovers laws may discourage takeover offers being made for us or may discourage the acquisition of large numbers of our shares. This may have the ancillary effect of entrenching our board of directors and may deprive or limit strategic opportunities of our shareholders and ADS holders to sell their shares and may restrict the ability of our shareholders and ADS holders to obtain a premium from such transactions. See Item 10B - "Change of Control."

***Risks Related to our ADSs or Ordinary Shares***

**The price of our ordinary shares is highly volatile and could decline significantly.**

The market price of our ordinary shares historically has been, and we expect will continue to be, subject to significant fluctuations over short periods of time. These fluctuations may be due to factors specific to us, to changes in analysts' recommendations and earnings estimates, to arbitrage between our Australian quoted shares and our ADSs, to changes in exchange rates, or to factors affecting the biopharmaceutical industry or the securities markets in general. For example, from our initial quotation on the Australian Stock Exchange on November 10, 2003 until December 9, 2005, the closing price per share of our ordinary shares ranged from a low of A\$0.34 on November 27, 2003 to a high of A\$3.12 on October 4, 2005. We may experience a material decline in the market price of our shares, regardless of our operating performance. Therefore, a holder of our ordinary shares or ADSs may not be able to sell those ordinary shares or ADSs at or above the price paid by such holder for such shares or ADSs.

declines in our ordinary shares or ADSs could result from a variety of factors, including many outside our control. These factors include:

- adverse or inconclusive results or delays in our clinical trial programs;
- unforeseen safety issues or adverse side effects resulting from the clinical trials or the commercial use of any of our products;
- regulatory actions in respect of any of our products or the products of any of our competitors;
- failure of any of our product candidates, such as Aridol, if approved, to achieve commercial success;
- announcements of the introduction of new products by us or our competitors;
- market conditions, including market conditions in the pharmaceutical and biotechnology sectors;
- increases in our costs or decreases in our revenues due to unfavorable movements in foreign currency exchange rates;
- developments or litigation concerning patents, licenses and other intellectual property rights;
- litigation or public concern about the safety of our potential products;
- changes in recommendations or earnings estimates by securities analysts;
- actual and anticipated fluctuations in our quarterly operating results;
- deviations in our operating results from the estimates of securities analysts;
- rumors relating to us or our competitors;
- additions or departures of key personnel;
- changes in third-party reimbursement policies; and
- developments concerning current or future strategic alliances or acquisitions.

Class action litigation has been brought in the past against companies which have experienced volatility in the market price of their securities. We may become involved in this type of litigation in the future. Litigation of this type is often extremely expensive and diverts management's attention and company's resources.

In addition, since the initial listing of our ADSs on Nasdaq National Market trading volume in our ADSs has been limited and a significant portion of the ownership of our ADSs is concentrated with a small number of holders. The limited trading volume may adversely affect the prices at which the ADSs may be bought or sold. There can be no assurance that a more active trading market in our ADSs will develop in the U.S.

**Rights as a holder of ordinary shares are governed by Australian law and our Constitution and differ from the rights of shareholders under U.S. law. Holders of our ordinary shares or ADSs may have difficulty in effecting service of process in the United States or enforcing judgments obtained in the United States.**

Pharmaxis Ltd is a public company incorporated under the laws of Australia. Therefore, the rights of holders of our ordinary shares are governed by Australian law and our Constitution. These rights differ from the typical rights of shareholders in U.S. corporations. The rights of holders of ADSs are affected by Australian law and our Constitution but are governed by U.S. law. Circumstances that under U.S. law may entitle a shareholder in a U.S. company to claim damages may also give rise to a cause of action under Australian law entitling a shareholder in an Australian company to claim damages. However, this will not always be the case.

Holders of our ordinary shares or ADSs may have difficulties enforcing, in actions brought in courts in jurisdictions located outside the U.S., liabilities under U.S. securities laws. In particular, if such a holder sought to bring proceedings in Australia based on U.S. securities laws, the Australian court might consider:

- that it did not have jurisdiction; and/or

- that it was not an appropriate forum for such proceedings; and/or
- that, applying Australian conflict of laws rule, U.S. law (including U.S. securities laws) did not apply to the relationship between holders of our ordinary shares or ADSs and us or our directors and officers; and/or
- that the U.S. securities laws were of a public or penal nature and should not be enforced by the Australian court.

All of our directors and executive officers are residents of countries other than the U.S. Furthermore, all or a substantial portion of their assets and our assets are located outside the U.S. As a result, it may not be possible for a holder of our ordinary shares or ADSs to:

- effect service of process within the U.S. upon any of our directors and executive officers or on us;
- enforce in U.S. courts judgments obtained against any of our directors and executive officers or us in the U.S. courts in any action, including actions under the civil liability provisions of U.S. securities laws;
- enforce in U.S. courts judgments obtained against any of our directors and senior management or us in courts of jurisdictions outside the United States in any action, including actions under the civil liability provisions of U.S. securities laws; or
- bring an original action in an Australian court to enforce liabilities against any of our directors and executive officers or us based upon U.S. securities laws.

Holders of our ordinary shares and ADSs may also have difficulties enforcing in courts outside the U.S. judgments obtained in the U.S. courts against any of our directors and executive officers or us, including actions under the civil liability provisions of the U.S. securities laws.

**Shares or ADSs eligible for public sale could adversely affect the price of our ordinary shares.**

The market price for our shares or ADSs could decline as a result of sales by our existing shareholders or management of ordinary shares or the perceptions that these sales could occur. These sales may also make it difficult for us to sell equity securities in the future at a time and at a price when we deem appropriate.

**Currency fluctuations may adversely affect the price of the ADSs relative to the price of our ordinary shares.**

The price of our ordinary shares is quoted in Australian dollars and the price of our ADSs is quoted in U.S. dollars. Movements in the Australian dollar/U.S. dollar exchange rate may adversely affect the U.S. dollar price of our ADSs and the U.S. dollar equivalent of the price of our ordinary shares. In the last two years, the Australian dollar has as a general trend appreciated against the U.S. dollar. Any continuation of this trend may positively affect the U.S. dollar price of our ADSs and the U.S. dollar equivalent of the price of our ordinary shares, even if the price of our ordinary shares in Australian dollars increases or remains unchanged. However, this trend may not continue and may be reversed. If the Australian dollar weakens against the U.S. dollar, the U.S. dollar price of the ADSs could decline, even if the price of our ordinary shares in Australian dollars increases or remains unchanged. If dividends are payable, we will likely calculate and pay any cash dividends in Australian dollars and, as a result, exchange rate movements will affect the U.S. dollar amount of any dividends holders of our ADSs will receive from the depository (as defined below in “Description of American Depositary Shares”). As previously noted, in the last two years, the Australian dollar has as a general trend appreciated against the U.S. dollar. Any continuation of this trend may positively affect the U.S. dollar value of our dividends. However, this trend may not continue and may be reversed. A reversal of this trend may result in a decrease in the U.S. dollar value of our dividends. The potential impact that currency fluctuations may have on our earnings is set out in the risk factor “Risks Related to Our Business – Currency fluctuations may expose us to increased costs and revenue decreases.”

**We may become a passive foreign investment company, or a PFIC, for U.S. federal income tax purposes, which could result in negative tax consequences to the holders of our ordinary shares or ADSs.**

Based on an analysis of our assets and gross income, we do not believe that we will be a PFIC for our current tax year or that we were a PFIC for our tax year ended June 30, 2005. However, our analysis indicates that we may have been a PFIC for our tax year ended June 30, 2004. We have not conducted a PFIC analysis for any tax year prior to our tax year ended June 30, 2004. The determination of whether we are, or at any time in the past have been, a PFIC is made annually on a taxable year basis and depends on factors such as the composition of our income and the value of our assets. Therefore, it is possible that we could be classified as a PFIC for our current taxable year and any future taxable year. If we are classified as a PFIC in any taxable year that a “U.S. Holder” (as defined in the section entitled “Taxation”) owns our ordinary shares or ADSs, we generally will continue to be treated as a PFIC for that U.S. Holder in all succeeding years. Such U.S. Holder would be subject to additional taxes on any “excess distributions” received from us and any gain realized from the sale or other disposition of our ordinary shares or ADSs. We urge U.S. investors to consult their own tax advisors about the application of the PFIC rules and certain elections that may help to minimize adverse U.S. federal income tax consequences in their particular circumstances. For a further discussion of the U.S. federal income tax consequences of investing in a PFIC, see the discussion below under the section entitled Item 10D - “Taxation.”

**We have never paid a dividend and we do not intend to pay dividends in the foreseeable future which means that holders of shares and ADSs may not receive any return on their investment from dividends.**

To date, we have not declared or paid any cash dividends on our ordinary shares and currently intend to retain any future earnings for funding growth. We do not anticipate paying any dividends in the foreseeable future. Dividends may only be paid out of our profits. Payment of cash dividends, if any, in the future will be at the discretion of our board of directors or, if our directors do not exercise their power to issue dividends, our shareholders in a general meeting may. Our holders of shares and ADSs may not receive any return on their investment from dividends.

**Our ADR holders are not shareholders and do not have shareholder rights.**

The Bank of New York, as depositary, executes and delivers our American Depositary Receipts, or ADRs, on our behalf. Each ADR is a certificate evidencing a specific number of American Depositary Shares, also referred to as ADSs. Our ADR holders will *not* be treated as shareholders and do not have the rights of shareholders. The depositary will be the holder of the shares underlying our ADRs. Holders of our ADRs will have ADR holder rights. A deposit agreement among us, the depositary and our ADR holders, and the beneficial owners of ADRs, sets out ADR holder rights as well as the rights and obligations of the depositary. New York law governs the deposit agreement and the ADRs.

Our ADR holders do not have the same voting rights as our shareholders. Shareholders are entitled to our notices of general meetings and to attend and vote at our general meetings of shareholders. At a general meeting, every shareholder present (in person or by proxy, attorney or representative) and entitled to vote has one vote on a show of hands. Every shareholder present (in person or by proxy, attorney or representative) and entitled to vote has one vote per fully paid ordinary share on a poll. This is subject to any other rights or restrictions which may be attached to any shares. Our ADR holders may instruct the depositary to vote the ordinary shares underlying their ADRs, but only if we ask the depositary to ask for their instructions. If we do not ask the depositary to ask for the instructions, our ADR holders are not entitled to receive our notices of general meeting or instruct the depositary how to vote. Our ADR holders will not be entitled to attend and vote at a general meeting unless they withdraw the ordinary shares. However, our ADR holders may not know about the meeting enough in advance to withdraw the shares. If we ask for our ADR holders’ instructions, the depositary will notify our ADR holders of the upcoming vote and arrange to deliver our voting materials and form of notice to them. The depositary will try, as far as practical, subject to Australian law and the provisions of the depositary agreement, to vote the shares as our ADR holders instruct. The depositary will not vote or attempt to exercise the right to vote other than in accordance with the instructions of the ADR holders. We cannot assure our ADR holders that they will receive the voting materials in time to ensure that they can instruct the depositary to vote their shares. In addition, there may be other circumstances in which our ADR holders may not be able to exercise voting rights.

Our ADR holders do not have the same rights to receive dividends or other distributions as our shareholders. Subject to any special rights or restrictions attached to a share, the directors may determine that a dividend will be payable on a share and fix the amount, the time for payment and the method for payment (although we have never declared or paid any cash dividends on our ordinary stock and we do not anticipate paying any cash dividends in the foreseeable future). Dividends may be paid on shares of one class but not another and at different rates for different classes. Dividends and other distributions payable to our shareholders with respect to our ordinary shares generally will be payable directly to them. Any dividends or distributions payable with respect to ordinary shares will be paid to the depositary, which has agreed to pay to our ADR holders the cash dividends or other distributions it or the custodian receives on shares or other deposited securities, after deducting its fees and expenses. Our ADR holders will receive these distributions in proportion to the number of shares their ADSs represent. In addition, there may be certain circumstances in which the depositary may not pay to our ADR holders amounts distributed by us as a dividend or distribution.

**There are circumstances where it may be unlawful or impractical to make distributions to the holders of our ADRs.**

The deposit agreement with the depositary allows the depositary to distribute the foreign currency only to those ADR holders to whom it is possible to do so. If a distribution is payable by us in Australian dollars, the depositary will hold the foreign currency it cannot convert for the account of the ADR holders who have not been paid. It will not invest the foreign currency and it will not be liable for any interest. If the exchange rates fluctuate during a time when the depositary cannot convert the foreign currency, our ADR holders may lose some of the value of the distribution.

The depositary is not responsible if it decides that it is unlawful or impractical to make a distribution available to any ADR holders. This means that our ADR holders may not receive the distributions we make on our shares or any value for them if it is illegal or impractical for us or the depositary to make them available to them.

**ITEM 4. INFORMATION ON THE COMPANY**

**A. History and Development of the Company**

Pharmaxis Ltd is a public company limited by shares, is domiciled in Australia and operates under, and is subject to, the Corporations Act 2001 (Commonwealth of Australia).

The Company was incorporated on May 29, 1998 in the Australian Capital Territory, Australia under the name “Praxis Pharmaceuticals Australia Pty Ltd.” On June 6, 2002, the Company changed its name from “Praxis Pharmaceuticals Australia Pty Ltd” to “Pharmaxis Pty Ltd.” On September 5, 2003, the Company changed its name from “Pharmaxis Pty Ltd” to “Pharmaxis Ltd” to reflect the change of company type from a proprietary company limited by shares to a public company limited by shares undertaken at that time. The Company’s ordinary shares are quoted on the Australian Stock Exchange Ltd and our American Depositary Shares (“ADS”) are quoted on the Nasdaq National Market. Each ADS represents 15 ordinary shares. In November, 2005 we completed a public offering of 1.3 million ADSs and a placement of 19.9 million ordinary shares and received US\$ 58.5 million therefrom after payment of underwriting fees and offering expenses. A total of 176,000 additional ADSs were sold by selling shareholders pursuant to partial exercise of the overallotment option by the underwriter. The Company received no proceeds from these shares.

Our principal place of business is Unit 2, 10 Rodborough Road, Frenchs Forest, NSW 2086, Australia, and the Company’s telephone number is 011 61 2 9454 7200.

**B. Business Overview**

We are an Australian specialty pharmaceutical company focused on the development of new products for the diagnosis and treatment of chronic respiratory and inflammatory/autoimmune diseases. We are developing Aridol, our lead product candidate, as a novel tool for the diagnosis of asthma and management of asthma and chronic obstructive pulmonary disease, or COPD. The Aridol test mimics the bronchoconstriction that occurs in

inflamed airways during asthma episodes and may be used to assist with the identification of people who have airway hyperresponsiveness, a hallmark of untreated or poorly controlled asthma. Aridol may also be used to determine the appropriate doses of anti-inflammatory medicine such as inhaled corticosteroids. We recently completed a pivotal Phase III clinical trial to determine the selectivity and specificity of Aridol as a test for the detection of airway inflammation in patients diagnosed with asthma. Based on results from this study, we have applied for marketing approval in Australia and the European Union, or the E.U., under the mutual recognition procedure. Later this year, we plan to initiate a pivotal Phase III trial of Aridol in the U.S. We are also developing Bronchitol, our proprietary inhaled dry powder mannitol formulation, for the treatment of cystic fibrosis, or CF, as well as COPD, an umbrella term for diseases such as bronchiectasis and chronic bronchitis. We recently completed a Phase II clinical trial of Bronchitol in patients with CF and demonstrated a statistically significant improvement in lung function relative to placebo over a two week treatment period. We have also completed a Phase II clinical trial of Bronchitol in bronchiectasis patients and demonstrated a clinically meaningful increase in patients' quality of life relative to placebo. We plan to conduct Phase III clinical trials of Bronchitol for the treatment of CF and for bronchiectasis. The U.S. Food and Drug Administration, or the FDA, has granted Orphan Drug designation to Bronchitol for the treatment of both bronchiectasis and CF for patients at risk for developing bronchiectasis. We believe that all CF patients are at risk for developing bronchiectasis. Our preclinical pipeline is focused on novel treatments for other inflammatory/autoimmune diseases, including rheumatoid arthritis and multiple sclerosis.

The following table summarizes our current product candidates. All time periods set forth in the table below refer to calendar years.

<u>Product Candidate / Indication</u>	<u>Development Status</u>	<u>Next Anticipated Milestones</u>
<b>Aridol</b>		
<i>Asthma: diagnosis and management</i>	<ul style="list-style-type: none"> <li>• Australian and E.U. marketing applications submitted</li> <li>• Commenced U.S. Phase III clinical trial</li> </ul>	<ul style="list-style-type: none"> <li>• Potential marketing authorization in Australia and Europe (1H06)</li> </ul>
<i>COPD: patient responsiveness to inhaled steroids</i>	<ul style="list-style-type: none"> <li>• Completed Phase IIa clinical trial (investigator-sponsored)</li> <li>• Commenced Phase II clinical trial</li> </ul>	<ul style="list-style-type: none"> <li>• Complete Phase II clinical trial (2H06)</li> </ul>
<b>Bronchitol</b>		
<i>Cystic fibrosis</i>	<ul style="list-style-type: none"> <li>• Phase II trial complete</li> <li>• Granted Orphan Drug designation by the U.S. FDA</li> <li>• Commenced Phase II dose-ranging clinical trial</li> <li>• Commenced enrollment of Phase II clinical trial with Pulmozyme®</li> </ul>	<ul style="list-style-type: none"> <li>• Initiate multi-national Phase III clinical trials (2006)</li> </ul>
<i>COPD – bronchiectasis</i>	<ul style="list-style-type: none"> <li>• Phase II trial complete</li> <li>• Granted Orphan Drug designation by the U.S. FDA</li> </ul>	<ul style="list-style-type: none"> <li>• Initiate international Phase III clinical trial (4Q05 – 1Q06) and U.S. Phase III clinical trial (Mid-2006)</li> </ul>
<i>COPD – chronic bronchitis</i>	<ul style="list-style-type: none"> <li>• Completed pilot clinical trial</li> </ul>	<ul style="list-style-type: none"> <li>• Initiate Phase II clinical trial (2006)</li> </ul>
<b>PXS25/64</b>		
<i>Multiple sclerosis</i>	<ul style="list-style-type: none"> <li>• Pre-clinical toxicology studies</li> </ul>	<ul style="list-style-type: none"> <li>• File U.S. IND (1H06)</li> </ul>
<b>PXS2076</b>		
<i>Rheumatoid arthritis</i>	<ul style="list-style-type: none"> <li>• Research to identify preclinical development candidate</li> </ul>	<ul style="list-style-type: none"> <li>• Nominate IND candidate (2006)</li> </ul>

## Lung Disease Overview

Our lead product candidates are for the diagnosis and treatment of chronic respiratory diseases, including asthma, cystic fibrosis and COPD. Several of these diseases share similar biology and pathology, such as the airway inflammation in both asthma and chronic bronchitis, as well as difficulty with normal clearance of lung mucus in patients with cystic fibrosis and bronchiectasis.

### *Asthma*

Asthma is a chronic inflammatory disease of the lungs where the airways narrow in response to a variety of stimuli. Published estimates indicate that this disease affects over 20 million people in the U.S. and approximately 51 million people in the seven major pharmaceutical markets of the U.S., Germany, France, United Kingdom, Italy, Spain, and Japan. Based on published studies, we estimate that each year in the U.S., 4.7 out of every 1,000 people under the age of 16 are newly diagnosed with asthma and two out of every 1,000 people aged 16 to 44 are newly diagnosed with the disease.

Many patients with asthma are not currently diagnosed with the disease. Sufferers and even physicians often attribute common asthma symptoms, such as cough and breathlessness, to smoking, lack of fitness or old age. Moreover, according to a recent publication, 34% of individuals diagnosed as asthmatic by their primary care physician do not have the disease. Even when accurately diagnosed, many patients do not receive the most appropriate therapy according to published guidelines. Physicians can underestimate the severity of the disease, and prescribe only bronchodilators, whereas the addition of an inhaled corticosteroid is the recommended course of action according to the Global Initiative for Asthma, or GINA, guidelines. We estimate that only about 30% of asthma patients in the U.S. receive inhaled corticosteroids despite evidence that uncontrolled asthma is common. Poorly controlled asthma can lead to irreversible damage to the airways. Therefore, the goal of treatment is to provide sufficient anti-inflammatory medication to control inflammation and airway remodeling. However, using high doses of medication can lead to unwanted side effects such as developmental difficulties in children and glaucoma in adults. Hence, selecting the right dose for individual patients remains a clinical problem.

To diagnose asthma and to evaluate patient response to treatment, pulmonary specialists will introduce an aerosolized substance directly into the lungs, and subsequently test lung function. These tests fall into two categories. The first category, known as “direct” challenge tests, use either histamine or methacholine to directly cause airway narrowing. These substances act on receptors on bronchial smooth muscle to cause contraction. The second category, known as “indirect” challenge tests, involve stimuli such as exercise, rapid breathing of dry air, or inhalation of salt solutions or adenosine monophosphate. This more closely mimics an asthmatic process, and can cause the release of chemicals from inflammatory cells within the lungs, resulting in airway contraction and narrowing.

The only FDA-approved direct test is Provocholine<sup>®</sup> (methacholine), marketed by Methapharm Inc. We believe that the disadvantage of direct tests are that the airway narrowing caused by histamine or methacholine is not dependent on the presence of inflammatory cells, and thus has a poor sensitivity for identifying people with active airway inflammation. Moreover, a positive response is not specific for identifying asthma and can occur in healthy people with no symptoms, smokers, and those with other diseases of the lung. Despite these limitations, we believe that over 200,000 direct tests are performed each year in the U.S., based on information reported by Solucient LLC in 2003. However, this represents only a small fraction of the potential market.

We believe that the indirect tests have a much lower false positive rate for asthma and increased sensitivity. However, each of them suffer from limitations. For example, tests involving exercise and rapid breathing of dry air require a lengthy period of time to complete and they require complicated equipment. Furthermore, these tests are limited to identifying exercise induced asthma and are not useful for determining the severity of airway inflammation. Hypertonic saline, which is delivered by a nebuliser during administration of the test, is uncomfortable for the patient, determination of the administered dose is difficult and this procedure is unsuitable for managing anti-inflammatory drug treatment. Adenosine monophosphate is unstable, also delivered by a nebuliser and its use is restricted to specialist research laboratories.



### *Lung Congestion*

The inside lining of the airways is covered by millions of fine hair-like structures called cilia, which are in turn covered by a surface liquid and a thin layer of mucus, secreted by the lungs to defend against germs, dust particles and other foreign bodies. The cilia move continuously and propel the mucus up to the throat. This constant process, which is barely noticeable in healthy people, cleans the airways, permits clean air to pass freely through the lungs and removes bacteria, thereby limiting infectious episodes.

Patients with COPD or with CF are generally affected by a breakdown in natural mechanisms of creating, hydrating, and clearing this mucus. These patients face the ongoing challenge of clearing excessive and thickened secretions from their congested lungs, usually by constant coughing. A key therapeutic goal for clinicians treating these patients is to assist the natural process of keeping the mucus hydrated and clearing it from the lungs.

### *Cystic Fibrosis*

CF is an inherited, progressive and fatal disease that affects epithelial surfaces including the airways, pancreas, sweat ducts, reproductive system and intestinal tract. The lungs of CF produce copious amounts of thick, tenacious secretions which are not cleared effectively by the lungs. Such changes are known to be present from birth and inevitably result in airway obstruction and bacterial infection. This generally leads to progressive lung deterioration, and eventually respiratory failure, the primary cause of death in adult CF patients.

According to Datamonitor, in 2002 there were 30,000 diagnosed CF patients in the U.S. and 65,000 in the seven major pharmaceutical markets. While this patient population is relatively small, the problem of sputum clearance is common to all sufferers and is a chronic lifelong problem. We believe that the annual healthcare cost for patients in the United States is over U.S.\$1 billion.

There is no cure for CF. Maintaining a reasonable quality of life for these patients is a significant challenge. Problems include breathing difficulties, respiratory infections, poor sleep, general discomfort, lifestyle limitations and gradual deterioration of lung function over time. Although the life expectancy of CF sufferers has increased over the past few decades due to better management of the disease, according to Datamonitor, the median life expectancy for patients with cystic fibrosis is only 31 years of age.

Physicians seek to improve lung function and reduce the number and severity of secondary lung infections by hydrating and breaking down the excessive, sticky mucus secretions, allowing it to be cleared from the lungs. Management of CF includes exercise, daily physiotherapy, postural drainage and chest percussion and can take several hours of at-home treatment every day. Medications to treat CF are limited, and few are very effective or convenient. Nebulised medications, delivered by aerosol or a facemask, are used to make the mucus less thick and sticky and open up the airways. Antibiotics may also be required to treat secondary infections, and are often used to prevent such infections.

Pulmozyme<sup>®</sup>, marketed by Genentech in the U.S., is the most widely used therapeutic for chronic use in CF to aid sputum clearance. According to Hoffman-La Roche Inc., worldwide sales of Pulmozyme were approximately U.S.\$265 million in 2004. We estimate that this represents a market penetration in developed countries and the seven major pharmaceutical markets of about 30%. Although Pulmozyme demonstrates lung function improvement in CF patients, similar benefit was not shown in other respiratory conditions, including bronchiectasis. Further, in previous clinical trials, Pulmozyme provided no increase in sputum clearance. Pulmozyme is unstable and is delivered by a nebulizer. Solutions have to be prepared by the patient before administration, the treatment periods are long and all equipment has to be sterilized after use.

### *Chronic Obstructive Pulmonary Disease*

Chronic Obstructive Pulmonary Disease, or COPD, encompasses a number of serious conditions affecting the lungs, including emphysema, chronic bronchitis and bronchiectasis. According to Datamonitor, there are 16 million people diagnosed with COPD in the U.S., and more than 30 million people are affected with COPD in the seven major pharmaceutical markets. In 2000, COPD was estimated to be responsible for the deaths of more than 119,000 people in the U.S. alone, making it the fourth leading cause of death after heart disease, cancer and stroke. In the United States, there are more than 10 million physician office visits and two million hospitalizations per year. The disease was estimated to cost the U.S. healthcare system U.S.\$30 billion in 2000.

Current management of COPD generally involves bronchodilators and steroids. However, only an estimated 20%-25% of patients respond positively to steroids and it is not practical to determine in advance which patients will respond to steroids. We believe that only half of moderate and severe COPD patients achieve an adequate treatment outcome. Therefore, as with asthma, we believe there is room to improve both the diagnosis and management of COPD.

### *Bronchiectasis*

In this disease the bronchial tubes become enlarged and distended, and the cilia do not function normally. Many patients with cystic fibrosis and asthma also have bronchiectasis. For other patients, bronchiectasis is a result of infections such as pneumonia, or the chronic inhalation of noxious substances. This results in poor clearing of mucus and predisposes the lung to more infections. The body repairs damaged lung tissue by forming tough, fibrous material, which leads to reduced lung function, lower lung efficiency, changes of the organization of blood vessels and increased blood flow through the lungs. These changes impair normal lung function and can ultimately lead to heart failure. Recurrent lung infections commonly reduce patients' quality of life and progressive respiratory insufficiency is the most common cause of death from this disease. Based on disease incidence data from a number of studies, we estimate that bronchiectasis affects about 580,000 people worldwide and about 100,000 people in the U.S.

Bronchiectasis treatment is aimed at controlling infections, increasing secretions, reducing airway obstructions and minimizing complications. Daily drainage to remove bronchial secretions is a routine part of treatment. Physicians often prescribe medications similar to those for chronic bronchitis, including inhaled bronchodilators to dilate the airways. Although antibiotics can be used to some effect to clear infections, no currently approved products effectively clear excess mucus secretions and improve the quality of life of these patients. Furthermore, because of the serious damage to lung tissue present in these patients, medications generally do not provide substantial improvement in lung function.

### *Chronic Bronchitis*

Patients with chronic bronchitis experience persistent airway inflammation and airflow obstruction, with symptoms including a chronic mucus-producing cough and shortness of breath. Due to the difficulties they have in clearing mucus from their lungs, sufferers are prone to periodic bacterial infections where their cough worsens, mucus production increases and breathing becomes more difficult. These episodes damage and scar the bronchial lining and contribute to continued chronic inflammation and immune-mediated cell damage as the body struggles to fight the infections. This cycle of infection and internal scarring causes a progressive decline in lung function, reducing quality of life and ultimately causing death.

Many of the deaths associated with chronic bronchitis are included in the COPD figure that now accounts for over 100,000 deaths a year in the U.S. The disease is predominately caused by inhaling some form of lung irritant repeatedly for many years, usually cigarette smoke. Chronic bronchitis is slow to develop and is often not diagnosed until the sufferer is in their 40s or 50s. The exact prevalence is not known but it may be as high as 10% of people over the age of 40.

Management of chronic bronchitis includes various general supportive measures such as giving up smoking, limiting exposure to dust and chemicals, avoiding sudden temperature changes, undertaking chest physiotherapy and deep-breathing exercises, and increasing fluid intake to keep the bronchial secretions thin. While there are a number of medications that dilate the airway and reduce airway inflammation, for chronic bronchitis sufferers, there are few therapeutic products available to effectively clear excess mucus secretions. This presents a major medical challenge, as ineffective mucus clearance is a major cause of infection and progression of the disease.

Treatments for chronic bronchitis include anti-cholinergic agents, steroids, antibiotics and oxygen. Anticholinergic agents, also known as antimuscarinics, are bronchodilators used for the relief of acute symptoms in both asthma and COPD, but tend to be more effective in COPD. Inhaled corticosteroids are less likely to cause systemic side effects than oral corticosteroids, and have been shown to be effective in asthmatics. However, the role of these agents in the management of COPD remains unclear. Inhaled steroids are effective in controlling inflammation in the lung, but based on a recent publication by Leuppi *et al.*, they are only effective in about 20%-25% of cases of patients with COPD.

## **Aridol**

We are initially developing Aridol as a more accurate and precise proprietary tool for physicians to use in the diagnosis and management of asthma and COPD. Physicians do not currently have rapid, accurate, safe and inexpensive tests to evaluate the presence or severity of these diseases. Aridol is a proprietary dry powder formulation of mannitol, delivered to the lungs through an inhaler. Mannitol is an osmotic agent which causes the release of certain mediators from inflammatory cells, which in turn cause a bronchoconstriction. This process mimics the changes that occur in the airways during exercise. Asthma patients who are not receiving adequate doses of anti-inflammatory medicine, such as an inhaled corticosteroid, experience airway narrowing and a drop in lung capacity when given the Aridol test. In contrast, healthy people or well-controlled asthma patients do not experience this narrowing and reduction in lung capacity. We recently completed a 646 patient, 12 center, Phase III clinical trial of Aridol. Based on the Phase III data, we have submitted applications to market in Australia and the European Union. We initiated a Phase III clinical trial in the fourth quarter of 2005 for approval in the U.S. If this trial is successful, we intend to file for approval with the FDA later next year. Based on independent research, we believe that the Aridol test will be reimbursable for the detection of airway hyperresponsiveness under current procedure codes in the United States. Based on independent market research commissioned by us, we believe that the addressable annual market for Aridol in the U.S., Europe and Australia includes the existing 400,000 bronchial challenge tests performed yearly, 2 million new tests for assisting the diagnosis of asthma, 16 million new tests performed by pulmonary specialists and primary care physicians for assisting the management of asthma, and 3 million new tests to evaluate steroid responsiveness in COPD patients.

Aridol has been used by over 1,200 people without serious adverse events, and is the subject of 30 peer-reviewed publications in international journals. We believe that Aridol may be superior to direct tests such as methacholine because Aridol is an indirect challenge test that relies on mediators released by inflammatory cells to cause a bronchoconstriction, thereby making Aridol a more accurate predictor of airway inflammation. We believe that Aridol's high degree of sensitivity and specificity for airway inflammation, combined with its ease of use, will make it possible for physicians to:

- diagnose asthma more accurately and objectively, and measure disease severity, with a high correlation to in-depth patient assessment by a pulmonary specialist physician;
- monitor the effectiveness of treatment, with a negative Aridol test indicating good control of asthma and a positive test indicating active airway inflammation and the need for more or different medication;
- determine the minimum required dose of steroids to achieve adequate disease control in a given patient, and predict the risk of exacerbation when reducing the steroid dose; and
- identify the 20%-25% of COPD patients who, according to a recent publication by Jörg Leuppi and colleagues, have a significant inflammatory component to their disease which will respond to inhaled corticosteroids. We believe this will allow the reduction of unnecessary steroid usage and potentially lower healthcare costs.

We have obtained from Central Sydney Area Health Service an exclusive, worldwide license to certain key intellectual property and patents relating to the use and formulation of Aridol.

### *Aridol for Asthma*

In our Phase II and Phase III clinical trials, patients use a dry powder inhaler to take progressively higher doses of Aridol (from 5 mg to 635 mg, nine steps in all). After each inhalation the patient's lung capacity is determined by a spirometer, an instrument to measure airflow and lung capacity. The Aridol test is stopped when a patient has a 15% fall in lung capacity, indicating the presence of active airway inflammation. Only those patients with active airway inflammation will experience a drop in lung capacity. On average, the procedure takes 17 minutes for a positive test and 26 minutes for a negative test. The only equipment required is a standard spirometer to record lung capacity.

A large number of investigator-sponsored, open-label Phase I and Phase II clinical trials have been conducted with Aridol. The results show that use of Aridol can identify subjects with asthma who are also responsive to inhaled salt solutions, inhaling dry air and exercise. Aridol also identifies both adults and children with

currently active asthma who are responsive to methacholine, as well as others who are not responsive to methacholine. The Aridol test demonstrates good repeatability in both adults and children, and responses are rapidly reversible using a standard dose of bronchodilator. Furthermore, Aridol can provide an assessment of the effectiveness of inhaled steroids in controlling the disease. Finally, Aridol response correlates with the symptoms and signs of exercise induced asthma, indicating that a negative response to Aridol may be a useful end point signifying adequate asthma control.

We recently completed a 12 center, 646 patient, Phase III clinical trial of Aridol to identify airway hyperresponsiveness in asthmatic patients, and to support filing for marketing authorization in Australia and the European Union. Airway hyperresponsiveness is a hallmark of untreated or poorly controlled asthma, and over time can lead to long-term changes in the lungs. However, long term use of inhaled corticosteroids can result in developmental abnormalities in children, and other complications. This trial included asthmatic patients who were currently treating their disease, patients with symptoms suggestive of asthma but without a clinical diagnosis, and healthy volunteers, including both children and adults. The goals of this trial were to:

- compare Aridol to hypertonic saline in identifying airway hyperresponsiveness in asthmatic subjects and non-asthmatic subjects;
- compare Aridol to standard clinical assessment in diagnosing asthma;
- compare asthma severity as determined by our Aridol test to the Severity of Asthma (Asthma Management Handbook 2002);
- evaluate the advantages of Aridol versus hypertonic saline with respect to simplicity, safety and patient and health care convenience; and
- further evaluate the safety profile of Aridol.

The primary endpoint was a comparison of the sensitivity and specificity of Aridol to that for an unapproved test, hypertonic saline, which is widely used in Australia. A secondary endpoint was a comparison of the sensitivity and specificity of Aridol to that of physician diagnosis. Sensitivity is a measure of the percentage of people correctly identified as having airway hyperresponsiveness by the test. Specificity is a measure of the percentage of people correctly identified as lacking airway hyperresponsiveness.

In this trial, sensitivity of Aridol against hypertonic saline was 81%, and specificity was 87%. This means that 81% of patients identified as having airway hyperresponsiveness by the hypertonic saline test were also identified as positive by the Aridol test and 87% of patients classified as lacking airway hyperresponsiveness were also identified as negative by Aridol. Conversely, the sensitivity of hypertonic saline against Aridol was 88%, and specificity was 79%. These numbers indicate good agreement between the two tests ( $p < 0.01$ ).

In comparison to physician diagnosis, Aridol had a sensitivity of 58%, and specificity was 94%. Significantly, of the 42% of patients identified as asthmatic by physician diagnosis, but lacking airway hyperresponsiveness as determined by Aridol, 85% were using inhaled corticosteroids at the time of the clinical trial. When the subjects who were Aridol negative and were using inhaled corticosteroids were removed from the analysis versus physician diagnosis, sensitivity was 89% and specificity was 95%. The increase in sensitivity underscores the utility of Aridol in managing patients on inhaled corticosteroid medication. We believe that these results, combined with data from dozens of other Aridol clinical trials, will provide the basis for approval in Australia and in the E.U.

As a result of this trial, we have filed a marketing authorization application through a contract research organization for Aridol for the identification of asthma with the Swedish Medical Products Agency in May 2005 as our entry to the mutual recognition procedure in the European Union. The Australian application for marketing authorization was submitted to Therapeutic Goods Administration, or the TGA, in January 2005. We believe that we could receive Australian and European regulatory authorizations to market Aridol during the first half of 2006. Currently, we intend to establish marketing partnerships in select territories for this product. We are supporting a number of investigator-sponsored trials to provide the basis for a rapid uptake of Aridol in the marketplace.

In the U.S., unlike Australia and Europe, a product, methacholine, is approved by the FDA to identify airway responsiveness in asthmatic patients. Based on discussions with the FDA, we are undertaking a Phase III

clinical trial comparing Aridol with methacholine and exercise challenge in patients with suspected asthma. The primary endpoint will be to compare the sensitivity and specificity of Aridol to identify exercise-induced bronchoconstriction. We initiated this trial in December 2005, and the results are due mid 2006.

Our initial target market for Aridol is the 50% of symptomatic asthmatics that are not diagnosed by physicians, as well as patients whose disease is poorly controlled. Because current use of objective lung function testing is low, we plan to focus initial Aridol marketing efforts on physician education regarding asthma diagnosis and disease control. We believe physicians who commonly diagnose asthma based only on patient history of asthma symptoms leads to sub-optimal control of this disease, falling far short of the goals of current clinical guidelines. We are also planning development and marketing efforts in new areas where challenge testing could be useful given the availability of an accurate, valid and easy to use test like Aridol. These include monitoring asthma therapy and assessing asthma prevalence in the community.

#### *Aridol for COPD*

We are also exploring the use of Aridol in the management of COPD. Treatment of COPD is difficult but approximately 20%-25% of patients with COPD can have a positive clinical outcome with the administration of inhaled steroids. A long standing problem has been that there is currently no test to identify those people that will respond clinically to inhaled steroids. A recent publication by Leuppi has shown that in an investigator-sponsored, Phase II clinical trial, those patients with COPD that have a positive response to an Aridol challenge test are likely to benefit from inhaled corticosteroids treatment. In this trial, all patients had a positive response to inhaled histamine (a lung challenge test) whereas only 23% had a response to inhaled Aridol. After three months treatment with steroids, only those patients who recorded a positive Aridol challenge test had an improvement in their lung capacity. The difference in response to treatment between the two groups was highly statistically significant ( $p=0.001$ ). We have commenced an additional Phase II clinical trial to determine if Aridol is a practical test to guide treatment of inhaled corticosteroids in COPD patients in the primary care setting. The objective of this trial will be to seek regulatory approval for the use of Aridol in COPD management.

#### **Bronchitol**

We are developing Bronchitol, our proprietary inhaled mannitol formulation, for the treatment of chronic obstructive lung diseases, including cystic fibrosis, bronchiectasis and chronic bronchitis. Mannitol is accepted as a food additive in the U.S. and is included in the FDA Inactive Ingredients Guide for drug products. We manufacture mannitol into a dry respirable powder and incorporate it into a capsule. The compound is delivered to a patient's lungs via a pocket-sized inhaler. In a Phase II trial involving 39 cystic fibrosis patients sponsored by us, Bronchitol provided a statistically significant reduction in airway obstruction and improvement in lung function. In a Phase II clinical trial sponsored by us and involving 60 patients with bronchiectasis, Bronchitol provided a statistically-significant increase in patients' quality of life relative to placebo and a highly statistically significant reduction in the symptoms of the disease. Patients have used Bronchitol at home successfully for up to 14 days without any serious adverse events. Based on the results from these two trials, we plan to initiate a pivotal Phase III clinical trial of Bronchitol for the treatment of bronchiectasis, the first of two planned for this indication, during the fourth quarter of 2005 or the first quarter of 2006, and pivotal Phase III clinical trials for cystic fibrosis during the first half of 2006. We have obtained from Central Sydney Area Health Service an exclusive, worldwide license to certain key intellectual property and patents relating to the use and formulation of Bronchitol.

#### *Mechanism and Early Data*

Bronchitol increases mucociliary clearance in asthmatic and healthy subjects. We have shown that a single inhalation of Bronchitol increases the clearance of mucus both acutely and over a 24 hour period in patients with bronchiectasis, and acutely in patients with cystic fibrosis. The precise mechanism whereby Bronchitol increases the clearance of mucus remains unclear. One possibility is that Bronchitol increases the clearance of mucus by changing its viscosity through disruption of hydrogen bonds that join the oligosaccharides in the mucin macromolecules. A second possibility is that Bronchitol increases hydration of the airway surface by drawing water towards the airway surface by osmotic pressure and enhancing cilia beat frequency. Our data in human patients and animal models support both hypotheses.

In an investigator-sponsored 19 patient, single-dose Phase II clinical trial of Bronchitol in patients diagnosed with bronchiectasis, an increase in whole lung mucus clearance was observed over a 75 minute period

beginning at the onset of intervention and this increase was statistically significant ( $p < 0.005$ ). There was an almost doubling of mucus clearance after Bronchitol treatment and most of this was in the central and intermediate regions of the lung. Over a 24 hour period after Bronchitol intervention the increase in mucus clearance was approximately 30% over control and this was statistically significant ( $p < 0.0001$ ).

#### *Bronchitol for CF*

In August 2005, we announced topline results from a Phase II clinical trial involving 39 patients with cystic fibrosis. Pharmaxis sponsored placebo-controlled trial was conducted at several sites in Australia and New Zealand. Patients were treated for two weeks with either Bronchitol or placebo. After a two week washout period where patients received neither drug or placebo, patients who previously received Bronchitol were treated with placebo, and vice versa. This crossover trial design allows each patient to act as their own control. The primary endpoint was change in Forced Expiratory Volume in 1 second, known as  $FEV_1$ . This is a quantitative measure of the volume of air a patient can exhale in one second, and is the most frequently used measure of the degree of airway obstruction. The secondary endpoints included quality of life, sputum microbiology, the physical properties of sputum, safety and additional lung function measurements. At the end of the treatment period, patients receiving Bronchitol had significantly better lung function compared to placebo as measured by  $FEV_1$  and for the maximum mid-expiratory flow, or MMEF, another measure of airway function. Approximately half the subjects were using Pulmozyme during the trial.

In this trial, Bronchitol had a positive impact on lung function. Patients who received Bronchitol had a 7% improvement in  $FEV_1$  as compared to placebo ( $p = 0.008$ ). MMEF increased by 15% while on Bronchitol treatment and this increase was significant compared to control ( $p < 0.01$ ). The MMEF reflects function in small airways and is an early abnormality in cystic fibrosis. Respiratory symptoms determined from a Likert scale self-assessment after Bronchitol treatment were significantly improved as compared to placebo ( $p < 0.02$ ). Additional longer term clinical studies will be required before Bronchitol can be considered for marketing and these additional studies may yield different results.

In August 2005, the FDA granted Orphan Drug designation to Bronchitol for the treatment of CF. We have initiated several additional clinical trials of Bronchitol for the treatment of this disease. In December 2005, we commenced a Phase II clinical trial to compare the effect on lung function of Bronchitol and Pulmozyme to either drug alone. We also started a dose-ranging Phase II trial to determine optimal dosing for Phase III clinical trials. Based on the outcome of these trials, in 2006 we plan to initiate the first pivotal Phase III clinical trial to provide the basis for applications for marketing authorization in the U.S., the E.U. and Australia.

#### *Bronchitol for Bronchiectasis*

We recently completed a proof of concept Phase II clinical trial of Bronchitol in 60 bronchiectasis patients. We began this placebo-controlled, crossover design trial at a single center in Sydney and later expanded it to include four centers in Australia and New Zealand. This trial was designed to explore the safety and efficacy of Bronchitol in bronchiectasis patients. Patients received 400 mg of Bronchitol or placebo, twice a day for 14 days. In this trial, placebo was a mannitol formulation with a larger particle size, which we anticipated to be most similar in patient experience to active Bronchitol, yet was intended not to enter the lungs to any significant degree. Endpoints of the study were to evaluate the effect of Bronchitol treatment on patient quality of life using a self-assessment known as the Likert scale, the St. Georges Hospital Respiratory Questionnaire, or SGRQ, which is another self assessed measure of quality of life, sleep quality as measured by the self assessed Epworth scale, exercise tolerance as measured by the 6 minute walk test, lung function as measured by two tests known as spirometry and flow oscillometry, sputum microbiology, the physical properties of sputum, the volume of sputum production over 24 hours and the safety profile of Bronchitol. The SGRQ includes changes in three components, symptom, activity and impact, as well as an overall score. Improvement in quality of life measures is indicated by a reduction in score.

Versus baseline, treatment with Bronchitol led to a significant reduction in the Likert scale score of 6.1 ( $p = 0.03$ ). Versus baseline and placebo, there was a statistically significant improvement in the Epworth sleep score ( $p < 0.02$  versus placebo). For patients receiving Bronchitol, 38% went from an unclear chest to a clear chest as compared to 17% on placebo ( $p < 0.05$ ). There was no statistically significant changes on lung function as measured by standard spirometry. Flow oscillometry showed a significant effect of Bronchitol compared to placebo ( $p < 0.05$ ). Flow oscillometry is considered to reflect changes in small airways.

However, the effect of Bronchitol was most pronounced in the 75% of patients who entered the study with an unclear chest, which indicates the most serious problems with normal clearance of lung mucus. There was a mean decrease of 10.2 in Likert scale score during Bronchitol treatment, compared to a mean decrease of 3.6 for placebo ( $p < 0.005$  versus placebo). Treatment with Bronchitol led to a significant improvement in the impact component of the SGRQ compared to placebo in those patients with an unclear chest. The improvement was clinically significant at 6.9 points. There was also a trend for an effect on the total score versus placebo but this did not reach significance ( $p = 0.15$ ). Compared to baseline, the overall score showed a strong trend with a clinically significant reduction of 5.6 ( $p = 0.055$ ).

In this exploratory trial, we saw statistically significant changes in several endpoints versus patient baseline, as well as statistically significant effects in the 75% patients with the most serious problems with normal clearance of lung mucus. For other endpoints, we saw effects ranging from strong trends to modest or no effect. We are currently evaluating the most appropriate control for Bronchitol clinical trials in bronchiectasis. No therapies to enhance mucus clearance in bronchiectasis patients have been approved in over 20 years in the U.S. Accordingly, we are currently in discussions with the FDA regarding appropriate primary endpoints for a pivotal clinical trial program.

In February 2005, the FDA granted Orphan Drug designation to Bronchitol for the treatment of bronchiectasis. We are currently supplying Bronchitol in Australia on an individual, named patient basis under a TGA-administered compassionate use program known as the Special Access Scheme. This program allows patients access to unapproved drugs where there are limited treatment options. We are planning pivotal Phase III clinical trials of Bronchitol for the treatment of this disease and intend to commence the first of these trials in Australia and Europe during the fourth quarter of 2005 or first quarter of 2006, and a second pivotal Phase III trial in the U.S. in mid-2006. If these trials are successful, we expect the data to provide part of the basis for marketing approval in the U.S., the E.U. and Australia.

#### *Bronchitol for Chronic Bronchitis*

Pilot data in patients with chronic bronchitis have shown that Bronchitol may also be beneficial in improving ciliary and cough clearance in these patients. We indirectly supported a small, investigator-sponsored Phase II clinical trial to determine the effects of Bronchitol on mucus clearance over a two hour period, and the effects on rate of clearance of a radiolabelled tracer over a 24 hour period. The trial was not powered nor suitably controlled for statistical analysis, but provided encouraging data.

We plan to conduct additional Phase II clinical trials in patients with chronic bronchitis. The objective of these trials will be to determine if Bronchitol assists in clearing mucus during an exacerbation and that Bronchitol has the ability to shorten the exacerbation period. The first trial will be a Phase II clinical trial involving 60 patients and will be conducted in hospitals in Australia and New Zealand.

#### **PXS25/PXS64**

We currently conduct an active research program designed to prevent the inappropriate migration of immune cells from blood to surrounding tissue. Our lead compounds in this program are PXS25 and PXS64, an analogue of PXS25 that is orally available in our preclinical studies. Our preclinical studies indicate that PXS25 prevents immune cell migration and is effective in animal models of multiple sclerosis and rheumatoid arthritis. We are currently developing PXS64 as an oral treatment for acute exacerbations of multiple sclerosis, and are evaluating its potential in other inflammatory/autoimmune diseases such as irritable bowel disease, lupus and psoriasis.

#### *Background*

A key element of inflammatory diseases such as multiple sclerosis and rheumatoid arthritis is the migration of white blood cells known as leukocytes from blood vessels into surrounding tissue. After binding to the inside surface of blood vessels, the leukocytes produce enzymes which allow them to move into adjacent tissue. These enzymes are anchored to the leukocytes through a protein found on the surface of these cells, known as the mannose phosphate receptor, or MPR.

Multiple sclerosis, or MS, is a disease where the patient's own immune system attacks the protective myelin sheath that insulates nerve fibres, or axons, in the central nervous system and the cells that make myelin, the

oligodendrocytes. The symptoms of MS result from both the loss of signal conduction in demyelinated axons and the loss of the axons themselves. Under normal circumstances the blood-brain barrier protects cells in the central nervous system, or CNS, from leukocytes found in the rest of the body. However, in MS patients activated T-cells are able to cross the blood-brain barrier by secreting proteases and thereby enter the CNS.

According to the Mayo Clinic, there are approximately 1 million people affected by MS worldwide. There is no cure for the disease, although treatments do exist to slow progression of the disease. There are currently five main drugs on the market for the treatment of MS: three beta interferons, Copaxone<sup>®</sup> marketed by Teva and Aventis, and Novantrone<sup>®</sup>, marketed by Serono. In addition, powerful immunosuppressive steroids such as prednisone are often used to treat patients in the acute relapsing phase of the disease, the most serious and debilitating. Based on sales and reported market share estimates published in the annual reports of a number of the manufacturers of these five main drugs, we estimate the worldwide market for drugs to treat MS to be approximately U.S.\$3 billion per year.

#### **PXS25/PXS64 for MS**

We believe drugs that prevent activated leukocytes from entering the CNS will inhibit lesion formation and slow the progression of MS. PXS25 is a small molecule that acts as a specific blocker of the mannose phosphate receptor, which blocks protease secretion and function *in vitro*. In animal models of MS, treatment with PXS25 has resulted in a reduction in peak severity of disease and accelerated recovery. PXS25 represents a new therapeutic approach to treating multiple sclerosis and we believe the compound could have broad utility in slowing the progression of the disease.

In our animal studies, PXS25 demonstrated significant activity when administered by injection. However, the oral bioavailability of PXS25 is low in several species of animals. Therefore, we have developed a prodrug of PXS25 which is metabolized to active PXS25 once absorbed by the body. This compound, PXS64, is now in preclinical development. Our goal is to file an IND in the first half of 2006 and to pursue an application for the treatment of multiple sclerosis.

#### **PXS2076**

We are developing PXS2076 for the treatment of rheumatoid arthritis. PXS2076 is an orally-available, selective agonist of the cannabinoid-2, or CB<sub>2</sub>, receptor. This receptor has been so named as it belongs to a family of cell surface receptors that are activated by chemicals found in the cannabis plant. The CB<sub>2</sub> receptor is found predominantly on inflammatory cells where its activation can prevent the release of inflammatory proteins that cause much of the tissue damage resulting from inflammation. In 2000, researchers from University College, London published a paper in *Nature* which demonstrated that activation of CB<sub>2</sub> receptors reduced MS symptoms, such as tremor and spasticity, in an animal model of the disease.

Collectively, this and other research has driven our interest in identifying selective CB<sub>2</sub> receptor agonists to address inflammatory/autoimmune diseases without the psychotropic side effects associated with cannabis use. PXS2076 is a synthetic, low molecular weight molecule invented in our research laboratories. It derives from a targeted medicinal chemistry program using computer modelling and assay directed synthesis to identify lead compounds for optimization. The lead compounds were derived from established non-selective ligands. PXS2076 activates the CB<sub>2</sub> receptor and inhibits the release of inflammatory proteins from inflammatory cells. PXS2076 is effective in an animal model of rheumatoid arthritis and we are evaluating its effects in other animal models of human diseases. We may choose to initiate a preclinical program in the fourth quarter of 2005.

#### **Our Strategy**

Our objective is to build a fully-integrated specialty pharmaceutical company focused on respiratory and inflammatory/autoimmune indications. Key aspects of our strategy include:

- *Obtain marketing authorization for Aridol.* We filed our marketing authorization of Aridol in Australia in January 2005 and in the E.U. in May 2005. Based on our pre-IND meeting with the FDA, we believe one additional successful clinical trial of Aridol would provide the basis for approval in the U.S.
- *Successfully complete the clinical development of Bronchitol.* We recently completed two successful Phase II trials of Bronchitol for the treatment of cystic fibrosis and bronchiectasis. We expect to initiate



our first pivotal Phase III trial in Australia and Europe with Bronchitol for the treatment of bronchiectasis in the fourth quarter of 2005 or the first quarter of 2006, and a second pivotal Phase III trial in the U.S. in mid-2006, and we expect to initiate pivotal Phase III clinical trials for cystic fibrosis in 2006.

- *Focus on attractive product opportunities in our core therapeutic areas.* We are developing products that address chronic respiratory and inflammatory/autoimmune diseases. These markets are attractive opportunities because of the limitations of current treatment, the severity of these diseases, and because they are treated by a relatively concentrated physician audience.
- *Develop sales and marketing capabilities in select markets.* We intend to retain commercial rights to our products in indications and territories where we believe we can effectively market them with a small specialized sales force. For all other indications and territories, we intend to pursue strategic collaborations.
- *Continue to expand our R&D pipeline.* We were founded in 1998 based on anti-inflammatory intellectual property we licensed from the Australian National University. In 2001, we acquired intellectual property that is the basis of our Aridol and Bronchitol product development programs. We will continue to build and strengthen our product pipeline and commercial capabilities, and we may acquire complementary technology and drug development candidates from research institutes, universities and private and public companies. These acquisitions may take the form of collaborations, licensing arrangements or outright purchase of intellectual property, research groups or corporate entities.

### **Sales and Marketing**

We have a small marketing group in Australia but currently have no sales or distribution capabilities. In order to commercialize any of our product candidates, we must further develop these capabilities internally or through collaborations with third parties. We intend to build our own sales force to market our products in Australia, and we intend to retain commercial rights to market our products to pulmonary specialists in the United States and potentially in Europe. Because the United States and European pulmonary specialist market is relatively concentrated, we believe we can effectively target it with a small specialized sales force. We may pursue strategic collaborations to commercialize our products in other territories and on a worldwide basis for indications treated by large physician populations, such as asthma or arthritis.

### **Manufacturing**

We manufacture both Aridol and Bronchitol in our production facility located in Sydney, Australia, under conditions of current Good Manufacturing Practice, known as cGMP. In early 2005, we completed an expansion of our manufacturing facility that approximately tripled our manufacturing capacity. Our manufacturing facility consists of a warehouse, adjoining office space, a cGMP laboratory for quality control and quality assurance, and clean rooms. Final packing of both Aridol and Bronchitol in blister packs is performed by a third party. The inhaler used in conjunction with Bronchitol is manufactured by Plastiapipe in Italy and is supplied to us on a non-exclusive basis through a standard supply agreement.

We believe that our manufacturing facility has ample operating capacity to produce adequate Aridol and Bronchitol to undertake the full clinical trial program through submission of an NDA in the U.S. for those product candidates and to support the commercial demand of Aridol two years after international launch. For larger-scale commercial manufacturing of these products, we may work with contract manufacturers, or we may choose further expansion of our current manufacturing capabilities.

Our cGMP facilities have been inspected and licensed by the Therapeutic Goods Administration, or TGA, the Australian regulator analogous to the FDA. Our facilities and those of any third-party manufacturers will be subject to periodic inspections confirming compliance with applicable law, including U.S. and European rules, should our product candidates receive approval in those jurisdictions. Our facilities must be cGMP certified before we can manufacture our drugs for commercial sale. Failure to comply with these requirements could result in the shutdown of our facilities or the assessment of fines or other penalties.

Mannitol is the key raw material required for the manufacture of both Aridol and Bronchitol. cGMP grade mannitol is available from a number of suppliers. Inhalers are also available from a number of suppliers.

PXS64 is a small molecular entity synthesized from inexpensive and readily available starting materials available from a number of different suppliers. The manufacture of PX64 is carried out by a contract manufacturer. We own all rights to the manufacturing process.

PXS2076 is a small molecular entity synthesized from inexpensive and readily available starting materials available from a number of different suppliers.

We have outsourced the manufacturing of cGMP grade PXS25 for preclinical trials and clinical trials as our manufacturing facilities are not suitable for the production of PXS25. Our contract manufacturers have the capacity to produce adequate PXS25 for clinical trials of PXS25. We believe we are also able to produce sufficient PXS2076 for research studies.

### **Competition**

We operate in highly competitive segments of the biotechnology and pharmaceutical markets. We face competition from many different sources, including commercial pharmaceutical and biotechnology enterprises, academic institutions, government agencies, and private and public research institutions. Many of our competitors have significantly greater financial, product development, manufacturing and marketing resources than do we. Large pharmaceutical companies have extensive experience in clinical testing and obtaining regulatory approval for drugs. These companies also have significantly greater research capabilities than do we. In addition, many universities and private and public research institutes are active in respiratory and autoimmune disease research, some in direct competition with us. We also compete with these organizations to recruit scientists and clinical development personnel. Smaller or early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies.

We are aware of many of our competitors in each of the markets we target. These products include approved and marketed products as well as products in development. We expect Aridol, if approved for the diagnosis and management of asthma, to compete with direct bronchiol provocation tests such as methacholine and histamine. We expect Bronchitol for CF to compete with or to be used in conjunction with Pulmozyme and other mucolytic agents. Although it has little market penetration, Mucomyst<sup>®</sup>, marketed by AstraZeneca, is used by some physicians to treat bronchiectasis, other forms of COPD and CF. Numerous other potential competing therapeutic products are in clinical treatment and preclinical development. In each of our development programs addressing indications for which there are therapies available, we intend to complete clinical trials designed to evaluate the potential advantages of our drug candidates as compared to, or in conjunction with, the current standard of care. Key differentiating elements affecting the success of all of our drug candidates are likely to be their efficacy, convenience and side-effect profile compared to commonly used therapies.

### **Intellectual Property**

We patent the technology, inventions and improvements that we consider important to the development of our business. As of December 9, 2005, we owned or had exclusive rights to 16 issued U.S. and foreign patents and ten pending U.S. and foreign patent applications. Of these, nine issued patents and three pending applications relate to Aridol and Bronchitol. The last of these issued patents are due to expire in 2021. One pending application in the form of a PCT international application relates to PXS25 and PXS64 and has now entered the national phase. The remaining patents and applications relate to other aspects of our technology or other drug discovery programs that have not yet entered a full development program. We intend to seek patent term extension for our eligible patents, including under the Hatch-Waxman Act, which provides up to five years of patent extension.

We have obtained from Central Sydney Area Health Service exclusive worldwide rights for certain key intellectual property and patents relating to the use of respirable dry powders for the assessment of bronchial hyper-responsiveness, a condition consistent with active asthma, for monitoring steroid use in asthma patients, and enhancing mucus clearance in diseases such as cystic fibrosis, bronchiectasis and chronic bronchitis. These exclusive rights, which form the basis for patent protection of both Aridol and Bronchitol, derive from one issued U.S. and eight issued foreign patents. The U.S. and most of the foreign patents covering Aridol and Bronchitol are due to expire in 2015. The latest expiring in any territory is 2021. The U.S. patent may be eligible for extension by up to an additional five years.

We also have an exclusive worldwide license from ANU Enterprises Pty Ltd. (formerly Anutech Pty Ltd.) to develop and commercialize intellectual property relating to the treatment of inflammatory or immune-mediated conditions in patients by administering a phosphosugar. These exclusive rights derive from two issued U.S. and four issued foreign patents covering the E.U. member states and Australia, as well as other major territories. The last of these patents are due to expire in 2017. The U.S. patents may be eligible for extension by up to an additional five years.

Our ability to build and maintain our proprietary position for our technology and drug candidates will depend on our success in obtaining effective claims and enforcing those claims once granted. The patent positions of biopharmaceutical companies like ours are generally uncertain and involve complex legal and factual questions for which important legal principles remain unresolved. Some countries in which we may sell our product candidates or license our intellectual property may fail to protect our intellectual property rights to the same extent as the protection that may be afforded in the United States or Australia. Some legal principles remain unresolved and there has not been a consistent policy regarding the breadth or interpretation of claims allowed in patents in the United States, the European Union, Australia or elsewhere. In addition, the specific content of patents and patent applications that are necessary to support and interpret patent claims is highly uncertain due to the complex nature of the relevant legal, scientific and factual issues. Changes in either patent laws or in interpretations of patent laws in the United States, the European Union or elsewhere may diminish the value of our intellectual property or narrow the scope of our patent protection.

We may not be able to develop patentable products or be able to obtain patents from pending patent applications. Even if patents are issued, those patents can be challenged by our competitors who can argue such patents are invalid. Patents also will not protect our products if competitors devise ways of making these product candidates without legally infringing our patents. The U.S. Federal Food, Drug and Cosmetic Act and FDA regulations and policies and equivalents in other jurisdictions provide incentives to manufacturers to challenge patent validity or create modified, noninfringing versions of a drug in order to facilitate the approval of abbreviated new drug applications for generic substitutes. In addition, patent applications filed before November 29, 2000 in the U.S. are maintained in secrecy until patents issue. Later filed U.S. applications and patent applications in most foreign countries generally are not published until at least 18 months after they are filed. Scientific and patent publication often occurs long after the date of the scientific discoveries disclosed in those publications. Accordingly, we cannot be certain that we were the first to invent the subject matter covered by any patent application or that we were the first to file a patent application for any inventions.

### **Government Regulation and Product Approval**

Regulation by governmental authorities in the United States and other countries is a significant factor in the development, manufacture and marketing of pharmaceuticals. All of our products will require regulatory approval by governmental agencies prior to commercialization. In particular, pharmaceutical drugs are subject to rigorous preclinical testing and clinical trials and other premarketing approval requirements by the FDA and regulatory authorities in other countries. In the United States, various federal, and in some cases state statutes and regulations, also govern or impact upon the manufacturing, safety, labeling, storage, record-keeping and marketing of pharmaceutical products. The lengthy process of seeking required approvals and the continuing need for compliance with applicable statutes and regulations require the expenditure of substantial resources. Regulatory approval, when and if obtained for any of our product candidates, may be limited in scope which may significantly limit the indicated uses for which our product candidates may be marketed. Further, approved drugs and manufacturers are subject to ongoing review and discovery of previously unknown problems that may result in restrictions on their manufacture, sale or use or in their withdrawal from the market.

#### *Preclinical Studies*

Before testing any compounds with potential therapeutic value in human subjects in the United States, stringent government requirements for preclinical data must be satisfied. Preclinical testing includes both *in vitro* and *in vivo* laboratory evaluation and characterization of the safety and efficacy of a drug and its formulation. Preclinical testing results obtained from studies in several animal species, as well as from *in vitro* studies, are submitted to the FDA as part of an investigational new drug application, or IND, and are reviewed by the FDA prior to the commencement of human clinical trials. These preclinical data must provide an adequate basis for evaluating both the safety and the scientific rationale for the initial trials in human volunteers.

### *Clinical Trials*

If a company wants to test a new drug in humans, an IND must be prepared and filed with the FDA. The IND becomes effective if not rejected or put on clinical hold by the FDA within 30 days. In addition, an Institutional Review Board comprised in part of physicians at the hospital or clinic where the proposed trials will be conducted must review and approve the trial protocol and monitor the trial on an ongoing basis. The FDA may, at any time during the 30-day period or at any time thereafter, impose a clinical hold on proposed or ongoing clinical trials. If the FDA imposes a clinical hold, clinical trials cannot commence or recommence without FDA authorization and then only under terms authorized by the FDA. In some instances, the IND application process can result in substantial delay and expense.

### *Clinical Trial Phases*

Clinical trials typically are conducted in three sequential phases, phases I, II and III, with phase IV trials potentially conducted after marketing approval. These phases may be compressed, may overlap or may be omitted in some circumstances.

- *Phase I clinical trials.* After an IND becomes effective, phase I human clinical trials can begin. These trials evaluate a drug's safety profile, and the range of safe dosages that can be administered to healthy volunteers and/or patients, including the maximum tolerated dose that can be given to a trial subject with the target disease or condition. Phase I trials also determine how a drug is absorbed, distributed, metabolized and excreted by the body, and duration of its action.
- *Phase II clinical trials.* Phase II clinical trials typically are designed to evaluate the potential effectiveness of the drug in patients and to further ascertain the safety of the drug at the dosage given in a larger patient population.
- *Phase III clinical trials.* In phase III clinical trials, the drug is usually tested in a controlled, randomized trial comparing the investigational new drug to an approved form of therapy in an expanded and well defined patient population and at multiple clinical sites. The goal of these trials is to obtain definitive statistical evidence of safety and effectiveness of the investigational new drug regime as compared to an approved standard therapy in defined patient populations with a given disease and stage of illness.

All clinical trials for our products have been conducted in accordance with the ICH (International Conference on Harmonization) guidance so that we can apply for marketing authorization in multiple jurisdictions.

### *New Drug Application*

After completion of clinical trials, if there is substantial evidence that the drug is safe and effective, a new drug application, or NDA, is prepared and submitted for the FDA to review. The NDA must contain all of the essential information on the drug gathered to that date, including data from preclinical and clinical trials, and the content and format of an NDA must conform with all FDA regulations and guidelines. Accordingly, the preparation and submission of an NDA is a major undertaking for a company.

The FDA reviews all NDAs submitted before it accepts them for filing and may request additional information from the sponsor rather than accepting an NDA for filing. In such an event, the NDA must be submitted with the additional information and, again, is subject to review before filing. Once the submission is accepted for filing, the FDA begins an in-depth review of the NDA. Typically, the FDA takes ten months to review and respond to the NDA. The FDA may refer the application to an appropriate advisory committee, typically a panel of clinicians, for review, evaluation and a recommendation as to whether the application should be approved. The FDA is not bound by the recommendation, but gives great weight to it. If the FDA evaluations of both the NDA and the manufacturing facilities are favorable, the FDA may issue either an approval letter or an approvable letter, which usually contains a number of conditions that must be satisfied in order to secure final approval. If the FDA's evaluation of the NDA submission or manufacturing facility is not favorable, the FDA may refuse to approve the NDA or issue a not approvable letter.

### *Other Regulatory Requirements*

Any products we manufacture or distribute under FDA approvals are subject to pervasive and continuing regulation by the FDA, including record-keeping requirements and reporting of adverse experiences with the products. Drug manufacturers and their subcontractors are required to register with the FDA and, where appropriate, state agencies, and are subject to periodic unannounced inspections by the FDA and state agencies for compliance with current GMP regulations which impose procedural and documentation requirements upon us and any third party manufacturers we utilize.

The FDA closely regulates the marketing and promotion of drugs. A company can make only those claims relating to safety and efficacy that are approved by the FDA. Failure to comply with these requirements can result in adverse publicity, warning letters, corrective advertising and potential civil and criminal penalties. Physicians may prescribe legally available drugs for uses that are not described in the product's labeling and that differ from those tested by us and approved by the FDA. Such off-label uses are common across medical specialties. Physicians may believe that such off-label uses are the best treatment for many patients in varied circumstances. The FDA does not regulate the behavior of physicians in their choice of treatments. The FDA does, however, restrict manufacturer's communications on the subject of off-label use.

The FDA's policies may change and additional government regulations may be enacted which could prevent or delay regulatory approval of our product candidates or approval of new indications for our existing products. We cannot predict the likelihood, nature or extent of adverse governmental regulations that might arise from future legislative or administrative action, either in the United States or abroad.

We received orphan drug designation for Bronchitol in August 2005 for the treatment of CF for patients at risk for developing bronchiectasis. Bronchiectasis is a major risk for CF patients. Under the Orphan Drug Act, the FDA may grant orphan drug designation to drugs intended to treat a rare disease or condition, which is generally a disease or condition that affects fewer than 200,000 individuals in the U.S. Orphan drug designation must be requested before submitting an NDA. After the FDA grants orphan drug designation, the identity of the applicant, the therapeutic agent and the designated orphan use are disclosed publicly by the FDA.

Orphan drug designation does not convey any advantage in or shorten the duration of the regulatory review and approval process. If a product that has orphan drug designation is the first such product to receive FDA approval for the disease for which it has such designation, the product is entitled to orphan product exclusivity, which means that the FDA may not approve any other applications to market the same drug for the same disease, except in limited circumstances, for seven years. The FDA may permit additional companies to market a drug for the designated condition if such companies can demonstrate clinical superiority. More than one product may also be approved by the FDA for the same orphan indication or disease as long as the products are different drugs. As a result, even if Bronchitol is approved to treat CF and receives orphan drug status, the FDA can still approve other drugs for use in treating the same indication or disease covered by Bronchitol, which could create a more competitive market for us. Moreover, if a competitor obtains approval of the same drug for the same indication or disease before us, we would be blocked from obtaining approval for our product for seven years, unless our product can be shown to be clinically superior.

### *Foreign Regulation*

In addition to regulations in the United States, we will be subject to a variety of foreign regulations governing clinical trials and commercial sales and distribution of our products. Whether or not we obtain FDA approval for a product, we must obtain approval of a product by the comparable regulatory authorities of foreign countries before we can commence clinical trials or marketing of the product in those countries. The approval process varies from country to country, and the time may be longer or shorter than that required for FDA approval. The requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement vary greatly from country to country.

Under European Union regulatory systems, marketing authorizations may be submitted either under a centralized procedure, a mutual recognition procedure or a decentralized procedure. The centralized procedure provides for the grant of a single marketing authorization that is valid for all European Union member states. The decentralized procedure provides for a joint assessment of safety and efficacy by a number of E.U. member states. The mutual recognition procedure provides for mutual recognition of national approval decisions. Under this procedure, the member state approving the first marketing authorization within the E.U. submits an application for

recognition to other E.U. member states. Within 90 days of receipt of the application and the first member state's report of the assessment of the drug, the other member states are supposed to recognize the marketing authorization of the first member state or refer the application to the Committee for Human Medicinal Products, or CHMP, for arbitration, if one or more member states believe there is a potential serious risk to public health, and the member states cannot reach agreement on the approval of the product. The CHMP is a scientific expert committee of the European Medicines Agency, or EMEA. The EMEA is responsible for the protection of public health in the E.U. through the coordination and evaluation and supervision of medicinal products, including administering the centralized procedure and performing a more limited role in the mutual recognition procedures. After member states agree to mutual recognition of the first marketing authorization, national marketing authorizations must still be issued in each member state which recognized it, including approval of translations, labeling and the like.

#### *Regulation in Australia*

In Australia, the relevant regulatory body responsible for the pharmaceutical industry is the Therapeutics Goods Administration, or TGA. The TGA maintains the Australian system of controls, safety, efficacy and availability of therapeutic goods used in Australia or exported from Australia.

Any products we manufacture in Australia or distribute in, or export from, Australia are subject to pervasive and continuing regulation by the TGA. Our products are subject to pre-market evaluation and approval by the TGA and must be entered on the Australian Register of Therapeutic Goods prior to commercial manufacture or sale. Our manufacturing facilities must be licensed by the TGA and our products must be manufactured in accordance with international standards of Good Manufacturing Practice. The TGA carries out a range of ongoing assessment and monitoring activities, including sampling, adverse event reporting, surveillance activities, and response to public inquiries and undertakes assessments of products for export. The TGA also regulates the advertising, labeling, product appearance and guidelines of our products.

The TGA's policies may change and additional governmental regulations may be enacted which could prevent or delay the regulatory approval of our product candidates or approval of new indications of our existing products. We cannot predict the likelihood, nature and extent or adverse governmental regulations that might arise from future legislative or administrative action.

In addition to regulations in Europe, Australia and the United States, we will be subject to a variety of foreign regulations governing clinical trials and commercial distribution of our products.

#### **Employees**

As of September 30, 2005, we had 32 employees, and 11 full time contractors, including 7 researchers based at the Australian National University, contracted completely to us by ANU Enterprises Pty Ltd under a service agreement between ANU Enterprises Pty Ltd and us. Thirty-three of our employees and full time contractors are engaged in research and development, with the remainder involved in administrative and marketing functions. Ten of our employees and full time contractors have doctoral degrees. We believe relations with our employees are generally good. None of our employees are covered by a collective bargaining agreement.

#### **Legal Proceedings**

We are not involved in any legal, arbitration or governmental proceedings which may have, or have had in the recent past, significant effects on our financial position or profitability. We are also not aware of any pending legal, arbitration or governmental proceedings against us which may have significant effects on our financial position or profitability.

#### **Research Grant Funding**

We have two research grants with the Commonwealth of Australia that assist us in funding certain of the research programs.

Under our AusIndustry P3 Pharmaceuticals Partnerships Program funding deed with the Commonwealth of Australia, subject to certain conditions, the Commonwealth of Australia may pay us a total amount of \$6.1 million between the July 2004 and June 2008 for eligible pharmaceutical research and development activities undertaken by

us in relation to the development of new treatments for autoimmune diseases and the development of new treatments for chronic respiratory diseases. For details regarding this research grant, see “Item 10. Additional Information – C. Material Contracts – AusIndustry P3 Pharmaceuticals Partnerships Program Funding Deed.”

Similarly, under our Start Grant Program funding deed with the Commonwealth of Australia, subject to certain conditions, the Commonwealth of Australia has provided the Company with a grant of 50% of the Company’s eligible expenditures on a project for the development of a new treatment for cystic fibrosis up to a maximum grant amount of \$3 million payable over the period from inception to December 31, 2005. For details regarding this research grant, see “Item 10. Additional Information – C. Material Contracts – Research and Development Start Program Grant Agreement.”

### **C. Organizational Structure**

Not applicable.

### **D. Property, Plant and Equipment**

As of June 30, 2005, we leased approximately 15,000 square feet of manufacturing, warehouse and office space at Unit 2, 10 Rodborough Road, Frenchs Forest, NSW 2086, Sydney, Australia. Our lease expires in June 2006, with an option to renew for a further five years thereafter. From July 1, 2001 to June 30, 2005, we spent approximately A\$2.8 million related to the establishment of this manufacturing facility. While the current manufacturing capacity expansion has been accommodated within this facility, we will require additional space and facilities as our business expands.

## **ITEM 5. OPERATING AND FINANCIAL REVIEW AND PROSPECTS**

*The following discussion and analysis should be read in conjunction with our financial statements and related notes included elsewhere in this report. This discussion and analysis contains forward-looking statements based upon current expectations that involve risks and uncertainties. Our actual results and the timing of events could differ materially from those anticipated in these forward-looking statements as a result of several factors, including those set forth under “Risk Factors” and elsewhere in this report. Please also see the section entitled “Special Note Regarding Forward-Looking Statements.” Our fiscal year ends on June 30. We designate our fiscal year by the year in which that fiscal year ends; e.g., fiscal year 2005 refers to our fiscal year ended June 30, 2005.*

### **A. Operating Results**

#### ***Overview***

We are a specialist pharmaceutical company involved in the research, development and commercialization of therapeutic products for chronic respiratory and autoimmune diseases. Research is in progress into new treatments for autoimmune diseases including multiple sclerosis and rheumatoid arthritis. We are most advanced in the development of products for asthma and for chronic obstructive pulmonary diseases, including bronchiectasis and chronic bronchitis, and cystic fibrosis. At June 30, 2005, we had one product in the marketing application stage in Europe and Australia, four projects at clinical trial stage (in patients), one project in pre-clinical evaluation (prior to being administered to volunteers or patients), and two research projects to identify a compound for development. Our development program has been designed to produce a series of products for large world markets over the coming years.

We were incorporated in May 1998 and in October 1999 obtained a license to a series of patents in the autoimmune area owned by the Australian National University, or ANU. We issued 11.2 million ordinary shares valued at A\$1.4 million to acquire the license. Our area of focus remained the autoimmune diseases area until October 2001 when we licensed a series of patents from the Central Sydney Area Health Service, or CSAHS, covering new treatments for chronic lung diseases and for the measurement of lung function. Our license with the ANU requires us to pay royalties based on sales revenue for products incorporating the licensed technology. Our current lead projects in the autoimmune area are not dependent on the technology licensed from the ANU. Our license agreement with the CSAHS requires us to pay royalties based on gross profit on product sales for products incorporating the licensed technology. Aridol and Bronchitol are derived from the CSAHS license.

We have incurred net losses since our inception. We recognized a net loss of A\$2.1 million, A\$6.6 million and A\$10.4 million in the years ended June 30, 2003, 2004 and 2005, respectively. Our accumulated losses from inception to June 30, 2005 are A\$20.9 million. We expect to incur increasing losses in the foreseeable future as we conduct clinical trials of our product candidates, expand our organization and prepare for the commercial launch of our products upon regulatory approval. We do not expect to generate revenue until we obtain regulatory approval to market and sell Aridol and sales of Aridol have commenced.

### **Research and Development**

Our research and development expenses consist primarily of salaries and related employee benefits, costs associated with our clinical trials, non-clinical activities such as toxicology testing and scale-up synthesis, regulatory activities, the manufacture of material for clinical trials, development of manufacturing processes and research-related overhead expenses. Our most significant costs are for clinical trials, preclinical development and regulatory filings. These expenses include regulatory consultants, clinical supplies and payments to external vendors such as hospitals and investigators. We expense all research and development costs as they are incurred. We expect our research and development expenses to increase significantly in the future as we continue to move our product candidates through the development pipeline.

We classify our research and development expenses into four components:

1. Our research unit based at the John Curtin School of Medical Research within the Australian National University, which is focused on autoimmune diseases.
2. Our preclinical development group located at Frenchs Forest, which is managing the outsourced safety/toxicology studies of the Aridol and Bronchitol products and the preclinical development of lead compounds in the autoimmune area (PXS25/64 and PXS2076).
3. Our clinical trials group located at Frenchs Forest, which designs and monitors our clinical trials.
4. Our Australian Therapeutic Goods Administration, or TGA, registered manufacturing facility at Frenchs Forest focused on producing material for clinical trials and developing enhanced manufacturing processes. It is therefore classified as a research and development expenditure.

We have spent, before the recognition of any research grants received, approximately A\$0.7 million, A\$1.4 million and A\$1.3 million and A\$5.8 million on our autoimmune research and development activities in the years ended June 30, 2003, 2004, and 2005, and the period from inception to June 30, 2005, respectively. Our autoimmune research and development activities have been primarily directed at the diseases of multiple sclerosis and rheumatoid arthritis, and have explored a range of approaches. Our current autoimmune product candidates for these diseases are PXS25/64 and PXS2076, respectively. We have spent, before the recognition of any research grants received, approximately A\$1.0 million, A\$4.5 million, A\$7.7 million and A\$13.4 million on our respiratory research and development activities in the years ended June 30, 2003, 2004, and 2005, and the period from inception to June 30, 2005, respectively. Our respiratory research and development activities have been directed at asthma, cystic fibrosis and the chronic obstructive pulmonary diseases of bronchiectasis and chronic bronchitis, and have been exclusively focused on the development of our two current respiratory product candidates of Aridol and Bronchitol. Research and development expenses for these periods, both before and after the recognition of research grants received, are presented in additional detail in the following table:

	Year ended June 30,			Period from inception (May 29, 1998) to June 30, 2005
	2003	2004	2005	
	A\$	A\$	A\$	A\$
	(in thousands)			
Total research and development expenses before research grants received	\$ 1,676	\$ 5,911	\$ 9,017	\$ 19,215
Research grants recognized against related expenses	(751)	(1,105)	(1,132)	(4,551)
<b>Net research and development</b>	<b>\$ 925</b>	<b>\$ 4,806</b>	<b>\$ 7,885</b>	<b>\$ 14,664</b>
	(unaudited)	(unaudited)	(unaudited)	(unaudited)
Autoimmune research and development	\$ 668	\$ 1,409	\$ 1,276	\$ 5,816
Respiratory research and development	1,008	4,502	7,741	13,399
	<b>\$ 1,676</b>	<b>\$ 5,911</b>	<b>\$ 9,017</b>	<b>\$ 19,215</b>



We expect to continue to incur significant costs in the foreseeable future as we pursue these activities. Although we cannot accurately forecast or reasonably estimate the additional costs that will be required to complete all of these activities, or the exact timing for their completion due to the potential failure risks and other uncertainties inherent in the development of new drugs, such as unsuccessful clinical trials, unsuccessful development and/or commercialization and delayed regulatory approvals, amongst others, we are able (except as noted below), in relation to the following clinical trials where the trial protocols have been finalized and negotiations with clinical research organizations and participating trial sites are sufficiently advanced, to reasonably estimate the costs and timeframes of the next anticipated milestones described below:

- The cost to complete our U.S. Phase III study of Aridol used for the diagnosis and management of asthma is currently estimated to be approximately A\$4 million. We commenced this trial during the fourth quarter of calendar 2005 and we expect to complete this trial in the second half of calendar 2006.
- The cost to complete our Australian Phase II study of Aridol used to predict chronic obstructive pulmonary disease, or COPD, patient responsiveness to inhaled steroids is currently estimated to be approximately A\$1 million. We commenced dosing for this trial in September 2005 and we expect to complete this trial in the second half of calendar 2006.
- The cost to complete our Phase II dose-ranging study of Bronchitol for cystic fibrosis is currently estimated to be approximately A\$1 million. We commenced this trial during the fourth quarter of calendar 2005 and we expect to complete this trial in the first half of calendar 2006.
- The cost to complete our Phase II trial of Bronchitol for cystic fibrosis versus Pulmozyme<sup>®</sup> is currently estimated to be approximately A\$1 million. We commenced enrollment of this trial during the fourth quarter of calendar 2005 or first quarter of calendar 2006 and we expect to complete this trial in calendar 2007.
- The cost to complete our European and Australian Phase III trial of Bronchitol for bronchiectasis is currently estimated to be approximately A\$6 million. We expect to commence this trial, the first of two planned for this indication, during the fourth quarter of calendar 2005 or the first quarter of calendar 2006 and we expect to complete this trial in calendar 2007.

We expect to commence our multi-national Phase III trials of Bronchitol for cystic fibrosis during the first half of calendar 2006. We expect to commence our Phase II trial of Bronchitol for chronic bronchitis during calendar 2006. We expect to commence our Phase III trial of Bronchitol for bronchiectasis in the U.S. during the middle of calendar 2006. The cost and timeframe to complete these trials cannot be reasonably estimated at this time.

We do not expect to complete the U.S. clinical trials for Aridol before the end of 2005, or any of the Bronchitol research and development projects before the end of 2006 and, therefore, we do not expect to receive any sales revenues in the U.S. prior to the completion of these projects. We anticipate that we will make determinations as to which research and development projects to pursue and how much funding to direct to each project on an on-going basis in response to the scientific and clinical success of each product candidate and available funds.

We have two research grants with the Commonwealth of Australia that assist us in funding certain of the research programs.

Under our AusIndustry P3 Pharmaceuticals Partnerships Program funding deed with the Commonwealth of Australia, subject to certain conditions, the Commonwealth of Australia may, as of June 30, 2005, pay us a total amount of A\$4.5 million between July 2005 and June 2008 for eligible pharmaceutical research and development activities undertaken by us in relation to the development of new treatments for autoimmune diseases and the development of new treatments for chronic respiratory diseases. For details regarding this research grant, see "Item

#### 10. Additional Information – C. Material Contracts - AusIndustry P3 Pharmaceuticals Partnerships Program Funding Deed.”

Similarly, under our Start Grant Program funding deed with the Commonwealth of Australia, subject to certain conditions, the Commonwealth of Australia has provided us with a grant of 50% of our eligible expenditures on a project for the development of a new treatment for cystic fibrosis up to a maximum grant amount of A\$3.0 million payable over the period from inception to December 31, 2005. For details regarding this research grant, see “Item 10. Additional Information – C. Material Contracts– Research and Development Start Program Grant Agreement.”

#### ***General and Administrative***

General and administrative expenses consist primarily of salaries and related expenses and professional services fees and includes accounting, administration, office and public company costs. We anticipate that general and administrative expenses will increase as a result of the expected expansion of our operations, facilities and other activities associated with the planned expansion of our business. As an Australian listed company also recently listed in the U.S., we operate in an increasingly demanding regulatory environment which requires us to comply with the Sarbanes-Oxley Act of 2002 and the related rules and regulations of the Securities and Exchange Commission, expanded disclosures, accelerated reporting requirements and more complex accounting rules. Responsibilities required by the Sarbanes-Oxley Act include establishing corporate oversight and adequate internal control. We also expect to incur additional costs for insurance and other professional fees associated with the quotation of our ordinary shares in the form of ADSs and becoming a public company in the U.S.

#### ***Commercial***

Our commercial expenses consist of salaries and professional fees related to the commercial launch of Aridol in Australia, Europe and the U.S. We anticipate that commercial expenses will increase as we launch Aridol, and will expand to include other selling and marketing costs.

#### **Critical Accounting Policies and Estimates**

Our management’s discussion and analysis of our financial condition and results of operations are based on our financial statements which have been prepared in accordance with accounting principles generally accepted in the U.S. The preparation of financial statements in conformity with accounting principles generally accepted in the U.S. requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, disclosures of contingent assets and liabilities at the date of the financial statements and the reported amounts of revenues and expenses during the reporting period. Significant management judgment is required to make estimates in relation to: the carrying value and recoverability of intangible assets; the recoverability of deferred income taxes; stock-based compensation; the accruing of liabilities for clinical and preclinical development activities; and research development and grants. Actual results could differ from those estimates. Our significant accounting policies are more fully described in Note 3 to our financial statements appearing elsewhere in this annual report. The following accounting policies are important in fully understanding and evaluating our reported financial results.

#### ***Intangible Assets***

We review our capital assets, including patents and licenses, for impairment whenever events or changes in business circumstances indicate that the carrying amount of the assets may not be fully recoverable. In performing the review, we estimate undiscounted cash flows from products under development that are covered by these patents and licenses. An impairment loss would be recognized when estimated undiscounted future cash flows expected to result from the use of the asset and its eventual disposition is less than the carrying amount of the asset. Impairment, if any, is measured as the amount by which the carrying amount of the asset exceeds its fair value. Impairment, if any, is assessed using discounted cash flows. Related patents are grouped in estimating future cash flows to determine whether patents are impaired and in measuring the amount of the impairment. Through June 30, 2005, there have been no such impairments.

### ***Income Taxes***

We apply Statement of Financial Accounting Standards No. 109 – Accounting for Income Taxes (SFAS 109) which establishes financial accounting and reporting standards for the effects of income taxes that result from our activities during the current and preceding years. Deferred tax assets and liabilities are recognized for the future tax consequences attributable to temporary differences between the financial statement carrying amounts of assets and liabilities and their respective tax bases. Deferred tax assets also are recognized for credit carryforwards. Deferred tax assets and liabilities are measured using the enacted rates applicable to taxable income in the years in which the temporary differences are expected to reverse and the credits are expected to be used. The effect on deferred tax assets and liabilities of a change in tax rates is recognized in income in the period that includes the enactment date. An assessment is made as to whether or not it is more likely than not that deferred tax assets are recoverable. This assessment includes anticipating future taxable income and our tax planning strategies and is made on an ongoing basis. Consequently, future material changes in the valuation allowance are possible.

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

We account for stock-based employee compensation arrangements using the fair value based method as prescribed in accordance with the provisions of Statement of Financial Accounting Standards No. 123 – Accounting for Stock-Based Compensation (SFAS 123). The terms of option issues are determined by the Board. Options are generally granted for no consideration, have a life of ten years and generally vest equally over a four year period. For options granted after January 1, 2003, the annual vesting is subject to approval by the Remuneration and Nomination Committee of the Board. The Committee gives its approval for vesting based on the achievement of individual employee's personal annual objectives. In accordance with SFAS 123, the fair values of the option grants are estimated on the date of each grant using the Black-Scholes option pricing model, which requires the use of certain significant assumptions. Changes to these assumptions, which include volatility, risk free interest rate, dividend yield, expected life, exercise price and the market price, could cause a material difference in the fair value of the options and the amount of compensation expense recognized.

### ***Clinical Development Expenses***

Clinical trial costs are a component of research and development expense. These expenses include fees paid to participating hospitals and other service providers, which conduct certain product development activities on our behalf. Depending on the timing of payments to the service providers and the level of service provided, we record prepaid or accrued expenses relating to these costs. Accrued clinical development liabilities are separately disclosed on the balance sheet.

These prepaid or accrued expenses are based on estimates of the work performed under service agreements, milestones achieved, patient enrollment and experience with similar contracts. Changes in these estimates are recognized in income in the period of change. Amounts recorded as changes of estimates have been immaterial, being A\$0, A\$0 and A\$0 and A\$0 for the years ended June 30, 2003, 2004 and 2005, and since inception to June 30, 2005, respectively. Subsequent to June 30, 2005, there were no changes in the estimates in relation to the accrued clinical development liabilities recorded at June 30, 2005.

### ***Research and Development Grants***

We receive grant funding under government research grant agreements to undertake work on the applicable grant projects, including the purchase of plant and equipment. In order to receive the grant funding, our existing grant agreements require us to incur specified eligible expenditure in the conduct of the applicable grant project. There are circumstances where grant funding may not be payable and there are certain limited circumstances, such as when we fail to use our best endeavors to commercialize the project within a reasonable time of completion of the project or upon termination of a grant due to our breach of the agreement or our insolvency, where we may be required to repay some or all of the research grants. Grants in relation to expenditure are recognized against the related research and development expenses as and when the relevant research expenditure is incurred. Grants related to plant and equipment are recognized against the acquisition costs of the related plant and equipment as and when the related assets are purchased. Grants received in advance of incurring the relevant expenditure are treated as deferred research grants and included in Other Accrued Liabilities on the balance sheet as we do not control the monies until the relevant expenditure has been incurred.

## Results of Operations

### Comparison of fiscal years ended June 30, 2004 and June 30, 2005

*Research and Development Expenses.* Research and development expenses for fiscal 2004 and 2005, respectively, are shown in the table below net of research grants received.

	Years ended June 30,	
	2004	2005
	A\$	A\$
	(in thousands)	
Total research and development expenses	\$ 5,911	\$ 9,017
Related research grants	(1,105)	(1,132)
Net research and development expenses	\$ 4,806	\$ 7,885

Grant revenue in fiscal 2004 and fiscal 2005 relates to an Australian Government R&D Start Grant awarded to us in June 2003 to develop new treatments for cystic fibrosis. The grant is payable based on underlying expenditure on the research project, which has been at consistent levels in fiscal 2004 and fiscal 2005. The grant runs until December 31, 2005. There are certain limited circumstances where we may be required to repay grant funding.

Total research and development expenses increased from A\$5.9 million in fiscal 2004 to A\$9.0 million in fiscal 2005. There are four components to the research and development expenses:

1. Our research unit based at the John Curtin School of Medical Research within the Australian National University accounted for approximately 8 percent of our total research and development expenditure in the current year. The research unit is focused on autoimmune diseases. The level of expenditure in fiscal 2005 increased by approximately 8 percent or A\$0.1 million, compared to fiscal 2004.
2. Our preclinical development group located at our Frenchs Forest facility accounted for approximately 16 percent of our total research and development expenditure in the current fiscal year and increased by approximately 7 percent or A\$0.1 million compared to fiscal 2004. This group is managing the outsourced safety/toxicology studies of the Aridol and Bronchitol products and the preclinical development of lead compounds in the autoimmune area (PXS25 and PXS2076). Approximately 60 percent of expenditure in fiscal 2005 related to the Aridol and Bronchitol studies. This area of research accounted for approximately 3 percent or A\$0.1 million of the increase in overall research and development expenditure during fiscal 2005.
3. Our clinical group located at our Frenchs Forest facility accounted for approximately 51 percent of our total research and development expenditure in fiscal 2005 and increased by approximately 82 percent or A\$2.1 million compared to fiscal 2004. The clinical group designs and monitors the clinical trials run by us. The majority of the expenditures of this group are directed at hospitals and other services related to the conduct and analysis of clinical trials. This significant increase in expenditure reflects the number of clinical trials ongoing during fiscal 2005. There were three trials ongoing during the period, including the 646 patient Aridol Phase III trial, and a number of trials in planning. This area of research accounted for approximately 66 percent or A\$2.1 million of the increase in overall research and development expenditure during fiscal 2005.
4. Our TGA registered manufacturing facility at Frenchs Forest is focused on producing material for clinical trials and developing enhanced manufacturing processes. It is therefore classified as a research and development expenditure. Manufacturing accounted for approximately 25 percent of our total research and development expenditure in fiscal 2005 and increased by approximately 67 percent or A\$0.9 million compared to fiscal 2004, reflecting manufacturing performance/yield innovation and product stability studies required to support registration applications. This area of expenditure accounted for approximately 30 percent or A\$0.9 million of the increase in overall research and development expenditure during fiscal 2005.

*General and Administrative Expenses.* General and administrative expenses increased from A\$2.2 million in fiscal 2004 to A\$3.1 million in fiscal 2005. The fiscal 2005 expenses include increased public company costs of approximately A\$0.3 million. A number of these costs were only present for the seven months of fiscal 2004 after our initial public offering in Australia in November 2003. However, the large increase in administration expenses is mainly attributable to costs of A\$0.9 million incurred in preparing the SEC filings necessary for us to list on the Nasdaq National Market.

*Commercial expenses.* Commercial expenses were A\$0 in fiscal 2004 compared to A\$0.8 million in fiscal 2005 as the commercial group was established at the beginning of fiscal 2005. It is preparing for the launch of Aridol in Australia and Europe.

*Stock Compensation Expenses.* Stock compensation expenses decreased from A\$0.5 million in fiscal 2004 to A\$0.3 million in fiscal 2005, reflecting the full year amortization in fiscal 2004 of a large grant of stock options issued in May 2003 following the employment of a number of employees, executives and directors.

*Interest and Other Income and Expense.* Interest and other income increased from A\$1.1 million in fiscal 2004 to A\$1.7 million in fiscal 2005. We started fiscal 2005 with A\$25.1 million of cash and cash equivalents on which interest was earned. This was increased by A\$19 million as a result of our private placement and share purchase plan in November and December 2004. By contrast, we started fiscal 2004 with A\$7.4 million of cash and cash equivalents on which interest was earned which increased by A\$22.9 million due to our Australian initial public offering in November 2003. The increase in interest income, while mainly attributable to the greater level of funds invested during the year, was to a lesser extent the result of a slightly higher prevailing interest rate in fiscal 2005.

*Net Loss.* Net loss increased from A\$6.6 million in fiscal 2004 to A\$10.4 million in fiscal 2005. The significant increase in operating expenses discussed above was only partly offset by the increase in interest and other income.

*Basic and diluted net loss per share.* Basic and diluted net loss per share decreased from A\$0.09 in fiscal 2004 to A\$0.08 in fiscal 2005 due to the issue of 26.4 million new ordinary shares in a share placement and share purchase plan in November and December 2004, and the full year effect on the calculation of the 50 million shares issued at the time of the Australian initial public offering in November 2003.

#### **Comparison of fiscal years ended June 30, 2003 and June 30, 2004**

*Research and Development Expenses.* Research and development expenses for fiscal 2003 and 2004, respectively, are shown in the table below net of research grants received.

	Years ended June 30,	
	2003	2004
	A\$	A\$
	(in thousands)	
Total research and development expenses	\$ 1,676	\$ 5,911
Related research grants	(751)	(1,105)
<b>Net research and development expenses</b>	<b>\$ 925</b>	<b>\$ 4,806</b>

In addition, we recognized A\$0.2 million research grant funding against the acquisition cost of related plant and equipment in fiscal 2003.

Substantially all of the fiscal 2004 grant revenue relates to an Australian Government R&D Start Grant awarded to us in June 2003 to develop new treatments for cystic fibrosis. In fiscal 2003, grant revenue related to a number of government grants for projects in both the autoimmune and respiratory areas, which were completed during fiscal 2003, except for the cystic fibrosis R&D Start Grant. The amount of research expenditure on projects funded by government grants was greater in fiscal 2004 than fiscal 2003. There are certain limited circumstances where we may be required to repay grant funding. For a summary of the grants see "Item 10. Additional Information – C. Material Contracts."

Total research and development expenses increased from A\$1.7 million in fiscal 2003 to A\$5.9 million in fiscal 2004. There are four components to the research and development expenses:

1. Our research unit based at the John Curtin School of Medical Research within the Australian National University, which is focused on autoimmune diseases. The level of expenditure in fiscal 2004 for this research unit has not changed materially from fiscal 2003, decreasing by A\$33,000.
2. Our preclinical development group, which relocated from our Canberra office to our Frenchs Forest office in January 2004. This group is managing the outsourced safety/toxicology studies of the Aridol and Bronchitol products and the preclinical development of lead compounds in the autoimmune area (PXS25 and PXS2076). This area of expenditure accounted for approximately 30 percent or A\$1.3 million of the increase in overall research and development expenditure reflecting the initiation of work in these areas.
3. Our clinical trials group, which also relocated from Canberra to Frenchs Forest in January 2004. This internal clinical trial group designs and monitors the clinical trials run by us. The majority of the expenditures of this group are directed at hospitals and other services related to the conduct and analysis of clinical trials. During fiscal 2004, expenditure commenced on three clinical trials. Approximately 50 percent or A\$2.1 million of the increase in overall research and development expenditure is attributable to the increased expenditure on clinical trials.
4. Our TGA registered manufacturing facility at Frenchs Forest is focused on producing material for clinical trials and developing enhanced manufacturing processes. It is therefore classified as a research and development expenditure. This area of expenditure also accounted for approximately 20 percent or A\$0.9 million of the increase in overall research and development expenditure, reflecting particularly the increased activity in clinical trials.

*General and Administrative Expenses.* General and administrative expenses increased from A\$1.0 million in fiscal 2003 to A\$2.2 million in fiscal 2004. Until November 2002, the level of general and administrative services we required was provided by outside providers, and we did not have our own premises. Since November 2002, when we first leased our facilities at Frenchs Forest, we have also employed the staff required to establish administrative capabilities. In fiscal 2004, we therefore incurred our first full fiscal year of general and administrative expenses, resulting in increases of approximately A\$0.2 million in lease rental costs for our facilities and A\$0.6 million in salaries. The salary increases also includes the payment in fiscal 2004 of senior management incentive payments for both fiscal 2003 and 2004. Our listing on the Australian Stock Exchange in November 2003 resulted in costs associated with being a listed public company of approximately A\$0.2 million. During fiscal 2004, we also incurred additional costs of A\$0.1 million to relocate a number of staff members to Sydney.

*Stock Compensation Expenses.* Stock compensation expenses increased from A\$0.4 million in fiscal 2003 to A\$0.5 million in fiscal 2004, reflecting the full year amortization of a large grant of stock options in May 2003 following the employment of a number of employees, executives and directors.

*Interest and Other Income and Expense.* Interest and other income increased from A\$0.3 million in fiscal 2003 to A\$1.1 million in fiscal 2004. We started fiscal 2004 with A\$7.4 million of cash and cash equivalents on which interest was earned. The A\$22.9 million raised in our Australian initial public offering in November 2003 added significantly to these invested funds. By contrast, we started fiscal 2003 with A\$751,000 of cash and cash equivalents. This amount was increased as a result of a A\$9.4 million private venture capital equity round in late August 2002. The increase in interest income, while mainly attributable to the greater level of funds invested during the year, was to a lesser extent the result of a board decision to invest in higher yielding securities and also rising interest rates.

*Net Loss.* Net loss increased from A\$2.1 million in fiscal 2003 to A\$6.6 million in fiscal 2004. The significant increase in operating expenses discussed above was only partly offset by the increase in interest and other income.

*Basic and diluted net loss per share.* Basic and diluted net loss per share decreased from A\$0.19 in fiscal 2003 to A\$0.09 in fiscal 2004 due to the conversion of 46,816,000 convertible redeemable preference shares to

ordinary shares and issue of 50 million new ordinary shares at the time of the Australian initial public offering in November 2003.

## **B. Liquidity and Capital Resources**

### **Liquidity and Capital Resources**

Since our inception, our operations have mainly been financed through the issuance of equity securities and convertible redeemable preference shares. Additional funding has come through research grants, interest on investments, and rent received from a former subtenant. Through June 30, 2005, we had received net cash proceeds from the following: (a) A\$41.9 million from the issuance of ordinary shares; (b) A\$11.4 million from the issuance of convertible redeemable preference shares; and (c) approximately A\$4.7 million in research grants. We have incurred significant losses since our inception. We incurred losses of A\$2.1 million, A\$6.6 million and A\$10.4 million and A\$20.9 million in fiscal 2003, 2004 and 2005, and since inception to June 30, 2005, respectively. As of June 30, 2005, we had cash and cash equivalents of A\$33.3 million and additionally have ongoing research grants with a total of A\$5.0 million of funding potentially available.

For fiscal 2005, we used net cash of A\$9.3 million for operating activities. This consisted of a net loss for the period of A\$10.4 million, which included A\$0.6 million of non-cash depreciation and amortization, and non-cash stock option expense of A\$0.3 million, and other working capital movements of A\$0.3 million. Net cash used in investing activities during fiscal 2005 was A\$1.6 million, which included purchase of plant and equipment of A\$1.3 million reflecting the manufacturing expansion. Net cash provided by financing activities during fiscal 2005 was A\$19.0 million resulting from the issue and sale of our ordinary shares in a share placement to Australian qualified institutional and sophisticated investors, and a share purchase plan available to existing shareholders.

For fiscal 2004, we used net cash of A\$4.8 million for operating activities. This consisted of a net loss for the period of A\$6.6 million, which included A\$0.6 million of non-cash depreciation and amortization, non-cash stock option expense of A\$0.5 million, and significant accrued clinical development liabilities at June 30, 2004 of A\$0.8 million. Net cash used in investing activities during fiscal 2004 was A\$0.4 million, which included purchases of plant and equipment of A\$0.4 million. Net cash provided by financing activities during fiscal 2004 was A\$22.9 million resulting from the issue and sale of our ordinary shares at the time of our initial public offering and listing in Australia.

For fiscal 2003, we used net cash of A\$1.3 million for operating activities. This consisted of a net loss for the period of A\$2.1 million, which included A\$0.3 million of non-cash depreciation and amortization and non-cash stock option expense of A\$0.4 million. Net cash used in investing activities during fiscal 2003 was A\$1.4 million, which included purchases of plant and equipment for A\$1.3 million primarily related to the establishment of the manufacturing facility and headquarters at Frenchs Forest. Net cash provided by financing activities during fiscal 2003 was A\$9.4 million resulting from the issue and sale of our "B" class convertible redeemable preference shares.

At June 30, 2005, we had cash and cash equivalents of A\$33.3 million as compared to A\$25.1 million as of June 30, 2004. This overall increase was primarily due to the receipt of net proceeds of A\$19.0 million related to the issue and sale of our ordinary shares in a share placement to Australian qualified institutional buyers and sophisticated investors, and a share purchase plan available to existing shareholders.

At September 30, 2005, we had cash and cash equivalents on an unaudited basis equal to A\$30.0 million.

We believe that our cash and cash equivalents will be sufficient to meet our capital requirements for at least the next 12 months. However, our forecast of the period of time through which our financial resources will be adequate to support our operations is a forward-looking statement that involves risks and uncertainties, and actual results could vary materially. If we are unable to raise additional capital when required or on acceptable terms, we may have to significantly delay, scale back or discontinue one or more of our clinical trials or our operations.

We expect to continue to incur substantial losses. Our future capital requirements are difficult to forecast and will depend on many factors, including:

- the costs of establishing sales, marketing and distribution capabilities;

- the scope, results and timing of preclinical studies and clinical trials;
- the costs and timing of regulatory approvals; and
- the cost of filing, prosecuting, defending and enforcing any patent claims and other intellectual property rights.

We anticipate that a substantial portion of our capital resources and efforts in the foreseeable future will be focused on:

- to fund clinical development for Bronchitol in patients with cystic fibrosis;
- to fund clinical development of Aridol for management of asthma and COPD;
- prepare for the commercial launch of Aridol for the management of asthma;
- to fund clinical development for Bronchitol in patients with bronchiectasis and chronic bronchitis;
- to fund pre-clinical development of our product pipeline; and
- to fund further expansion of our manufacturing facility.

### **Quantitative and Qualitative Disclosures about Market Risk**

Our exposure to market risk is limited to interest income sensitivity, which is affected by changes in the general level of Australia interest rates, particularly because the majority of our investments are in cash and cash equivalents. The primary objective of our investment activities is to preserve principal while at the same time maximizing the income we receive without significantly increasing risk. Our investment portfolio is subject to interest rate risk and will fall in value in the event market interest rates increase. Due to the short duration of our investment portfolio, we believe an immediate 10% change in interest rates would not be material to our financial condition or results of operations. We do not have any foreign currency or derivative financial instruments.

### **Income Taxes**

As of June 30, 2005, we had net operating loss carry forwards of A\$21.6 million (A\$8.9 million as of June 30, 2004). While these losses do not expire, our utilization will depend upon our ability to derive future taxable income of a nature and of an amount sufficient to enable the deduction for the losses to be realized, our continued compliance with the conditions for deductibility imposed by tax legislation, and the absence of changes in tax legislation that adversely affect our ability to utilize the loss.

As of June 30, 2004 and 2005, we did not record a benefit for the deferred tax assets because realization of the deferred tax assets was not more likely than not and, accordingly, a valuation allowance is provided to completely offset the deferred tax assets.

### **Recently Issued Accounting Announcements**

In December 2004, the Financial Accounting Standards Board issued Statement of Financial Accounting Standards No. 123 revised – Share-Based Payment (SFAS 123R), an amendment of Statements No. 123 (SFAS 123) and 95 (SFAS 95) that addressed the accounting for share-based payment transactions in which an enterprise receives employee services in exchange for either equity instruments of the enterprise or liabilities that are based on the fair value of the enterprise's equity instruments or that may be settled by the issuance of such equity instruments. This statement eliminates the ability to account for share-based compensation transactions using the intrinsic value method as prescribed by Accounting Principles Board Opinion No. 25 – Accounting for Stock Issued to Employees (APB 25), and requires that such transactions be accounted for using a fair-value-based method and recognized as expenses in the statement of operations. As of the required effective date, the standard requires that the modified prospective method be used, which requires that the fair value of new awards granted on or following the effective date (plus unvested awards as of the effective date) be expensed over the vesting period. In addition, the statement encourages the use of the "binomial" approach to value stock options, which differs from the Black-Scholes option pricing model that we currently use. Further, in March 2005, SEC has issued Staff Accounting Bulletin 107 (SAB



107) providing guidance on the application of SFAS 123R. Further, as per a new rule approved by SEC in April 2005, SFAS 123R will be effective for public companies' annual, rather than interim, periods that begin after June 15, 2005. The adoption of SFAS 123R and SAB 107 is not expected to have a significant impact on our statement of operations as we currently expense the fair value of our stock option grants.

In May 2005, the Financial Accounting Standards Board issued Statement of Financial Accounting Standards No. 154 – Accounting Changes and Error Corrections (SFAS 154), a replacement of Accounting Principles Board Opinion No. 20 (APB 20) and Statement No. 3 (SFAS 3), which previously addressed accounting changes. SFAS 154 establishes, unless impracticable, retrospective application as the required method for reporting a change in accounting principle in the absence of explicit transition requirements specific to the newly adopted accounting principle. SFAS 154 also provides guidance for determining whether retrospective application of a change in accounting principle is impracticable and for reporting a change when retrospective application is impracticable. SFAS 154 carries forward without change the guidance in APB 20 for reporting the correction of an error in previously issued financial statements. SFAS 154 will be effective for accounting changes and corrections of errors made in fiscal years beginning after December 15, 2005. The adoption of this standard will not have a material impact on our financial statements.

C. See Item 5A - “Operating Results – Research and Development.”

D. See Item 5A – “Operating Results” and Item 5B – “Liquidity and Capital Resources.”

#### E. Off-Balance Sheet Arrangements

We have no off-balance sheet arrangements that have or are reasonably likely to have a current or future effect on our financial condition, changes in financial condition, revenues or expenses, results of operations, liquidity, capital expenditures or capital resources that are material to investors.

#### F. Contractual Obligations and Commitments

The following table summarizes financial data for our contractual obligations and other commercial commitments, including interest obligations, as of June 30, 2005 (in thousands):

Contractual Obligations	Payments due by Period				
	Total	Less than 1 year	1 - 3 years	3 - 5 years	More than 5 years
	A\$	A\$	A\$	A\$	A\$
Long-Term Debt Obligations	\$—	\$ —	\$ —	\$ —	\$ —
Capital (Finance) Lease Obligations	—	—	—	—	—
Operating Lease Obligations	322	322	—	—	—
Purchase Obligations	—	—	—	—	—
Other Long-Term Liabilities Reflected on our Balance Sheet under the GAAP of the primary financial statements	—	—	—	—	—
<b>Total</b>	<b>\$322</b>	<b>\$ 322</b>	<b>\$ —</b>	<b>\$ —</b>	<b>\$ —</b>

The contractual summary above reflects only payment obligations that are fixed and determinable. We have additional contractual payment obligations that are contingent on future events. Our operating lease obligations relate to the lease for our headquarters in Frenchs Forest, Sydney which expires in June 2006 but contains an option to renew for a further five years thereafter. We also have agreements with clinical sites, and contract research organizations, for the conduct of our clinical trials and other research activities.

## ITEM 6. DIRECTORS, SENIOR MANAGEMENT AND EMPLOYEES

### Management

#### A. Directors and Senior Management

The following table presents information about our executive officers and directors as of September 30, 2005.

<u>Name</u>	<u>Age</u>	<u>Position</u>
Alan D. Robertson, Ph.D.	49	Chief Executive Officer and Managing Director
Brett Charlton, Ph.D.	49	Medical Director and Director
William B. Cowden, Ph.D.	56	Chief Scientific Officer
John F. Crapper	53	Chief Operations Officer
Ian A. McDonald, Ph.D.	58	Chief Technical Officer
David M. McGarvey, C.A., C.P.A.	49	Chief Financial Officer, Company Secretary
Gary J. Phillips	44	Head of Commercial Development
Denis M. Hanley(1)(2)	58	Chairman of the Board
Carmel J. Hillyard, Ph.D.(2)	56	Director
Charles P.H. Kiefel(1)	50	Director
Malcolm J. McComas(1)	51	Director
Brigitte H. Smith(2)	38	Director

(1) Member of the Audit Committee.

(2) Member of the Remuneration and Nomination Committee.

The business address for our executive officers and directors is c/o Pharmaxis Ltd, Unit 2, 10 Rodborough Road, Frenchs Forest, NSW Australia 2086.

#### Executive Officers and Directors

*Alan D. Robertson, Ph.D.*, has been our Chief Executive Officer since December 1999 and a member of our board of directors since July 2000. Dr. Robertson has more than two decades of experience in drug discovery and product development with leading pharmaceutical companies, including spending 8 years with Wellcome plc in London and thereafter with two Australian companies, Faulding Ltd and Amrad Ltd. Dr. Robertson has been actively involved in the discovery, development and marketing of various compounds, including new treatments for migraine and cardiovascular disease. Dr. Robertson is the co-inventor of 18 patents and author of more than 35 scientific papers, and was the inventor of the migraine therapeutic Zomig that is marketed worldwide by AstraZeneca. Dr. Robertson holds a B.Sc. and a Ph.D. in Synthetic Organic Chemistry from the University of Glasgow.

*Brett Charlton, Ph.D.*, is a co-founder of Pharmaxis and has been our Medical Director and a member of our board of directors since June 1998. Dr. Charlton is the author of more than 60 scientific papers and has over 15 years of experience in clinical trial design and management. Dr. Charlton was founding Medical Director of the National Health Sciences Centre and established its Clinical Trials Unit. Prior to joining us, Dr. Charlton held various positions with the Australian National University, Stanford University, the Baxter Centre for Medical Research, Royal Melbourne Hospital, and the Walter and Eliza Hall Institute. Dr. Charlton holds a M.B.B.S. with honors from the University of New South Wales and a Ph.D. from the University of New South Wales.

*William B. Cowden, Ph.D.*, is a co-founder of Pharmaxis and has been our Chief Scientific Officer since June 1998. Dr. Cowden has two decades of experience in researching and developing therapeutic compounds to treat cancer and infectious and inflammatory diseases. Dr. Cowden is the co-inventor of 12 patents and author of more than 150 scientific papers. Dr. Cowden has a long association with the John Curtin School of Medical Research at the Australian National University, and has held senior research positions with that university's departments of medical chemistry, experimental pathology, and cell biology and virology. Dr. Cowden is Head of the Immunopathology Research Group and directs our research into autoimmune compounds for multiple sclerosis and rheumatoid arthritis. Dr. Cowden holds a B.S. (honors) from Troy State University and a Ph.D. in Medical Chemistry from the University of Queensland.

*John F. Crapper* has been our Chief Operations Officer since July 2003. Mr. Crapper has over three decades of experience in manufacturing and operations. From 1987 to 2003, Mr. Crapper held various positions

within the Memtec Limited/Memcor organization most recently as Senior Vice-President and General Manager of Memcor International, and Managing Director of Memcor Australia Pty Ltd, a leader in the design and manufacture of microfiltration membranes and systems. During his 15 years at Memcor, Mr. Crapper managed the scale-up of manufacturing equipment and processes from the company's research and development group, created full-scale production operations, and managed the establishment of Quality Assurance and Enterprise Resource Planning systems. From 1980 to 1987, Mr. Crapper served as Operations Director of the Animal Health Division at Syntex Pharmaceutical. From 1971 to 1980, Mr. Crapper served as Production Manager at VR Laboratories, a private veterinary pharmaceutical company. Mr. Crapper holds a B.S. in Applied Chemistry from the University of Technology, Sydney and an M.B.A from Macquarie University.

*Ian A. McDonald, Ph.D.*, has been our Chief Technical Officer since April 2005. Dr. McDonald has over 25 years of experience in managing drug discovery and design teams in Europe and the U.S. From 2002 to 2004, Dr. McDonald served as Vice President of Drug Discovery at Structural GenomiX Inc. (now SGX Pharmaceuticals Inc.). From 2001 to 2002, Dr. McDonald served as Vice President of Drug Discovery at Structural Bioinformatics Inc. (now Cengent Therapeutics). From 1993 to 2000, Dr. McDonald served as Director, then Vice President of Chemistry at SIBIA Neuroscience (now part of Merck Research Laboratories) and was responsible for medicinal and bio-chemistry research. From 1978 to 1993, Dr. McDonald served in various capacities as a research chemist at Merrell Dow (now part of Sanofi-Aventis). Dr. McDonald is the co-inventor of 39 U.S. patents and co-author of 77 peer-reviewed manuscripts and book chapters. Dr. McDonald holds B.S. and Ph.D. degrees in Organic Chemistry from the University of Western Australia.

*David M. McGarvey, C.A., C.P.A.*, has been our Chief Financial Officer and Company Secretary since December 2002. Mr. McGarvey has two decades of experience in overseeing the financial affairs of different Australian companies. From 1998 to 2002, Mr. McGarvey served as Chief Financial Officer of the Filtration and Separations Group of U.S. Filter Corporation where he managed over 20 merger and acquisition transactions, including the sale of the Filtration and Separations Group to Pall Corporation in 2002. From 1985 to 1997, Mr. McGarvey served as Chief Financial Officer of Memtec Limited. While at Memtec, Mr. McGarvey oversaw the U.S. listing of Memtec on the Nasdaq National Market and the New York Stock Exchange and managed over 10 merger and acquisition transactions, including the acquisition of Memtec by U.S. Filter. From 1975 to 1985, Mr. McGarvey held various positions at PricewaterhouseCoopers. Mr. McGarvey holds a B.A. in Accounting from Macquarie University and was admitted to the Institute of Chartered Accountants in Australia in 1981, and to the Australian Society of Certified Practising Accountants in 1993.

*Gary J. Phillips* has been our Head of Commercial Development since December 2003. Mr. Phillips has over two decades of operational management experience in the pharmaceutical and healthcare industry in Europe, Asia and Australia. From 1998 to 2003, Mr. Phillips held various positions within Novartis Asia, most recently as Chief Executive Officer of Novartis Pharmaceuticals Australia Pty Ltd, where he successfully launched leading oncology and ophthalmology products and relaunched newly acquired primary care products. From 1992 to 1998, Mr. Phillips served as Chief Executive Officer at Ciba Geigy in Hungary. Mr. Phillips holds a B. Pharm. in Pharmacy with honors from Nottingham University in the U.K. and an M.B.A. from Henly Management College.

*Denis M. Hanley* has been the Chairman of our board of directors since October 2001. From 1983 to 1997, Mr. Hanley served as Chief Executive Officer of Memtec Limited, a leader in the design and manufacture of microfiltration membrane systems. From 1971 to 1982, Mr. Hanley held various positions within Baxter Healthcare, most recently as Australian Managing Director. Mr. Hanley has served on the Australian Industry Research and Development Board and various technology councils and roundtables. Mr. Hanley serves on the board of directors of Lochard Limited and CathRx Ltd, and is a member of the Australian Government's Cooperative Research Centre Committee. Mr. Hanley holds an M.B.A. with high distinction from the Harvard Graduate School of Business Administration, where he was named a Baker Scholar.

*Carmel J. Hillyard, Ph.D.*, has been a member of our board of directors since August 2002. Dr. Hillyard has more than three decades of experience in managing all aspects of drug discovery and development at life science companies. Dr. Hillyard's career extends from research in cancer and endocrinology at London University, through patenting and developing novel diagnostic technologies, to assisting entrepreneurs and early-stage life science companies. From 1997 to present, Dr. Hillyard has served as a founder and Partner at CM Capital Investments, a venture capital fund manager, where she heads the life sciences practice. Dr. Hillyard is the inventor of six patent families, and has provided strategic guidance to pharmaceutical and biotechnology companies on licensing

technology, managing collaborations and obtaining venture funding. Dr. Hillyard has also advised the Australian government on science and technology matters and is a board member of the Australian Nuclear Science and Technology Organisation. Dr. Hillyard serves on the board of directors of CathRx Ltd. Dr. Hillyard holds a joint B.Sc. in biochemistry and biology with honors from the University of London and a Ph.D. in medicine from the Royal Postgraduate Medical School in the U.K. Dr. Hillyard is a Fellow of the Academy of Technological Sciences and Engineering and the recipient of the Centenary medal from the Australian Government.

*Charles P.H. Kiefel* has been a member of our board of directors since May 2003. Mr. Kiefel has more than two decades of experience in finance and investment banking. From 1990 to 2000, Mr. Kiefel was Managing Director of Corporate Finance at ANZ Investment Bank. From 1986 to 1990, Mr. Kiefel was Director of Corporate Finance at Ord Minnett. From 1985 to 1986, Mr. Kiefel held positions at Lazard Brothers & Co. Ltd in the U.K. and Lazard Frere in New York. Mr. Kiefel serves on the board of directors of Military Superannuation & Benefits Board, Wilson HTM Charities Ltd, Hyperion Asset Management Ltd and Hyperion Holdings Limited. Mr. Kiefel has a B. Com. from the University of New South Wales and is a Fellow of the Institute of Chartered Accountants in Australia.

*Malcolm J. McComas* has been a member of our board of directors since July 2003. Mr. McComas has more than two decades of experience in investment banking, particularly in equity and debt finance, mergers and acquisitions, and privatizations. From 1999 to 2004, Mr. McComas was a director of Grant Samuel and currently serves as a consultant to their corporate advisory, property services and funds management group. During 1998, Mr. McComas served as a Managing Director at Salomon Smith Barney. From 1988 to 1997, Mr. McComas served as a Managing Director at County NatWest. Mr. McComas serves as a non-executive director of Australasian Investors Limited and non-executive chairman of Sunshine Heart Inc. Mr. McComas holds a B.Ec. and a Bachelor of Laws from Monash University.

*Brigitte H. Smith* has been a member of our board of directors since October 1999. Ms. Smith is a venture capital investor with more than ten years of experience in strategic management consulting and working with early stage technology-based businesses in the U.S. and Australia. Since 2002, Ms. Smith has served as a Managing Director of GBS Venture Partners, a specialist life science venture capital business she co-founded after completing a management buy-out from Rothschild. A former Fulbright Scholar, Ms. Smith is also an Adjunct Senior Lecturer at Melbourne Business School, where she teaches Entrepreneurial Finance. Ms. Smith holds a B.S. in Chemical Engineering with honors from the University of Melbourne, an M.A. from the Fletcher School of Law and Diplomacy and an M.B.A with honors from the Harvard Graduate School of Business Administration.

There are no family relationships between any of our executive officers or directors.

### **Composition of the Board of Directors**

We regard Messrs. Kiefel, McComas and Hanley as independent non-executive directors for purposes of applicable Australian law, as described by the Pharmaxis Corporate Governance Framework, and in accordance with the Principles of Good Corporate Governance and Best Practice Recommendations issued by the Australian Stock Exchange.

We do not regard Dr. Robertson and Dr. Charlton as independent directors under applicable Australian law as they are Pharmaxis executives. We do not consider Ms. Smith and Dr. Hillyard as independent directors under applicable Australian law because they are associated with shareholders that own a significant number of shares in Pharmaxis.

### **Scientific Advisory Board**

The members of the Pharmaxis Scientific Advisory Board play an important role advising us in their areas of expertise.

*Sandra Anderson, B.Sc., Ph.D., D.Sc., FANZSRS*, is an expert in the diagnosis and treatment of asthma. She is a world authority in the measurement, management and mechanisms of exercise-induced asthma, and has developed a variety of tests for identifying asthma, including Aridol. A prolific author and the recipient of numerous awards for her work, Dr. Anderson is Principal Hospital Scientist in the Department of Respiratory Medicine of the Royal Prince Alfred Hospital, Sydney. She is a Vice President of Asthma NSW and Co-Chairman of their Research

Advisory Committee. Dr. Anderson has served on various international taskforces and committees and is currently part of an independent panel of the International Olympic Committee Medical Commission. She is actively engaged in our development, participating in technical presentations to various opinion leaders and regulatory authorities around the world. Dr. Anderson holds a Bachelor of Science in Physiology from the University of Sydney and a Ph.D. in Medicine from the University of London.

*Norbert Berend, M.B., B.S., M.D., FRACP*, is Director of the Woolcock Institute of Medical Research at Royal Prince Alfred Hospital, Sydney and is internationally recognized for his work in chronic obstructive pulmonary disease. Dr. Berend is active in national and international peer groups, is a member of the COPD Guidelines Working Party, and serves on the Respiratory Clinical Expert Reference Committee of the NSW Department of Health. In addition, he is a Senior Investigator for the Cooperative Research Centre, or CRC, for Asthma and a Director of the CRC for Chronic Inflammatory Diseases and is the author of more than 95 publications on airways disease, emphysema and infection in COPD. Dr. Berend was a principal investigator at one site participating in the Aridol trial as well as serving on trial related safety committees.

*Malcolm Fisher, A.O., M.B., Ch.B., M.D.*, is renowned for his work in critical care medicine, having received numerous awards and being named an officer in the Order of Australia. Based in Sydney, Dr. Fisher is a Staff Specialist in the Intensive Care Unit of Royal North Shore Hospital, and Area Director of Intensive Care and Clinical Professor in Intensive Care Medicine in the Departments of Medicine and Anaesthesia at the University of Sydney. He is a past President of the World Federation of Intensive and Critical Care Medicine Societies, and its Australasian chapter, ANZICS. He is the author of two books and more than 130 scientific articles.

*Richard J.I. Morgan, C.Biol., MIBiol., DRCPATH*, has more than 25 years' experience in pharmaceutical research and development, and has been involved in the development of a large number of successful, marketed pharmaceutical products. He has held senior management positions within preclinical safety (a vital precursor to human clinical trials), including Head of Toxicology at the pharmaceutical company Wellcome and International Head of Toxicology and Preclinical Outsourcing for GlaxoWellcome (later GlaxoSmithKline). He has been responsible for evaluating the preclinical safety of more than 100 new chemical entities, ranging from anti-infectives and anti-parasitics to cancer compounds and vaccines. He currently advises U.K. and Australian companies on toxicology and preclinical discovery and development. Mr. Morgan consults to Pharmaxis on the preclinical safety aspects of developing products.

## **B. Compensation**

*Principles used to determine the nature and amount of remuneration.* As a company building an international pharmaceutical business, we require a board of directors and executive team that have both the technical capability and relevant experience to execute our business plan. We consider options to purchase ordinary shares a key tool in attracting the required talented individuals to our board of directors and executive team while staying within the fiscal constraints of a growing company. Director and executive compensation includes a mix of short and long-term components. Compensation of executive directors and other executives include a meaningful proportion that varies with individual performance. Cash incentives and the vesting of options are subject to performance assessment by the Remuneration and Nomination Committee. Performance targets primarily relate to objectives and milestones assigned to individual executives from our annual business plan. Individual performance targets are agreed by the Remuneration and Nomination Committee and the full board each year. Annual performance of each executive is reviewed by the Remuneration and Nomination Committee each year. Compensation of executive directors and other executives are reviewed each year with changes typically taking effect on January 1 each year. As non-executive directors assess individual and company performance, their remuneration does not have a variable performance related component.

*Non-executive directors.* Fees and payments to non-executive directors reflect the demands that are made on, and the responsibilities of, the directors. Non-executive directors' fees and payments are reviewed annually by the Remuneration and Nomination Committee of our board of directors. There are four components to the fees:

- a base fee, as of January 1, 2005, of A\$52,500 for the Chairman and A\$25,625 for other non-executive directors;

- an additional flat annual fee of A\$5,000, as of January 1, 2005, for non-executive directors serving on committees;
- statutory superannuation fund contributions for directors who we consider independent for purposes of Australian law, which are similar in nature to retirement fund contributions, currently equal to 9% of the non-executive's base fee per year for the non-executive directors we deem independent under applicable Australian guidelines; and
- options to purchase ordinary shares under the Pharmaxis Employee Option Plan. Options vest over approximately four years from the grant date. Note that options are not granted to Ms. Smith or Dr. Hillyard who are principals of their respective venture capital firms that manage funds which are significant shareholders of Pharmaxis.

Non-executive directors' fees (including statutory superannuation) are determined within an aggregate directors' fee pool limit, which is periodically recommended for approval by shareholders. The pool of remuneration approved by shareholders at a general meeting currently stands at a maximum of A\$300,000 per annum in total. The amount paid to non-executive directors in 2004 was A\$149,846. Subject to the requirements of Australian law, termination payments apply only to executive directors, as discussed below.

*Executive directors and executive officers.* There are four components to executive director and executive officer remuneration:

- annual base salary paid in cash or packaged at the executive's discretion within Australian fringe benefits tax guidelines as a total cost package;
- annual statutory superannuation fund contributions up to 9% of the executive's base salary;
- a variable incentive component payable annually dependent upon achievement of performance targets set and approved by the Remuneration and Nomination Committee; and
- options to purchase ordinary shares under the Pharmaxis Employee Option Plan. Options typically vest over a four-year time frame. For options granted after January 1, 2003, the number of an individual executive director's or officer's options vesting is subject to achievement of the performance targets set and approved by the Remuneration and Nomination Committee. The Remuneration and Nomination Committee may approve the vesting of all or only a portion of the relevant options. Founder options were granted in 2003 to the founding scientists, Dr. Cowden and Dr. Charlton. These options became fully vested at June 30, 2003. Sign-on options were granted to Mr. McGarvey in 2003, to Mr. Crapper and Mr. Phillips in 2004 and Dr. McDonald in 2005. Sign-on options fully vest on the first anniversary of the executive commencing employment with us.

Base pay for executive directors and executive officers is reviewed annually to ensure the executive's pay is competitive with the market. An executive's pay is also reviewed on promotion. The employment contracts for each of the above-listed executive directors and executive officers, which are described below, can be terminated immediately by us for serious misconduct, with one month's prior notice if the employee becomes mentally or physically unfit to perform or carry out their employment, with two months prior notice if the executive's position ceases to exist, and with three months prior notice without cause. No additional payments apply on termination.

## Executive Compensation

The following table presents all of the compensation awarded to, earned by or paid to each individual who served as a member of our board of directors and as an executive officer in fiscal 2005.

### Summary Compensation Table

Name	Primary				Total
	Cash salary or Directors' Fees	Cash Incentive	Super- annuation	Equity Options(1)	
	A\$	A\$	A\$	A\$	
<b>Directors</b>					
Alan D. Robertson, Ph.D.(2)	\$ 194,750	\$68,000	\$15,750	\$ 31,702	\$310,202
Denis M. Hanley	57,500*	—	5,175	13,209	75,884
Brett Charlton, Ph.D.(2)	160,750	36,000	11,700	15,851	224,301
Brigitte H. Smith	30,625*	—	—	—	30,625
Charles P.H. Kiefel	30,625*	—	2,756	8,634	42,015
Carrie J. Hillyard, Ph.D.	30,625*	—	—	—	30,625
Malcolm J. McComas	30,625*	—	2,756	9,135	42,516
<b>Executive Officers</b>					
William B. Cowden, Ph.D.	139,913	20,000	11,700	15,851	187,464
John F. Crapper	182,963	22,500	15,300	21,853	242,616
Ian A. McDonald(3)	42,628	—	3,837	10,187	56,652
David M. McGarvey	193,722	40,000	16,200	15,851	265,773
Gary J. Phillips	189,625	36,000	16,650	41,413	283,688

(1) The fair value of option grants were estimated on the date of each grant using the Black-Scholes option pricing model.

(2) Individual also serves as an executive officer.

(3) Dr. McDonald joined Pharmaxis on April 4, 2005.

\* Directors' fees.

### Stock Option Grants in Fiscal 2005

Details of options to purchase our ordinary shares provided during the fiscal year ended June 30, 2005 as remuneration to each director of Pharmaxis and each of our executive officers are set out below. When exercisable, each option is convertible into one ordinary share of Pharmaxis. Options are issued at a zero purchase price. Vesting details are set out in the subsequent table.

Name	Number of Options Granted During the Year	Exercise Price	Expiration Date
<b>Directors</b>			
—	—	\$ —	—
<b>Executive Officers</b>			
Ian A. McDonald	200,000	1.147	May 11, 2015

## Stock Option Grants since June 30, 2005

On August 5, 2005 the board of directors proposed, subsequent to a review of employee and director performance for the year ended June 30, 2005, to grant 760,000 options under the Pharmaxis Employee Option Plan to executive officers and independent directors and 194,500 options to other employees. The grant of options to directors requires shareholder approval and 335,000 of the proposed option grants were approved by shareholders at our annual meeting on November 15, 2005. The remaining 425,000 executive officer options were granted together with the 194,500 options to other employees on August 5, 2005. Details of grants to executive officers and directors are presented below. When exercisable, each option is convertible into one ordinary share of Pharmaxis. Options are issued at a zero purchase price. Vesting details for granted options are presented in the subsequent table.

Name	Number of Options Granted	Exercise Price	Expiration Date
A\$			
<i>Directors</i>			
Alan D. Robertson, Ph.D.(1)	150,000	\$ 1.79	08/04/2015
Brett Charlton, Ph.D.(1)	105,000	1.79	08/04/2015
Denis M. Hanley	40,000	1.79	08/04/2015
Charles P.H. Kiefel	20,000	1.79	08/04/2015
Malcolm. M. McComas	20,000	1.79	08/04/2015

Name	Number of Options Granted	Exercise Price	Expiration Date
A\$			
<i>Executive Officers</i>			
William B. Cowden, Ph.D.	100,000	\$ 1.79	08/04/2015
John F. Crapper	100,000	1.79	08/04/2015
Ian A. McDonald, Ph.D.	20,000	1.79	08/04/2015
David M. McGarvey	100,000	1.79	08/04/2015
Gary J. Phillips	105,000	1.79	08/04/2015

(1) Individual also serves as an executive officer.

## Stock Option Values

### Option holdings

The numbers of options to purchase our ordinary shares held at December 9, 2005 by each director of Pharmaxis and each of the executive officers are listed below. When exercisable, each option is convertible into one ordinary share of Pharmaxis. Options are issued at a zero purchase price.

Name	Number of Securities	Exercise Price	Expiration Date	Vesting
A\$				
<i>Directors</i>				
Alan D. Robertson, Ph.D.(1)	1,120,000	\$0.1250	11/30/2009	280,000 at each of June 30, 2000, 2001, 2002 and 2003(3)
	960,000	0.3125	06/30/2012	240,000 at each of June 30, 2003, 2004, 2005 and 2006(2)(3)
	150,000	1.790	08/04/2015	37,500 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
Denis M. Hanley	640,000	0.3125	08/30/2011	640,000 at August 30, 2002(3)
	400,000	0.3125	06/30/2012	100,000 at each of June 30, 2003, 2004, 2005 and 2006(3)
	40,000	1.790	08/04/2015	10,000 at each of June 30, 2006, 2007, 2008 and 2009(3)



<u>Name</u>	<u>Number of Securities</u>	<u>Exercise Price</u>	<u>Expiration Date</u>	<u>Vesting</u>
		A\$		
Brett Charlton, Ph.D.(1)	640,000	\$0.1250	11/30/2009	160,000 at each of June 30, 2000, 2001, 2002 and 2003(3)
	480,000	0.3125	06/30/2012	480,000 at June 30, 2003(3)
	480,000	0.3125	06/30/2012	120,000 at each of June 30, 2003, 2004, 2005 and 2006(1)(3)
	105,000	1.790	08/04/2015	26,250 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
Charles P.H. Kiefel	200,000	0.3125	04/30/2013	50,000 at each of June 30, 2004, 2005, 2006 and 2007(3)
	20,000	1.790	08/04/2015	5,000 at each of June 30, 2006, 2007, 2008 and 2009(3)
Malcolm. M. McComas	200,000	0.3125	07/03/2013	50,000 at each of June 30, 2004, 2005, 2006 and 2007(3)
	20,000	1.790	08/04/2015	5,000 at each of June 30, 2006, 2007, 2008 and 2009(3)
<i>Executive Officers</i>				
William B. Cowden, Ph.D.	640,000	0.1250	11/30/2009	160,000 at each of June 30, 2000, 2001, 2002 and 2003(3)
	480,000	0.3125	06/30/2012	480,000 at June 30, 2003(3)
	480,000	0.3125	06/30/2012	120,000 at each of June 30, 2003, 2004, 2005 and 2006(2)(3)
	100,000	1.7900	08/04/2015	25,000 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
John F. Crapper	480,000	0.3125	06/30/2013	480,000 at July 1, 2004 (3)
	480,000	0.3125	06/30/2013	120,000 at each of June 30, 2004, 2005, 2006 and 2007(2)(3)
	100,000	1.7900	08/04/2015	25,000 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
Ian A. McDonald, Ph.D.	50,000	1.1470	05/11/2015	April 3, 2006 (3)
	150,000	1.1470	05/11/2015	37,500 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
	20,000	1.7900	08/04/2015	5,000 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
David M. McGarvey	480,000	0.3125	06/30/2012	120,000 at each of June 30, 2003, 2004, 2005 and 2006(2)(3)
	480,000	0.3125	11/30/2012	480,000 at December 1, 2003 (3)
	100,000	1.7900	08/04/2015	25,000 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)
Gary J. Phillips	250,000	0.3760	11/30/2013	62,500 at each of June 30, 2004, 2005, 2006 and 2007(2)(3)
	250,000	0.3760	11/30/2013	250,000 at December 1, 2004 (3)
	105,000	1.7900	08/04/2015	26,250 at each of June 30, 2006, 2007, 2008 and 2009(2)(3)

- (1) Individual also serves as an executive officer.
- (2) Vesting is subject to approval of the Remuneration and Nomination Committee.
- (3) Options granted to directors and executive officers are escrowed until February 6, 2006.

### Employment Agreements

The following executive directors and executive officers have employment agreements with Pharmaxis. Each of these agreements provide for the provision of performance-related cash incentives and participation, when eligible, in the Pharmaxis Employee Option Plan. These agreements also contain certain confidentiality, intellectual property and noncompetition provisions that serve to protect our intellectual property rights and other proprietary information.

The employment agreements can only be terminated by us without notice if for cause. Should an employee become mentally or physically unfit to fulfill his duties, we may terminate his employment upon one month's advance notice. If termination is due to the elimination of such employee's position, then two months advance notice is required prior to any termination. For any other

termination without cause, we are required to provide the

employee three months advance notice. During the above noted notice periods, the employee is entitled to his base salary and other benefits. Upon termination, the employee is also entitled to payment of any accrued annual leave benefits.

In addition to their respective base salaries, each of the following executive officers may be awarded an annual performance bonus upon satisfaction of certain milestones upon the sole discretion of our Remuneration and Nomination Committee and as approved by the board of directors.

Other material terms of each of these agreements are identified below.

Alan D. Robertson, Ph.D., *Chief Executive Officer and Managing Director*

- Subject to earlier termination by us, the term of Dr. Robertson's employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Dr. Robertson enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$220,000, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation fund contributions equal to 9% of the annual base salary, or A\$19,800, per year for the benefit of Dr. Robertson.

Brett Charlton, Ph.D., *Medical Director and Director*

- Subject to earlier termination by us, the term of Dr. Charlton's employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Dr. Charlton enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$185,000, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$16,650, per year for the benefit of Dr. Charlton.

William B. Cowden, Ph.D., *Chief Scientific Officer*

- Subject to earlier termination by us, the term of Dr. Cowden's employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Dr. Cowden enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$143,500, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$12,915, per year for the benefit of Dr. Cowden.

John F. Crapper, *Chief Operations Officer*

- Subject to earlier termination by us, the term of Mr. Crapper's employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Mr. Crapper enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$187,500, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and

- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$16,875, per year for the benefit of Mr. Crapper.

Ian A, McDonald, Ph.D., *Chief Technical Officer*

- Subject to earlier termination by us, the term of Dr. McDonald's employment will last until June 30, 2007, unless the term of the employment agreement is either extended or Dr. McDonald enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$175,000, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$15,750, per year for the benefit of Dr. McDonald.

David M. McGarvey, C.A., C.P.A., *Chief Financial Officer and Secretary*

- Subject to earlier termination by us, the term of Mr. McGarvey's employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Mr. McGarvey enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$198,500, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$17,865, per year for the benefit of Mr. McGarvey.

Gary J. Phillips, *Head of Commercial Development*

- Subject to earlier termination by us, the term of Mr. Phillips' employment will last until June 30, 2008, unless the term of the employment agreement is either extended or Mr. Phillips' enters into a new employment agreement with us;
- The employment agreement provides for an annual base salary of A\$194,500, which may be subject to increase upon review annually by our Remuneration and Nomination Committee and as approved by the board of directors; and
- We are also required to make superannuation contributions equal to 9% of the annual base salary, or A\$17,505, per year for the benefit of Mr. Phillips.

### **Limitation of Liability and Indemnification Matters**

Our Constitution provides that, except to the extent prohibited by the Corporations Act 2001, each of our officers shall be indemnified out of our funds against any liability incurred by such person in his or her capacity as an officer in defending any legal proceedings, whether civil or criminal, in which judgment is given in such person's favor or where such officer is acquitted in connection with any application under the Corporations Act 2001 and relief is granted to such officer by a court.

We have entered into Deeds of Access to Documents and Indemnity agreements to indemnify our directors and executive officers and to provide contractual indemnification in addition to the indemnification provided for in our Constitution. We believe that these provisions and agreements are necessary to attract and retain qualified directors and executive officers. Our Constitution also permits us, to the extent permitted by law, to secure insurance on behalf of any officer for any liability arising out of his or her actions.

At present, there is no pending litigation or proceeding involving any of our directors, officers, employees or agents where indemnification by us will be required or permitted, and we are not aware of any threatened litigation or proceeding that may result in a claim for such indemnification.

We maintain directors' and officers' liability insurance providing for the indemnification of our directors and officers against certain liabilities incurred as a director or officer, including costs and expenses associated in successfully defending legal proceedings. We intend to continue to maintain this insurance in the future.

### **Severance and Change of Control Agreements**

Details of the severance payments owed to directors upon termination are set out above. See Item 6— "Directors, Senior Management and Employees."

### **Employee Option Plan**

Any person considered to be an employee of Pharmaxis, or any subsidiary of Pharmaxis, by our board of directors including the executive directors and the non-executive directors are eligible to participate in the Pharmaxis Employee Option Plan, but do so at the invitation of our board of directors. Under the Pharmaxis Employee Option Plan, the board of directors may issue options to purchase our ordinary shares on such terms, including the issue price, the exercise price and the vesting conditions, as it determines. The maximum number of options available to be issued under the Pharmaxis Employee Option Plan at any given time is 15% of our total issued shares and other securities convertible into shares at such time, or such number as is consistent with any Listing Rules or laws or regulations that apply to Pharmaxis.

Any vesting conditions determined by our board of directors must be satisfied before the employee options vest and become exercisable. Options are generally granted for no consideration and vest in equal tranches over a four-year period. For options granted after January 1, 2003, the annual vesting is subject to approval by the Remuneration and Nomination Committee of our board of directors. The Remuneration and Nomination Committee gives its approval for vesting based on the achievement of individual employee's personal annual objectives. If a takeover offer is made for Pharmaxis, all options which have not yet vested, vest.

When exercisable, each option issued under the Pharmaxis Employee Option Plan entitles the holder to subscribe for one fully paid ordinary share. Each ordinary share issued on exercise of an option will rank equally with all other ordinary shares then issued.

The exercise price of the employee options is set by our board of directors. Before we listed on the Australian Stock Exchange in November 2003, our board of directors set the exercise price based on its assessment of the market value of the underlying shares at the time of grant. Since listing on the Australian Stock Exchange, the exercise price is set by our board of directors as the average closing price of our ordinary shares on the Australian Stock Exchange during the five business days prior to the grant of the options.

The employee options lapse on such date as determined by the board of directors at the time of grant. If an optionholder ceases to be regarded as an employee by our board of directors, all of his or her options which have not yet vested lapse and all options which have already vested lapse, if not exercised, within 30 days of such determination. If an employee is terminated for cause, dishonesty or fraud, his or her options lapse immediately on ceasing to be an employee. If an employee dies, all options which have not vested lapse and all options which have vested, lapse on the date 12 months after the death of the employee (to the extent that they are not exercised by the estate of the former employee).

The employee options which have not been exercised do not confer a right to notices of general meetings (except as may be required by law) or a right to attend, speak or vote at general meetings.

A holder of employee options may only participate in new issues of securities with respect to options which have been exercised and ordinary shares issued prior to the record date.

In the event of a consolidation, subdivision or similar reconstruction of our issued share capital, the number of shares to which a holder of options is entitled on exercise of an option will be adjusted in the same proportion as our issued share capital is consolidated, subdivided or reconstructed (as applicable) and an appropriate adjustment will be made to the exercise price with the effect that the total amount payable on an exercise of all options by each holder will not change.

If any pro-rata offer is made by us to at least all holders of shares, the exercise price of the relevant employee options will be reduced according to a formula set out in the Pharmaxis Employee Option Plan and consistent with the Listing Rules of the Australian Stock Exchange.

If we make a bonus issue of ordinary shares to our shareholders, the number of ordinary shares over which the employee options are exercisable may be increased by the number of shares the relevant option holder would have received if the option had been exercised prior to the record date of the bonus issue.

If we make a return of capital to our shareholders generally, the exercise price of the employee options will be proportionately reduced by the amount of the return of capital.

Except by transmission on death or with the prior written consent of our board of directors, employee options may not be transferred, encumbered, assigned or otherwise disposed of by the relevant employee. Shares issued upon exercise of options are freely transferable and we seek quotation of any such shares on the Australian Stock Exchange.

The Pharmaxis Employee Option Plan may be amended with any necessary approvals under the Corporations Act 2001 and the Listing Rules of the Australian Stock Exchange. The Corporations Act 2001 and the Listing Rules of the Australian Stock Exchange prevail over the Pharmaxis Employee Option Plan to the extent of any inconsistency. The Pharmaxis Employee Option Plan is administered by the board of directors and the Remuneration and Nomination Committee.

Presented below are summaries of options granted under the Pharmaxis Employee Option Plan during fiscal 2004 and fiscal 2005. During the period between July 1, 2005 and December 9, 2005 options to purchase 1,109,500 ordinary shares were granted, options to purchase 283,500 ordinary shares were exercised, and options to purchase 20,000 ordinary shares lapsed. As of July 1, 2005, options to purchase 10,914,000 ordinary shares were outstanding, and as of December 9, 2005, options to purchase 11,720,000 ordinary shares were outstanding.

**Year ended June 30, 2005**

<u>Grant date</u>	<u>Expiration Date</u>	<u>Exercise Price</u>	<u>Balance at Start of the Period</u>	<u>Issued During the Period</u>	<u>Exercised During the Period</u>	<u>Lapsed During the Period</u>	<u>Balance at End of the Period</u>
		A\$					
12/01/1999	11/30/2009	\$ 0.1250	2,400,000	—	—	—	2,400,000
07/01/2000	06/30/2010	0.1250	384,000	—	224,000	—	160,000
01/01/2001	12/31/2010	0.1250	96,000	—	96,000	—	—
09/01/2001	08/30/2011	0.3125	640,000	—	—	—	640,000
12/02/2001	11/30/2011	0.1250	160,000	—	—	—	160,000
05/12/2003	06/30/2012	0.3125	4,640,000	—	72,000	20,000	4,548,000
05/12/2003	11/30/2012	0.3125	480,000	—	—	—	480,000
05/12/2003	04/30/2013	0.3125	216,000	—	—	—	216,000
07/01/2003	06/30/2013	0.3125	960,000	—	—	—	960,000
07/04/2003	07/03/2013	0.3125	200,000	—	—	—	200,000
12/09/2003	11/30/2013	0.3760	500,000	—	—	—	500,000
04/25/2004	04/24/2014	0.5080	60,000	—	—	30,000	30,000
06/04/2004	06/03/2014	0.4260	15,000	—	—	—	15,000
02/02/2005	02/01/2015	0.8340	—	275,000	—	—	275,000
05/12/2005	05/11/2015	1.1470	—	330,000	—	—	330,000
			<u>10,751,000</u>	<u>605,000</u>	<u>392,000</u>	<u>50,000</u>	<u>10,914,000</u>

## Year ended June 30, 2004

Grant date	Expiration Date	Exercise Price	Balance at Start of the Period	Issued During the Period	Exercised During the Period	Lapsed During the Period	Balance at End of the Period
		(A\$)					
12/01/1999	11/30/2009	\$0.1250	2,400,000	—	—	—	2,400,000
07/01/2000	06/30/2010	0.1250	384,000	—	—	—	384,000
01/01/2001	12/31/2010	0.1250	96,000	—	—	—	96,000
09/01/2001	08/30/2011	0.3125	640,000	—	—	—	640,000
12/02/2001	11/30/2011	0.1250	160,000	—	—	—	160,000
05/12/2003	06/30/2012	0.3125	4,640,000	—	—	—	4,640,000
05/12/2003	11/30/2012	0.3125	480,000	—	—	—	480,000
05/12/2003	04/30/2013	0.3125	224,000	—	—	8,000	216,000
07/01/2003	06/30/2013	0.3125	—	960,000	—	—	960,000
07/04/2003	07/03/2013	0.3125	—	200,000	—	—	200,000
12/09/2003	11/30/2013	0.3760	—	500,000	—	—	500,000
04/25/2004	04/24/2014	0.5080	—	75,000	—	15,000	60,000
06/04/2004	06/03/2014	0.4260	—	15,000	—	—	15,000
			<u>9,024,000</u>	<u>1,750,000</u>	<u>—</u>	<u>23,000</u>	<u>10,751,000</u>

### Retirement Benefits

As required by Australian law, we contribute to standard defined contribution superannuation funds on behalf of all employees at an amount of 9% of the employee's salary. We permit employees to choose the superannuation fund into which the contributions are paid, provided the fund is appropriately registered.

We contributed A\$69,000, A\$125,000 and A\$196,000, and A\$391,000 for the fiscal years ended June 30, 2003, 2004 and 2005, and the period from inception (May 29, 1998) to June 30, 2005.

### C. Board Practices

For the date of expiration of the current term of office of our directors and senior management and the period during which the person has served in that office, see "Item 6. Directors, Senior Management and Employees – A. Directors and Senior Management."

For details of directors' service contracts with the Company providing for benefits upon termination of employment, see "Item 6. Directors, Senior Management and Employees – B. Compensation – Employment Agreements."

### Board of Directors

Our board of directors currently consists of seven directors, including five non-executive directors, of which one is non-executive chairman. Under our Constitution, the number of directors will not, unless otherwise determined by an ordinary resolution of Pharmaxis, be less than three nor more than nine. A director need not be a shareholder of Pharmaxis. Only a person over the age of 18 may be appointed as a director.

Our directors are subject to periodic retirement and re-election by shareholders in accordance with our Constitution and the Listing Rules of the Australian Stock Exchange. At each annual general meeting, one-third of our directors who are subject to retirement by rotation or, if their number is not a multiple of three, the nearest to one-third but not exceeding one-third, retire from office. Any director appointed by the directors since the last annual general meeting or for whom it would be their third annual general meeting must also retire from office. Any retiring director is eligible for reappointment. Generally, the effect of the retirement by rotation provisions is that the directors retire and are subject to re-election at staggered intervals. Messrs. McComas and Kiefel and Dr. Hillyard are subject to re-election at our annual general meeting to be held on November 15, 2005.

The directors may appoint one of themselves as a managing director, for any period and on any terms as the directors decide. The retirement by rotation provisions do not apply to the managing director. A person ceases to be a director if the Corporations Act 2001 so provides, if the director resigns by notice to Pharmaxis, if the shareholders in a general meeting remove the director, if the director is absent without the consent of the board of directors from all directors' meetings during any six-month period, if the director becomes mentally incapable and the director's estate or property has had a personal representative or trustee appointed to administer it, or if the director is an executive and he or she ceases to be an executive of Pharmaxis.

A director may appoint an alternate for a specified period with the consent of the directors. If the appointor of the alternate is not present, the alternate may attend the directors' meeting, count in the quorum, speak, and vote in the place of the appointor and exercise any other powers (except the power to appoint an alternate) that the appointor may exercise. We may pay an alternate any remuneration the directors decide, in reduction of the appointor's remuneration. We do not currently have any alternate directors.

The directors may meet, adjourn and otherwise regulate their meetings as they decide. Any director may call a directors' meeting. The quorum for a directors' meeting is two directors, unless the board of directors decides otherwise. If a person appointed as an alternate director is already a director, he or she must be counted as a director and separately as an alternate for quorum purposes. If a person is an alternate for more than one director, he or she must be counted separately for each appointment for quorum purposes. If there are not enough directors in office to form a quorum, the remaining directors may only act to increase the number of directors, to call a general meeting of shareholders or in an emergency.

Subject to the Corporations Act 2001, each director has one vote. If a director is also an alternate, the director has one vote as a director and one vote as an alternate. If a person is an alternate for more than one director, the person has one vote for each appointment. A resolution of the directors is passed by a majority of votes cast. Subject to the Listing Rules of the Australian Stock Exchange, the chairman has a deciding vote.

Our board of directors are given all our powers to manage our business except for any powers that the Corporations Act 2001 or our Constitution require Pharmaxis to exercise in a general meeting. The directors may execute documents on behalf of Pharmaxis, execute negotiable instruments, delegate any of their powers to a committee of directors or to one director and may appoint any person to be our attorney and agent.

Our directors are not prohibited from entering into proposals, arrangements and contracts in which they are interested. A director must declare to Pharmaxis the nature of their material personal interest. This notification may be a standing notification. A director is not required to give notice of an interest if the interest:

- arises because the director hold shares in Pharmaxis in common with other shareholders;
- arises in relation to the director's remuneration as a director of Pharmaxis;
- relates to a contract we are proposing to enter into that is subject to approval by our shareholders and will not impose any obligation on Pharmaxis if it is not approved by shareholders;
- arises merely because the director is a guarantor or has given an indemnity or security for all or part of a loan (or proposed loan) to Pharmaxis or arises merely because the director has a right of subrogation in relation to a guarantee or indemnity;
- relates to a contract that insures, or would insure, the directors against liabilities the director incurs as an officer of a company (but only if the contract does not make the company or related corporate entity the insurer);
- relates to any payment by Pharmaxis or a related corporate entity in respect of an indemnity permitted under the Corporations Act 2001 or a contract relating to such indemnity;
- is in a contract, or proposed contract, with, or for the benefit of, or on behalf of, a related corporate entity of Pharmaxis and arises merely because the director is a director of the related corporate entity; or



- the director has already given a standing notice of the nature and extent of the interest and the materiality of the interest has not increased above that disclosed in the notice.

A director who has a material personal interest in a proposal, arrangement or contract that is being considered at a directors' meeting must not be present while the matter is being considered at the meeting or vote on the matter and may not be counted in a quorum unless the Corporations Act 2001 provides otherwise. The director may be present and vote at such a directors' meeting if the directors who do not have a material personal interest in the matter have passed a resolution that identifies the director, the nature and extent of the director's interest in the matter and its relation to our affairs and states that those directors are satisfied that the interest should not disqualify the director from being present or voting.

At a shareholders meeting, we will disregard any votes cast by a director or any associate of a director who is voting in his or her capacity as a shareholder on a resolution relating to a proposal, arrangement or contract in which the director has a material personal interest if required to do so by the Listing Rules of the Australian Stock Exchange. The Listing Rules of the Australian Stock Exchange provide that the votes of certain shareholders must be disregarded in a number of circumstances. Generally, a shareholder's vote must be disregarded if the person may benefit from the transaction that is the subject of the resolution (unless that benefit is received in their capacity as a shareholder in common with other shareholders). For example, a director's vote in his or her capacity as a shareholder may be disregarded in respect of:

- issues of shares or options requiring the approval of shareholders, if the director is entitled to acquire securities under the issue or has acquired securities under the issue (subject to a range of exceptions including in respect of a pro-rata offer made to all shareholders);
- the issue of options to the director;
- an increase in the remuneration payable to the directors;
- termination benefits payable to directors;
- related party transactions, if the director or his or her associates may receive a benefit under a related party transaction; or
- any transaction if, in the view of the Australian Stock Exchange, the director's vote should be disregarded.

We may not be required to disregard the vote of the director if the director is voting as a proxy for a person who is entitled to vote.

We may remunerate each director as the board of directors decides, but the total amount of the remuneration of non-executive directors may not exceed the amount fixed by the shareholders in a general meeting. Other amounts may be payable by Pharmaxis to directors under our Constitution, including the payment of reasonable costs and expenses incurred in the performance of their duties or amounts paid in respect of indemnity obligations. In addition, shareholder approval may also be required in relation to the remuneration of executive directors unless the remuneration would be reasonable given our circumstances and the role of the director. Our Remuneration and Nomination Committee is responsible for recommending to the board of directors the remuneration to be paid to both non-executive directors and executive directors.

In order to loan money or give similar financial benefits to a director, we must either obtain the approval of shareholders or the financial benefit must fall under an approved exception under the Corporations Act 2001. In the case of a loan, shareholder approval may not be required if the terms of the loan are reasonable in the circumstances (as though the parties were dealing on arms length terms) or the terms are less favorable to the director than reasonable notional arms length terms or if the financial benefit is an amount which does not exceed A\$2,000. Our policies, however, ban loans to directors and executive officers.

## **Committees of the Board of Directors**

Our board of directors has established the committees described below and may establish others from time to time.

*Audit Committee.* The members of our audit committee are Messrs. Kiefel, McComas and Hanley. Mr. Kiefel chairs the audit committee. Our audit committee is responsible for:

- our financial reporting process including annual and half-year financial reports and all other financial information published by us;
- our system of internal control and management of risk;
- integrity of our financial reporting system;
- the internal and external audit process;
- our process for monitoring compliance with laws and regulations and our own Code of Conduct;
- reviewing related party transactions;
- appointing the external auditor; and
- overseeing the independence of the external auditor.

*Remuneration and Nomination Committee.* The members of our remuneration and nomination committee are Mr. Hanley, Ms. Smith, and Dr. Hillyard. Mr. Hanley chairs the committee. The purpose of our Remuneration and Nomination committee is:

- to assess the appropriate size, composition and skill mix of our board of directors;
- to evaluate the performance of our board of directors and individual directors;
- reviewing and recommending approval of compensation of directors and executive officers; and
- to administer the Pharmaxis Employee Option Plan.

## **Committees of the Board of Directors/Corporate Governance**

We have adopted a Corporate Governance Framework. In preparing the framework, we have been mindful of the Principles of Good Corporate Governance and Best Practice Recommendations issued by the Australian Stock Exchange Corporate Governance Council in March 2003. These recommendations are not mandatory, however departures from them are required to be fully disclosed in our annual report which is sent to shareholders and publicly disclosed through the Australian Stock Exchange. Conscious of the need for our policies to be appropriate for Pharmaxis, we have identified several areas where we are best served by policies that differ from the Best Practice Recommendations. We expect that our Corporate Governance Framework will alter over time as we implement our business plans, as we grow in operational complexity, as our shareholder base grows and to reflect the requirements of the Sarbanes-Oxley Act of 2002, rules adopted by the SEC and listing standards of the Nasdaq National Market.

*The Sarbanes-Oxley Act of 2002, SEC Rules and the Nasdaq National Market Marketplace Rules.* The Sarbanes-Oxley Act of 2002, as well as related new rules subsequently implemented by the SEC, require foreign private issuers, such as Pharmaxis, to comply with various corporate governance practices. In addition, Nasdaq has recently made certain changes to its corporate governance requirements for companies that are listed on the Nasdaq National Market. These changes allow us to follow Australian “home country” corporate governance practices in lieu of the otherwise applicable Nasdaq corporate governance standards, as long as we disclose each requirement of Rule 4350 that we do not follow and describe the home country practice we follow in lieu of the relevant Nasdaq corporate governance standards. We intend to take all actions necessary for Pharmaxis to maintain compliance with applicable corporate governance requirements of the Sarbanes-Oxley Act of 2002, rules adopted by the SEC and

listing standards of Nasdaq. We propose to follow Australian corporate governance practices in lieu of the corporate governance requirements of the Nasdaq Marketplace Rules in respect of:

- Nasdaq requirement under Rule 4350(f) that a quorum consist of holders of 33 1/3% of the outstanding ordinary shares – The Listing Rules of Australian Stock Exchange do not have an express requirement that each issuer listed with Australian Stock Exchange have a quorum of any particular number of the outstanding ordinary shares, but instead allow a listed issuer to establish its own quorum requirements. Our quorum is currently five persons who are entitled to vote. We believe this quorum requirement is consistent with the requirements of the Australian Stock Exchange and is appropriate and typical of generally accepted business practices in Australia.
- The Nasdaq requirements under Rules 4350(c)(1) and (2) relating to director independence, including the requirements that a majority of the board of directors must be comprised of independent directors and that independent directors must have regularly scheduled meetings at which only independent directors are present – The Nasdaq and Australian Stock Exchange definitions of what constitute an independent director are not identical and the requirements relating to the roles and obligations of independent directors are not identical. The Australian Stock Exchange, unlike Nasdaq, permits an issuer to establish its own materiality threshold for determining whether a transaction between a director and an issuer affects the director's status as independent and it does not require that a majority of the issuer's board of directors be independent, as long as the issuer publicly discloses this fact. In addition, the Australian Stock Exchange does not require that the independent directors have regularly scheduled meeting at which only independent directors are present. We believe that our board composition is consistent with the requirements of the Australian Stock Exchange and that it is appropriate and typical of generally accepted business practices in Australia.
- The Nasdaq requirements under Rule 4350(d) (other than Rule 4350(d)(2)(A)(ii), which we will comply with) relating to the composition of the audit committee and the audit committee charter – The Nasdaq and Australian Stock Exchange audit committee requirements are not identical. Moreover, differences in the requirements of Nasdaq and Australian Stock Exchange also arise because of the differences in the definitions of who constitutes an independent director, as discussed above. Issuers listed on the Australian Stock Exchange are required under applicable listing standards to establish an audit committee consisting only of non-executive directors, a majority of independent directors, an independent chair, and at least three members, and adopt a formal audit committee charter which sets out the roles and responsibilities, composition, structure and membership requirements of the audit committee, or publicly disclose that it has not done so. We have an audit committee and audit committee charter that are consistent with the requirements of the Australian Stock Exchange and which we believe are appropriate and typical of generally accepted business practices in Australia; and
- The Nasdaq requirements under Rules 4350(c)(3) and (4) that compensation of an issuer's officers must be determined, or recommended to the Board for determination, either by a majority of the independent directors, or a compensation committee comprised solely of independent directors, and that director nominees must either be selected, or recommended for the Board's selection, either by a majority of the independent directors, or a nominations committee comprised solely of independent directors – The Nasdaq compensation committee requirements are not identical to the Australian Stock Exchange remuneration and nomination committee requirements. Moreover, differences in the requirements of Nasdaq and Australian Stock Exchange with respect to these committees also arise because of the differences in the definitions of what constitutes an independent director, as discussed above. Issuers listed on the Australian Stock Exchange are required under applicable listing standards to establish a remuneration committee consisting of a majority of independent directors and an independent chairperson, or publicly disclose that it has not done so. We have a remuneration and nomination committee that is consistent with the requirements of the Australian Stock Exchange and which we believe is appropriate and typical of generally accepted business practices in Australia.

#### **Remuneration Committee Interlocks and Insider Participation**

As noted above, the remuneration and nomination committee of our board consists of Mr. Hanley, Ms. Smith and Dr. Hillyard. None of our executive officers serve as a member of the board of directors or compensation

committee of any entity that has one or more executive officers who serve on our board of directors or compensation committee.

#### D. Employees

The table below presents certain information regarding our employees and full time contractors as of June 30, 2003, 2004 and 2005, respectively.

	As of June 30,		
	2003	2004	2005
Research and development – Frenchs Forest	—	8	11
Research and development – Canberra	12	8	8
Manufacturing	3	8	15
Commercial	—	1	2
Administration	3	3	5
	<u>18</u>	<u>28</u>	<u>41</u>

Our main office facility at Frenchs Forest was established in November 2002. Apart from the research group based at the Australian National University, or ANU, all employees are based at Frenchs Forest. Seven of the researchers based at the ANU are contracted completely to Pharmaxis by ANU Enterprises Pty Ltd under a service agreement between ANU Enterprises Pty Ltd and Pharmaxis.

Each of our full time employees enter into an agreement with a term of employment of between two to three years. We also engage casual employees from time to time who enter into contracts of employment with us. We do not have any “at will” employees, as this concept is not customary in Australia. We may only terminate the employment of any of our employees in accordance with the relevant employee’s contract of employment. Our standard contract of employment for full time and part time employees provides that we can terminate the employment of an employee without notice for serious misconduct or with between one to three months notice without cause (as set out in the relevant employee’s contract of employment). Our standard contracts of employment for casual employees provide that we can terminate the employment of a casual employee without notice. For a summary of the key terms of employment of each of our executive officers, see “Item 6 – Directors, Senior Management and Employees.” Minimum notice periods may be prescribed for certain of our employees under applicable Australian law. The notice periods in our contracts of employment are equal to or exceed the minimum requirements.

On December 9, 2002, we entered into a services agreement with ANU Enterprises Pty Ltd (formerly Anutech Pty Ltd) pursuant to which ANU Enterprises Pty Ltd agrees to use its best endeavors to provide Pharmaxis with research staff on employment terms acceptable to Pharmaxis and human resource management services. We have an arrangement with the John Curtin School of Medical Research (part of the Australian National University) to provide the research staff engaged under the services agreement with access to the John Curtin School of Medical Research. Any intellectual property created by the research staff under the services agreement is owned by and vests in Pharmaxis. We have granted the Australian National University a royalty free, irrevocable and perpetual non-exclusive license to use the intellectual property and confidential information generated under the services agreement for non-commercial research purposes

None of our full time employees are represented by any collective bargaining unit. Certain of our employees may be subject under Australian law to what is known in Australia as an “industrial award” which prescribes certain minimum standards of working conditions and employment terms. The researchers contracted from the ANU Enterprises Pty Ltd are employed by the ANU Enterprises Pty Ltd on terms negotiated between the Australian National University and their employees.

We believe that we maintain good relations with all of our employees and contractors.

## E. Share Ownership

For information with respect to our ordinary shares held by the persons listed above in “Management – Directors and Executive Officers” and options granted to them on our ordinary shares, see Item 7A and Item 6—“Directors, Senior Management and Employees.”

For a description of any arrangements involving employees in the capital of Pharmaxis, see “Management – Employee Option Plan.”

## ITEM 7. MAJOR SHAREHOLDERS AND RELATED PARTY TRANSACTIONS

### A. Major Shareholders

The following table presents certain information regarding the beneficial ownership of our ordinary shares as of December 9, 2005 by the following persons:

- each person known by us to be the beneficial owner of more than 5% of our ordinary shares;
- our executive officers;
- our directors; and
- our executive officers and directors as a group.

Beneficial ownership is determined according to the rules of the Securities and Exchange Commission and generally means that a person has beneficial ownership of a security if he or she possesses sole or shared voting or investment power of that security, and includes options that are exercisable within 60 days. Information with respect to beneficial ownership has been furnished to us by each director, executive officer or 5% or more shareholder, as the case may be. Unless otherwise indicated, to our knowledge, each shareholder possesses sole voting and investment power over the shares listed, subject to community property laws where applicable. All holders of our ordinary shares have the same voting rights.

The below table lists applicable percentage ownership based on 174,453,592 ordinary shares outstanding as of December 9, 2005. Options to purchase our ordinary shares that are exercisable within 60 days of December 9, 2005 are deemed to be beneficially owned by the persons holding these options for the purpose of computing percentage ownership of that person, but are not treated as outstanding for the purpose of computing any other person’s ownership percentage.

Unless otherwise indicated in the footnotes to the table below, the address for each of the persons listed in the table below is c/o Pharmaxis Ltd, Unit 2, 10 Rodborough Road, Frenchs Forest, NSW 2086, Australia.

Individual/Group	Beneficial Ownership		Percentage of Shares Outstanding
	Shares Beneficially Owned(1)	Options Exercisable within 60 Days	
<b>5% Shareholders</b>			
Orbis Global Equity Fund Limited	19,766,000	—	11.3%
CM Capital Investments Pty Ltd(3)	14,825,000	—	8.5%
The Australian Bioscience Trust(4)	14,307,220	—	8.2%
Acorn Capital Limited(5)	9,750,932	—	5.6%
<b>Executive Officers and Directors</b>			
Alan D. Robertson, Ph.D.(6)	1,940,000	1,840,000	1.1%
William B. Cowden, Ph.D.	1,480,000	1,480,000	*
John F. Crapper	722,000	720,000	*
Ian A. McDonald, Ph.D.	—	—	*
David M. McGarvey(7)	885,000	840,000	*
Gary J. Phillips	381,664	375,000	*
Denis M. Hanley(8)	1,714,661	940,000	*
Brett Charlton, Ph.D.	1,500,000	1,480,000	*
Charles P.H. Kiefel(9)	450,000	100,000	*
Malcolm J. McComas(10)	239,999	100,000	*
Brigitte H. Smith(11)	21,789,110	—	12.5%
Carmel J. Hillyard, Ph.D.(12)	14,831,000	—	8.5%
All executive officers and directors as a group (11 persons)	45,933,434	7,875,000	26.3%

\* Represents beneficial ownership of less than one percent of our outstanding ordinary shares.



- (1) Includes ordinary shares issuable pursuant to options exercisable within 60 days of December 9, 2005. The figures represent the amounts last notified to Pharmaxis unless otherwise stated. The relevant shareholders may have acquired or disposed of share since the last notification that are not reflected. However, any such transaction that resulted in a change of one percent or greater would require the notification of such to Pharmaxis.
- (2) Consists of 3,265,000 ordinary shares and 1.1 million ADSs held by Orbis Global Equity Fund Limited and or companies associated with Orbis Global Equity Fund Limited.
- (3) Consists of (i) 11,189,044 ordinary shares held by CM Capital Investments Pty Ltd as trustee of the CM Capital Venture Trust No. 3, and (ii) 3,635,956 ordinary shares held by CIBC Australia VC Fund LLC, as general partner of the Australia Venture Capital Fund LP, of which CM Capital Investments Pty Ltd is a special limited partner.
- (4) All of these shares are held of record by Perpetual Trustees Nominees Limited, in its capacity as trustee of the Australian Bioscience Trust, and Perpetual Trustees Nominees Limited disclaims beneficial ownership of all such shares. GBS Venture Partners Ltd, or GBS, in its capacity as investment manager of the Australian Bioscience Trust, may be deemed to have beneficial ownership of all such shares. Ms. Smith, a member of our board of directors, is a Managing Director of GBS and has shared voting and dispositive power over such shares. Ms. Smith disclaims beneficial ownership of these shares, except to the extent of her pecuniary interest therein.
- (5) All of these shares are held of record by nominee and trustee companies on behalf of Acorn Capital Limited, in its capacity as discretionary investment manager to certain superannuation funds, pooled superannuation trusts, managed investment schemes and investment management agreements. Acorn Capital Limited has sole voting and dispositive power over these shares.
- (6) Includes 100,000 ordinary shares held by Dr. Robertson's spouse.
- (7) Includes 5,000 ordinary shares held by McGarvey Investments Pty Ltd., of which Mr. McGarvey's spouse is the sole director and shareholder. Mr. McGarvey disclaims beneficial ownership over the shares held by McGarvey Investments Pty Ltd.
- (8) Includes 151,333 ordinary shares held by a superannuation fund of which Mr. Hanley is the beneficiary. Also includes 16,664 ordinary shares held by Mr. Hanley's spouse.
- (9) Includes 150,000 ordinary shares held by Mr. Kiefel's spouse.
- (10) Includes (i) 100,000 ordinary shares held by Movilli Pty Ltd, and (ii) 26,666 ordinary shares held by the Bunyala Super Pty Ltd. Mr. McComas has shared voting and dispositive power over the shares held by these two entities. Also includes 13,333 ordinary shares held by Mr. McComas' spouse.

- (11) Consists of (i) 14,307,220 ordinary shares beneficially owned by The Australian Bioscience Trust over which GBS holds sole voting and dispositive power in its capacity as investment manager, and (ii) 7,481,890 ordinary shares beneficially owned by Bioscience Ventures II over which GBS holds sole voting and dispositive power in its capacity as trustee. Ms. Smith, a managing director of GBS, may be deemed to have an indirect pecuniary interest in an indeterminate portion of all such shares. Ms. Smith disclaims beneficial ownership of all such shares, except to the extent of her pecuniary interest therein.
- (12) Includes 6,000 ordinary shares held by Bionetworks Superannuation Fund over which Ms. Hillyard holds shared voting and dispositive power. Also includes (i) 11,189,044 ordinary shares held by CM Capital Investments Pty Ltd as trustee of the CM Capital Venture Trust No. 3, and (ii) 3,635,956 ordinary shares held by CIBC Australia VC Fund LLC, as general partner of the Australia Venture Capital Fund LP, of which CM Capital Investments Pty Ltd is a special limited partner. Ms. Hillyard is a founder and partner of CM Capital Investments Pty Ltd and may be deemed to have an indirect pecuniary interest in an indeterminate portion of all shares held by CM Capital Venture Trust No. 3 and CIBC Australia VC Fund LLC. Ms. Hillyard disclaims beneficial ownership of such shares, except to the extent of her pecuniary interest therein.

### Significant Changes to Percentage Ownership of Principal Shareholders

The following table presents information with respect to certain significant changes in percentage ownership of our ordinary shares held by beneficial holders of more than five percent of our ordinary shares since June 30, 2004.

<u>5% Shareholder</u>	<u>Date</u>	<u>% of Shares Beneficially Owned</u>	<u>Change in % of Shares Beneficially Owned</u>
The Australian Bioscience Trust	08/28/2002(1)	33.5%	(25.3)%
	11/10/2003(2)	18.7	(14.4)
	03/11/2005(3)	11.9	(3.1)
	11/30/2005(7)	8.2	(3.7)
CM Capital Investment Pty Ltd	08/28/2002(1)	16.8	16.8
	11/10/2003(2)	13.7	(2.8)
	11/11/2005(8)	8.5	(5.2)
Acorn Capital Limited	11/10/2003(2)	3.0	3.0
	02/27/2004(4)	5.2	2.2
	06/24/2004(5)	6.2	1.0
	09/13/2004(6)	7.2	1.0
	11/11/2005(9)	5.6	(1.6)
Orbis Global Equity Fund Limited	11/11/2005(10)	11.3	11.3

- (1) After giving effect to shares purchased by the listed entity as part of the first closing of the private placement of our "B" class convertible redeemable preference shares in August 2002.



- (2) After giving effect to shares purchased by the listed entity as part of the initial public offering of our ordinary shares on the Australian Stock Exchange in November 2003.
- (3) After giving effect to the sale of 4,204,800 ordinary shares by the listed entity on March 11, 2005.
- (4) After giving effect to (i) 3,200,000 ordinary shares purchased by Acorn Capital Limited as part of the initial public offering of our ordinary shares on the Australian Stock Exchange in November 2003 and (ii) 2,377,359 ordinary shares purchased in private transactions from September 24, 2003 to February 27, 2004.
- (5) After giving effect to the purchase of 1,109,441 ordinary shares in private transactions from March 1, 2004 to June 24, 2004.
- (6) After giving effect to the purchase of 1,130,559 shares in private transactions from June 28, 2004 to September 13, 2004.
- (7) After giving effect to the sale of 1,732,980 ordinary shares by the listed entity on November 30, 2005
- (8) After giving effect to the issue of shares by the company in November 2005 in its US public offering and Australian private placement
- (9) After giving effect to the issue of shares by the company in November 2005 in its US public offering and Australian private placement, and after giving effect to the purchase of 1,933,573 ordinary shares on October 1, 2004
- (10) After giving effect to shares purchased by the listed entity as part of the US public offering of our ADS and Australian private placement of our ordinary shares in November 2005.

We are not aware of any other significant changes in percentage ownership with respect to our principal shareholders resulting from their respective purchases or sales of our ordinary shares.

Our major shareholders do not have different voting rights.

As of November 30, 2005, two percent of our ordinary shares were held in the United States by eight holders of record, and 93% of our ordinary shares were held in Australia by 3,590 holders of record.

As of December 9, 2005, 1417,532 ADSs, representing 21,262,980 ordinary shares, were outstanding in the United States, all of which were held of record by Cede & Co.

## B. Related Party Transactions

### Share Sales

The following executive officers and directors purchased shares from us in the amounts and on the dates shown below since June 30, 2004:

<u>Executive Officers and Directors</u>	<u>Date</u>	<u>Number of Shares</u>	<u>Aggregate Purchase Price (A\$)</u>
John F. Crapper	12/16/2004(1)	20,000	15,000
Gary J. Phillips	12/14/2004(2)	6,664	4,998
Denis M. Hanley	12/16/2004(1)	157,997	118,498
Malcolm McComas	12/16/2004(1)	26,666	20,000

- (1) Shares purchased as part of the second closing of a private placement of our ordinary shares.

(2) Shares purchased as part of a share purchase plan conducted in December 2004.

#### **Stock Options Granted to Executive Officers and Directors**

See Item 6 - "Directors, Senior Management and Employees."

#### **Relationships and Transactions with The Principal Funds Management Pty Ltd**

Messrs. Hanley and Kiefel are shareholders of Principal Funds. Cameron Billingsley of PFM Legal Pty Ltd, our primary legal counsel in Australia, serves as a director of Principal Funds.

#### **Employment Agreements**

We have entered into employment agreements with our executive officers. For more information regarding these agreements, see Item 6 - "Directors, Senior Management and Employees."

#### **Director Indemnification**

We have on various dates entered into Deeds of Access to Documents and Indemnity agreements with certain executive officers and each of our directors. See Item 6 - "Directors, Senior Management and Employees."

#### **C. Interests of Experts and Counsel**

Not Applicable.

### **ITEM 8. FINANCIAL INFORMATION**

#### **A. Consolidated Statements and Other Financial Information**

Our financial statements are included as Item 18 of this registration statement on Form 20-F.

#### **Legal Proceedings**

We are not involved in any significant legal, arbitration or governmental proceedings. We are not aware of any pending significant legal, arbitration or governmental proceedings with respect to the Company.

#### **Dividend Policy**

We have never declared or paid any cash dividends on our ordinary stock and we do not anticipate paying any cash dividends in the foreseeable future. Dividends may only be paid out of our profits. Payment of cash dividends, if any, in the future will be at the discretion of our board of directors or, if our directors do not exercise their power to issue dividends, our shareholders in a general meeting may exercise the powers. See "Item 10. Additional Information – B. Constitution– Dividends."

#### **B. Significant Changes**

On November 11, 2005 we announced the completion of a Global Capital Raising with gross proceeds of A\$87 million (US\$63 million). A total of 39.4 million shares (including 1.3 million American Depositary Shares) were issued bringing total shares on issue after the raising to 174.4 million.

In the US, 1.3 million American Depositary Shares (ADSs) were issued to US institutional investors in a public offering at a price of US\$24.16 per ADS. CIBC World Markets Corp. acted as sole book-running manager of the offering. JPM Securities LLC served as co-manager of the offering.

In Australia, 19.9 million ordinary shares were issued to Australian and other non US institutional, sophisticated and professional investors at a price of A\$2.20 per ordinary share, equivalent to the ADS price. Wilson HTM served as sole agent for the Australian placement.

## ITEM 9. THE OFFER AND LISTING

### A. Offer and Listing Details – Price History

#### Ordinary Shares

The following tables present, for the periods indicated, the high and low market prices for our ordinary shares reported on the Australian Stock Exchange since November 10, 2003, the date on which our ordinary shares were initially quoted. Prior to the initial quotation of our ordinary shares on the Australian Stock Exchange on November 10, 2003, our ordinary shares were not regularly traded in any organized market and were not liquid.

		<u>High</u>	<u>Low</u>
		A\$	A\$
Fiscal Year 2004	First Quarter	\$ —	\$ —
	Second Quarter	0.570	0.340
	Third Quarter	0.550	0.360
	Fourth Quarter	0.510	0.415
	From November 10, 2003 to June 30, 2004	0.570	0.340
Fiscal Year 2005	First Quarter	0.860	0.485
	Second Quarter	0.940	0.730
	Third Quarter	1.310	0.730
	Fourth Quarter	1.850	1.020
	Full Year	1.850	0.485
Fiscal 2006 (through December 9, 2005)		3.280	1.530
Most Recent Six Months			
	June 2005	1.850	1.155
	July 2005	1.890	1.530
	August 2005	2.410	1.715
	September 2005	2.970	2.070
	October 2005	3.280	2.330
	November 2005	2.570	2.030

#### American Depositary Shares (ADS)

The following tables present, for the periods indicated, the high and low market prices for our ADSs as reported on the Nasdaq National Market since August 29, 2005, the date on which our ADSs were initially quoted. Prior to the initial quotation of our ADSs on the Nasdaq National Market on August 29, 2005, our ADSs were not regularly traded in any organized market and were not liquid.

	<u>High</u>	<u>Low</u>
	U.S.\$	U.S.\$
August 2005	\$30.230	\$26.400
September 2005	32.000	23.750
October 2005	35.980	28.360
November 2005	28.000	24.160
Fiscal 2006 (through December 9, 2005)	35.980	23.050

**B. Plan of Distribution**

Not applicable.

**C. Markets**

Our ordinary shares are traded on the Australian Stock Exchange Ltd. and are traded on the Nasdaq National Market.

**D. Selling Shareholders**

Not applicable.

**E. Dilution**

Not applicable.

**F. Expenses of the Issue**

Not applicable.

**ITEM 10. ADDITIONAL INFORMATION**

**A. Description of Share Capital**

Not applicable

**B. Constitution**

Our constituent document is a Constitution which is similar in nature to the by-laws of a company incorporated under the laws of the U.S. Our Constitution does not provide for or prescribe any specific objects or purposes of the Company. Our Constitution is subject to the terms of the Listing Rules of the Australian Stock Exchange and the Corporations Act 2001. Our Constitution may be amended or repealed and replaced by special resolution of shareholders, which is a resolution passed by at least 75% of the votes cast by shareholders entitled to vote on the resolution.

**Shareholders Meetings**

We must hold an annual general meeting within five months of the end of each fiscal year. Our end of fiscal year is currently June 30 each year. At the annual general meeting, shareholders typically consider the annual financial report, directors' report and auditors report and vote on matters, including the election of directors, the appointment of the auditor and fixing the non-executive directors' and auditor's remuneration. We may also hold other meetings of shareholders from time to time. The annual general meeting must be held in addition to any other meetings which we may hold.

A director or the board of directors may call and arrange a meeting of shareholders, when and where they decide. The directors must call a meeting of shareholders when requested by shareholders who hold at least 5% of the votes that may be cast at the meeting or at least 100 members who are entitled to vote at the meeting or as

otherwise required by the Corporations Act 2001. Shareholders with at least 5% of the votes in Pharmaxis may also call a general meeting at their own cost.

At least 28 calendar days notice must be given of a meeting of shareholders. A meeting of shareholders may be called on shorter notice if, in respect of the annual general meeting, all of the shareholders agree beforehand, or in respect of any other meeting of shareholders, if 95% of the shareholders agree beforehand.

Directors, auditors, shareholders, proxies, and attorneys and representatives of shareholders are entitled to attend general meetings. We may refuse admission to the meeting to anyone (other than a director) in accordance with our Constitution and applicable Australian law. For the purpose of determining who is a shareholder at a particular meeting, the directors will determine that shareholders at a specified time (typically this will be 48 hours before the meeting) are taken to be shareholders at the meeting.

The necessary quorum for a meeting of shareholders is five shareholders entitled to vote. We believe this quorum requirement is consistent with common practice for many listed Australian publicly listed companies.

Unless applicable law or our Constitution requires a special resolution, a resolution of shareholders is passed if more than 50% of the votes cast by shareholders entitled to vote are cast in favor of the resolution. A special resolution is passed if the notice of meeting sets out the intention to propose the special resolution and states the resolution and it is passed by at least 75% of the votes cast by shareholders entitled to vote on the resolution. A special resolution usually involves more important questions affecting Pharmaxis as a whole or the rights of some or all of our shareholders. Special resolutions are required in a variety of circumstances under our Constitution and the Corporations Act 2001, including without limitation:

- to change our name;
- to amend or repeal and replace our Constitution;
- to approve the terms of issue of preference shares;
- to approve the variation of class rights of any class of shareholders;
- to convert one class of shares into another class of shares;
- to approve certain buy backs of shares;
- to approve a selective capital reduction of our shares;
- to approve Pharmaxis financially assisting a person to acquire shares in Pharmaxis;
- to remove and replace our auditor;
- to approve the transfer of our place of registration to registration under a law of another state or territory of Australia;
- to change our company type;
- with the leave of an authorized Australian court, to approve our voluntary winding up;
- to confer on a liquidator of Pharmaxis either a general authority or a particular authority in respect of compensation arrangements of the liquidator; and
- to approve an arrangement entered into between a company about to be, or in the course of being, wound up.

### **Shareholder Voting Rights**

At a general meeting, every shareholder present (in person or by proxy, attorney or representative) and entitled to vote has one vote on a show of hands. Every shareholder present (in person or by proxy, attorney or representative) and entitled to vote has one vote per fully paid ordinary share on a poll. This is subject to any other rights or restrictions which may be attached to any shares. In the case of an equality of votes on a resolution at a

meeting (whether on a show of hands or on a poll), the chairman of the meeting has a deciding vote in addition to any vote that the chairman of the meeting has in respect of that resolution. A poll may be requested on any resolution by at least five shareholders entitled to vote on the resolution, by shareholders holding at least 5% of the votes that may be cast on the resolution or by the chairman.

The Listing Rules of the Australian Stock Exchange provide that the votes of certain shareholders must be disregarded in certain circumstances. Generally, a shareholder's vote may be disregarded if the person may benefit from the transaction that is the subject of the resolution (subject to certain exceptions, such as where the benefit is received in their capacity as a shareholder in common with other shareholders). Without limitation, a shareholder's vote may be disregarded in respect of:

- the issue of shares or options, if the shareholder is entitled to acquire securities under the issue or has acquired securities under the issue (subject to a range of exceptions including in respect of a pro-rata offer made to all shareholders) or is entitled to any other sort of benefit as a result of the issue (for example underwriting commissions);
- the amendment of the terms of options, if the shareholder holds the relevant options;
- if the shareholder is a director, to approve an increase in the remuneration payable to the directors;
- if the shareholder is a director, in respect of termination benefits payable to directors;
- the acquisition or disposal of a substantial asset;
- the issue of securities to specified related parties or anyone else the Australian Stock Exchange considers should not be entitled to vote; and
- significant transactions such as changes to the nature and scale of our operations or a change to our main undertaking.

The Australian Stock Exchange may also identify a person who in their view should not be entitled to vote.

### **Issue of Shares and Changes in Capital**

Subject to our Constitution, the Corporations Act 2001, the Listing Rules of the Australian Stock Exchange and any other applicable law, we may at any time issue shares and grant options or warrants on any terms, with preferred, deferred or other special rights and restrictions and for the consideration and other terms that the directors determine. Our power to issue shares includes the power to issue bonus shares (for which no consideration is payable to Pharmaxis), preference shares (including redeemable preference shares) and partly paid shares.

Subject to the requirements of our Constitution, the Corporations Act 2001, the Listing Rules of the Australian Stock Exchange and any other applicable law we may:

- consolidate or divide our share capital into a larger or smaller number by resolution passed by shareholders at a general meeting;
- may reduce our share capital by special resolution passed by at least 75% of the votes cast by shareholders entitled to vote on the resolution provided that the reduction is fair and reasonable to our shareholders as a whole, and does not materially prejudice our ability to pay creditors;
- undertake an equal access buyback of our ordinary shares by ordinary resolution of shareholders (although if we have bought back less than 10% of our shares over the period of the previous 12 months, shareholder approval may not be required); and
- undertake a selective buyback of certain shareholders' shares by special resolution passed by at least 75% of the votes cast by shareholders entitled to vote on the resolution, with no votes being cast in favor of the resolution by any person whose shares are proposed to be bought back or by their associates.

In certain circumstances, including the division of a class of shares into further classes of shares, the issue of additional shares or the issue of a new class of shares, we may require the approval of any class of shareholders whose rights are varied or are taken to be varied by special resolution of shareholders generally and by special resolution of the holder of shares in that class whose rights are varied or taken to be varied.

### **Dividends**

Subject to any special rights or restrictions attached to a share, we may pay dividends on our shares as the directors decide. Dividends may be only paid out of our profits.

Subject to any special rights or restrictions attached to a share, the directors may determine that a dividend will be payable on a share and fix the amount, the time for payment and the method for payment. Dividends may be paid on shares of one class but not another and at different rates for different classes. If the board of directors does not exercise their power to issue dividends, the shareholders in a general meeting may. Under our Constitution, a shareholder or shareholders holding the requisite number of shares required to convene a general meeting would be able to convene a meeting or require the directors to convene a meeting to consider whether we should pay a dividend. The proposed resolution to pay the dividend would need to be included in the notice of meeting and would be voted on by shareholders as an ordinary resolution. Any dividend payable would only be payable out of our profits.

### **Liquidation Rights**

Subject to any special rights or restrictions attached to shares, on a winding up, all available assets must be repaid to the shareholders and any surplus must be distributed among the shareholders in proportion to the number of fully paid shares held by them. For this purpose a partly paid share is treated as a fraction of a share equal to the proportion which the amount paid bears to the total issue price of the share before the winding up began.

If we experience financial problems, the directors may appoint an administrator to take over our operations to see if we can come to an arrangement with our creditors. If we cannot agree with our creditors, Pharmaxis may be wound up.

A receiver, or receiver and manager, may be appointed by order of a court or under an agreement with a secured creditor to take over some or all of the assets of a company. A receiver may be appointed, for example, because an amount owed to a secured creditor is overdue.

We may be wound up by order of a court, or voluntarily if our shareholders pass a special resolution to do so. A liquidator is appointed when a court orders a company to be wound up or the shareholders of a company pass a resolution to wind up the company. A liquidator is appointed to administer the winding up of a company.

### **Calls, Lien and Forfeiture in Respect of Partly Paid Shares**

Subject to any special rights or restrictions attached to shares, the board of directors may make calls on the holder of a share for any unpaid portion of the issue price of that share at any time. The directors may make a call payable by installments. If the amount called is not paid by the requisite time, the shareholder must pay Pharmaxis interest on the amount unpaid from the date the call becomes payable until and including the date of payment and our costs arising from the non-payment. Joint holders of a share and their respective personal representatives are all jointly and severally liable to pay all calls on the share. The board of directors may recover an amount presently payable as a result of a call by suing the shareholder for the debt, by enforcing the lien on the share or by declaring forfeit the share. The forfeiture of a share extinguishes the former shareholder's interest in the share. We have a first ranking lien on each share registered to a shareholder, dividends payable on a shares, proceeds on the sale of a share for an unpaid call or installment that is due but unpaid on the share, any amounts we are required by law to pay in respect of the shares of that shareholder, and in respect of any interest and costs presently payable to Pharmaxis by the shareholder. We may sell a share to enforce a lien in certain circumstances.

We do not currently have any partly paid shares outstanding.

## Limitations on Rights to Own Shares and ADSs

The Foreign Acquisitions and Takeovers Act 1975 regulates acquisitions of shares by non-Australian persons giving rise to substantial interests or controlling interests in an Australian companies. Some of the relevant terms of the Foreign Acquisitions and Takeovers Act 1975 are summarized below.

In general terms, the Foreign Acquisitions and Takeovers Act 1975 prohibits a foreign interest from acquiring shares or entering into an agreement to acquire shares or interests in shares if, after the acquisition or agreement, such foreign interest would hold a substantial interest or controlling interest in an Australian corporation, without first applying for approval by the Treasurer of the Australian Government and such approval being granted or 40 days having elapsed after such application was made.

For foreign investors other than U.S. investors, the notification obligation arises in relation to proposals to acquire a substantial interest or controlling interest in an Australian business, the value of whose assets exceeds A\$50 million or whose business is valued at over A\$50 million. Our business currently has a market valuation of greater than A\$50 million. However, in the case of U.S. investors, other than the U.S. government and other than when the investment proposal relates to investments in prescribed sensitive sectors, the requirement to notify the Australian Government of a proposal to acquire a substantial interest or a controlling interest in an Australian business arises when the value of the assets of the relevant Australian business exceeds A\$800 million or the value of the Australian business exceeds A\$800 million. A U.S. investor is defined as a national or permanent resident of the U.S., a U.S. enterprise, or a branch of an entity located in the U.S. and carrying on business activities in the U.S. We currently have assets and a market value less than A\$800 million and would not be regarded as a business in a sensitive sector.

A “foreign interest” is defined, in summary, as:

- a natural person not ordinarily resident in Australia;
- a company in which a natural person not ordinarily resident in Australia or a foreign company holds a substantial interest;
- a company in which two or more persons, each of whom is either a natural person not ordinarily resident in Australia or a foreign company, hold an aggregate substantial interest;
- the trustee of a trust estate in which a natural person not ordinarily resident in Australia or a foreign company holds a substantial interest; or
- the trustee of a trust estate in which two or more persons, each of whom is either a natural person not ordinarily resident in Australia or a foreign company, hold an aggregate substantial interest.

In summary, a person is taken to hold a substantial interest in a company if:

- the person, alone or together with any associate or associates of the person, is in a position to control not less than 15% of the voting power in the company or holds legal or equitable interests in not less than 15% of the issued shares in the company; or
- two or more persons are taken to hold an aggregate substantial interest in a company if they, together with any associate or associates of any of them, are in a position to control not less than 40% of the voting power in the company or hold legal or equitable interests in not less than 40% of the issued shares in the company.

Where a person holds a substantial interest in a company or two or more persons hold an aggregate substantial interest in a company, that person will be taken to hold a controlling interest in the company, or those persons will be taken to hold an aggregate controlling interest in the company, unless the Treasurer is satisfied that the person together with their associates (if any) are not in a position to determine the policy of the company.

The Treasurer may make an order prohibiting a proposed acquisition of shares or all or any of the proposed acquisitions. Where the Treasurer makes an order prohibiting a proposed acquisition of shares, it may also make an



order in relation to a specified foreign person and their associates prohibiting those persons from acquiring additional interests or voting rights in the company.

Where a person has acquired shares in a company, and the Treasurer is satisfied that the acquisition has had the result that the company becomes controlled by foreign persons, or in the case of a company that was previously controlled by foreign persons, includes a person who is not one of the foreign persons forming part of the existing foreign interest, and that result is contrary to Australia's national interest, the Treasurer may make an order directing the person who acquired the shares to dispose of those shares within a specified time to any person or persons approved in writing by the Australian government.

If a person or persons acquires shares or enters into an agreement to acquire shares or interests which requires the approval of the Treasurer, but the person or persons fails to get approval, the person or persons are guilty of an offence and may be liable to penalties and imprisonment. Among other things, orders are able to be made restraining the exercise of any rights attached to shares held by the foreign person or corporation and directing the disposal of shares.

Shareholders, potential shareholders and holders of ADSs and potential holders of ADSs are urged to get their own independent legal advice in relation to the application of the Foreign Acquisitions and Takeovers Act 1975.

## **Change of Control**

### *Corporations Act 2001*

Takeovers of listed Australian public companies, such as Pharmaxis, are regulated amongst other things by the Corporations Act 2001 which prohibits the acquisition of a relevant interest in issued voting shares in a listed company if the acquisition will lead to the person's or someone else's voting power in the company increasing from 20% or below to more than 20% or increasing from a starting point that is above 20% and below 90%, subject to a range of exceptions.

A relevant interest is defined very broadly to capture most forms of interest in shares and would include interests in our ADSs. Generally, and without limitation, a person will have a relevant interest in securities if they:

- are the holder of the securities;
- have power to exercise, or control the exercise of, a right to vote attached to the securities; or
- have power to dispose of, or control the exercise of a power to dispose of, the securities (including any indirect or direct power or control).

It does not matter how remote the relevant interest is or how it arises. If two or more people can jointly exercise one of these powers, each of them is taken to have that power.

If at a particular time a person has a relevant interest in issued securities and the person:

- has entered or enters into an agreement with another person with respect to the securities;
- has given or gives another person an enforceable right, or has been or is given an enforceable right by another person, in relation to the securities; or
- has granted or grants an option to, or has been or is granted an option by, another person with respect to the securities,

and the other person would have a relevant interest in the securities if the agreement were performed, the right enforced or the option exercised, the other person is taken to already have a relevant interest in the securities.

A person will also be regarded as having a relevant interest in voting shares in a company if the non-voting securities in which the person already had a relevant interest become voting shares in the company or there is an increase in the number of votes that may be cast on a poll attached to voting shares that the person already had a

relevant interest in. In these circumstances, the acquisition of the relevant interest will occur when the securities become voting shares or the number of votes increases.

There are a number of exceptions to the prohibition on acquiring a relevant interest in issued voting shares in a listed company if the acquisition will lead to the person's or someone else's voting power in the company increasing from 20% or below to more than 20% or increasing from a starting point that is above 20% and below 90%. In general terms, some of the more significant exceptions include:

- when the acquisition results from the acceptance of an offer under a formal takeover bid;
- when the acquisition is conducted on market by or on behalf of the bidder under a takeover bid and the acquisition occurs during the bid period;
- when shareholders of the company approve the takeover by resolution passed at general meeting;
- an acquisition by a person if, throughout the 6 months before the acquisition, that person, or any other person, has had voting power in the company of at least 19% and as a result of the acquisition, none of the relevant persons would have voting power in the company more than 3 percentage points higher than they had 6 months before the acquisition;
- as a result of a rights issue;
- as a result of dividend reinvestment schemes;
- as a result of underwriting arrangements;
- through operation of law;
- an acquisition which arises through the acquisition of a relevant interest in another listed company;
- arising from an auction of forfeited shares; or
- arising through a compromise, arrangement, liquidation or buyback.

Breaches of the takeovers provisions of the Corporations Act 2001 are criminal offences. The Australian Securities and Investments Commission and the Australian Takeover Panel have a wide range of powers relating to breaches of takeover provisions including the ability to make orders canceling contracts, freezing transfers of, and rights attached to, securities, and forcing a party to dispose of securities. There are certain defenses to breaches to the takeovers provisions provided in the Corporations Act 2001.

#### *Proportional Takeover*

Our Constitution contains what is known as a proportional takeover provision which provides that the registration of transfers giving effect to a takeover for only a specified proportion of Pharmaxis is prohibited until a resolution to approve the bid is passed by shareholders of the bid class of securities. The resolution is passed if the proportion of bid class shareholders accepting the resolution is greater than 50%. The proportional takeover provision in our Constitution expires in September 2006. Shareholders may prior to or after that time renew the applicability of the proportional takeover provision at a general meeting.

#### **Disclosure of Interests**

The Corporations Act 2001 requires that a person must give notice to Pharmaxis in the prescribed form within two business days (or in some cases by the next business day) if:

- the person begins to have, or ceases to have, a substantial holding in Pharmaxis. A substantial holding will arise if a person and their associates have a relevant interest in 5% or more of the votes in Pharmaxis or the person has made a takeover bid for the voting shares in Pharmaxis;
- if the person has a substantial holding in Pharmaxis and there is a movement of 1% in their holding; or
- if the person makes a takeover bid for Pharmaxis.

For the purposes of the notification obligation, a relevant interest in the voting shares is defined very broadly to capture most forms of interests in shares and would include interests in our ADSs. Generally, a person will have a relevant interest in securities if such person is the holder of the securities, has power to exercise, or control the exercise of, a right to vote attached to the securities or has power to dispose of, or control the exercise of a power to dispose of, the securities (including any indirect or direct control or power). Likewise, associates are defined broadly and includes:

- corporate entities owned or controlled by the person;
- corporate entities that control the person;
- corporate entities that are controlled by an entity which controls the person;
- persons with whom the person has or proposes to enter into agreements with which relate to the composition of our board; and
- persons with whom the person is acting or is proposing to act in concert.

The rights attaching to our shares for non-compliance with the disclosure of interest requirements may result in disenfranchisement, loss of entitlement to dividends and other payments and restrictions on transfer. A person who contravenes these obligations is liable to compensate a person for any loss or damage the person suffers because of the contravention.

### **C. Material Contracts**

Following is a summary of our material contracts, other than contracts entered into in the ordinary course of business, to which we are a party, for the two years immediately preceding the filing of this document.

#### *License Agreement with the Central Sydney Area Health Service*

On October 10, 2001, we entered into a license agreement with Central Sydney Area Health Service. Pursuant to the license agreement, Central Sydney Area Health Service grants us an exclusive, worldwide license, which is able to be sublicensed, to exploit certain key intellectual property and patents relating to the use of respirable dry powders for the assessment of bronchial hyper-responsiveness, a condition consistent with active asthma, for monitoring steroid use in asthma patients, and for the management of diseases such as cystic fibrosis, bronchiectasis and chronic bronchitis.

There is no fixed expiry date for the license agreement. The term of the license in each relevant country is for the longer of 10 years from the first commercial sale of products which exploits the Central Sydney Area Health Service intellectual property in that country or until the expiry of the last registered patent in that country. The license may be terminated earlier by either party if there is a breach of the agreement by a party and that party fails to remedy the breach within 30 days after receiving notice to do so, or if any party becomes insolvent or if we determine in our commercial judgment that it is not prudent to continue the license. If we decide not to obtain product approval in any country, we will not unreasonably refuse to convert the license into a non-exclusive license for that country.

We must bear the cost of maintaining the relevant registered Central Sydney Area Health Service intellectual property and must use our reasonable commercial endeavors to exploit and undertake research and development of the intellectual property.

We may at our own cost prosecute applications for any new patentable inventions arising in the course of exploiting the Central Sydney Area Health Service intellectual property, in our name. If we do not seek patent protection for the new patentable invention in any country, Central Sydney Area Health Service may at its own cost file patent applications.

For the term of the license, we are liable to pay the royalties described below to Central Sydney Area Health Service on the net sales of products and services which exploit the Central Sydney Area Health Service intellectual property.

In respect of the upper and lower airway function test application of the intellectual property:

- no royalties until aggregate net sales of products and services from all countries of A\$500,000 have been achieved;
- a royalty of 4% of the gross margin if the net sales of the products or services by us achieve a gross margin of 20% or less;
- a royalty of 8% of the gross margin if the net sales of the products or services by us achieve a gross margin between 20% and 40%;
- a royalty of 10% of the gross margin if the net sales of the products or services by us achieve a gross margin greater than 40%; and
- 20% of any royalty received from any sub-licensee.

In respect of the mucociliary clearance and sputum induction applications of the intellectual property:

- no royalties are payable until sales representing a gross margin of A\$1 million have been achieved then, when the gross margin achieved by the product sales is between A\$1 million and A\$25 million a royalty equal to 3% of the gross margin will apply, when it is between A\$25 million and A\$75 million a royalty equal to 2.5% of the gross margin will apply and when it is greater than A\$75 million a royalty equal to 2% of the gross margin will apply; and
- 20% of any royalty received from a sub-licensee.

To date, we have not paid Central Sydney Area Health Service any royalty or license fees. We are not able to accurately estimate the aggregate amount of potential payments that may be due to Central Sydney Area Health Service as this amount will be a function of the future sales of our applicable products and the percentage royalty and license fees set out above.

We have agreed to indemnify Central Sydney Area Health Service against all loss and damage that Central Sydney Area Health Service may sustain or incur as a result of any actions, claims, suits, proceedings or demands arising directly or indirectly out of the breach of the license by us. Both parties have agreed to indemnify the other party against all loss and damage that the other party may sustain or incur as a result of any damage to the other party's property or injury to or death of any of the other party's personnel arising out of the agreement.

Subject to a policy being available on commercially reasonable terms, we must maintain a product liability insurance policy naming Central Sydney Area Health Service, both during the term of the agreement and for a period of six years after the termination of the agreement.

#### *License Agreement with ANU Enterprises Pty Ltd*

On October 14, 1999, we entered into a license agreement with ANU Enterprises Pty Ltd (formerly called Anutech Pty Ltd). Pursuant to the license agreement, ANU Enterprises Pty Ltd, as agent for the Australian National University, grants us an exclusive, worldwide license, with the right to sub-license, to exploit intellectual property and patents relating to the use of phosphosugars and their analogues as anti-inflammatory agents in the field of ethical therapeutics. PXS25/64 is not covered by patents that are the subject of this agreement. We have granted to ANU Enterprises Pty Ltd a royalty-free non-exclusive license to use the intellectual property and patents licensed to us and any of our improvements to that intellectual property for their internal research purposes.

There is no fixed expiry date for the license agreement. The term of the license agreement in respect of each patent is for the life of each of the licensed patents. The license may be terminated earlier for breach of the agreement if the defaulting party fails to remedy a breach under the agreement within 90 days of receipt of written notice from the other party (or such longer period as may be agreed), or if the default is not capable of being remedied and the defaulting party does not agree to pay compensation for the loss being suffered as a result of the breach. The non-defaulting party may terminate the agreement if one party defaults in the payment of any money due and the defaulting party fails to remedy the breach within 30 days of receipt of notice from the non-defaulting party. The license agreement may also be terminated by one party if the other party becomes insolvent.

We must use our reasonable endeavors to exploit the intellectual property and licensed patents which are the subject of the agreement and have agreed to commence the sale of products which incorporate or arise from the whole or partial use of the intellectual property and licensed patents by 2011.

Any new intellectual property acquired or developed by us during the term of and in connection with the agreement is our property. ANU Enterprises Pty Ltd must advise us of the filing of any patent application or the issue of any patent which is legally or beneficially owned by the Australian National University or ANU Enterprises Pty Ltd which dominates or is dominated by the licensed patents or relates to the product arising wholly or partially from the intellectual property or licensed patents. For a period of three months after notification of any such intellectual property, we have the option of having any such intellectual property licensed to us under the agreement without any change to the royalty rate payable.

We must reimburse ANU Enterprises Pty Ltd for the costs incurred in filing, maintaining and renewing the Australian National University's intellectual property. A royalty of 2% of revenue (net of expenses) received by us in connection with its use of the intellectual property and licensed patents is payable by us to ANU Enterprises Pty Ltd. The obligation to pay the royalty survives termination of the agreement. To date, we have reimbursed ANU Enterprises Pty Ltd for certain patents costs and expenses but have not paid any royalties. We are not able to accurately estimate the aggregate amount of potential payments that may be due to ANU Enterprises Pty Ltd as this amount will be a function of the future sales of our applicable products (if any) and the percentage of royalty payments set out above.

We have agreed to indemnify the Australian National University and ANU Enterprises Pty Ltd and any of their directors, officers, employees, staff, students and agents against all loss, liability, damage, claim, cost and expense arising from or in connection with, amongst other things, breach by us of the agreement or our use of the intellectual property and licensed patents.

We must maintain a product liability insurance policy in respect of products related to the agreement.

#### *AusIndustry P3 Pharmaceuticals Partnerships Program Funding Deed*

On August 12, 2004, we entered into a funding deed with the Commonwealth of Australia under the AusIndustry P3 Pharmaceuticals Partnerships Program. The term of the funding deed is until June 30, 2008. Subject to us expending certain eligible expenditure and undertaking certain eligible activities in connection with projects for the development of new treatments for autoimmune diseases (multiple sclerosis and rheumatoid arthritis) and the development of new treatments for chronic respiratory disease, the Commonwealth of Australia may, as of June 30, 2005, pay us a total amount of A\$4.5 million between July 2005 and June 2008 for eligible pharmaceutical research and development activities undertaken by us in relation to the development of new treatments for autoimmune diseases and the development of new treatments for chronic respiratory diseases. As at June 30, 2005, we have received A\$55,481 under the funding deed.

Payments are made under the funding deed depending on the amount of eligible expenditure forecasted and actually made by us. Payment amounts are limited to specified amounts each year which may be increased or decreased in accordance with the terms of the funding deed. The amount to be paid under the funding agreement is reduced by other financial assistance received from the Commonwealth of Australia or any Australian state governments.

We are not entitled to funds under the funding agreement if the Commonwealth of Australia has insufficient funding for the program, if we fail to submit reports when required, if we have not otherwise complied with our obligations under the funding agreement or if the Commonwealth of Australia is entitled to terminate the deed. The Commonwealth of Australia may withhold payment if it considers that we are not entitled to the payment under the deed, for such time as the Commonwealth of Australia requires to make inquiries to determine whether we are entitled to payment. We may be required to repay the amount of any overpayment of the grant if the Commonwealth of Australia gives us notice that they have paid us more than we are entitled to be paid.

The Commonwealth of Australia does not assert any ownership of, or any right to, any of the intellectual property created under the funding deed. We have granted to the Commonwealth of Australia a permanent, irrevocable, royalty free, worldwide, non-exclusive license to use, reproduce, publish, transmit, adapt and modify

any documents and associated materials brought into existence for the purpose of us reporting on the performance of our obligations under the deed or otherwise used in connection with the grant program. This licensed material may only be used for the purposes of the Commonwealth of Australia's dissemination, reporting and accountability requirements, but not to commercially exploit such material.

We must provide a range of reports, including quarterly, annual and ad-hoc reports and must respond to requests for information, in relation to the performance of our obligations under the funding deed to the Commonwealth of Australia. We must also provide the Commonwealth with audited financial statements verifying actual expenditure. The Commonwealth of Australia is entitled to undertake assessments and reviews of our performance of our obligations at stipulated times under the funding deed and also on an ad-hoc basis.

The funding deed terminates on the earliest of the following:

- June 30, 2008;
- if we enter into a new funding deed with the Commonwealth of Australia in relation to the research and development;
- if we agree with the Commonwealth of Australia to terminate the funding deed; and
- if the Commonwealth of Australia has paid the maximum amounts to be paid under the deed.

The Commonwealth of Australia may terminate the funding deed:

- if the Commonwealth of Australia gives us not less than 30 days written notice of termination;
- if we are in breach of the funding deed and in the opinion of the Commonwealth of Australia the breach is not capable of being remedied or if capable of being remedied it is not remedied within 21 days of receipt of written notice;
- in certain circumstances, if we fail to submit reports;
- if our research and development activities or the quality of those activities do not satisfy the grant eligibility criteria;
- if there is a change of control of us; or
- if we become insolvent.

#### *Research and Development Start Program Grant Agreement*

On June 17, 2003, we entered into a grant agreement with the Commonwealth of Australia under the research and development Start Grant Program. The Commonwealth of Australia has provided us with a grant of 50% of our eligible expenditure on a project for the development of a new treatment for cystic fibrosis up to a maximum grant amount of A\$3.0 million payable over the period to December 31, 2005. At June 30, 2005, we had received A\$2.5 million of the grant out of an aggregate maximum of A\$3.0 million. The grant agreement formally terminates on December 31, 2010. We have ongoing reporting obligations beyond the project completion date until the termination of the grant agreement.

We may only use the grant solely for the project for the development of a new treatment for cystic fibrosis. We may not encumber our rights under the grant agreement. Grant payments are made in accordance with an agreed schedule subject to the Commonwealth of Australia having sufficient funding available and us making the necessary eligible expenditure, achieving satisfactory progress on a project for the development of a new treatment for cystic fibrosis and having submitted all progress reports due. We must use our best endeavors to commercialize the project for the development of a new treatment for cystic fibrosis on normal commercial terms within a reasonable time of completion of the project.

We must provide reports to the Commonwealth of Australia every three months.

The Commonwealth of Australia may terminate the grant agreement for breach of the agreement by us, for a failure by us to undertake the required research, if there is a change of control of us, or on the grounds of our insolvency. In certain limited circumstances where we fail to use our best endeavors to commercialize the project within a reasonable time of completion of the project or upon termination of a grant due to our breach of agreement or our insolvency, the Commonwealth of Australia may require us to repay some or all of the grant. We consider that the likelihood of being required to repay grant funding is remote while we continue to act in good faith with respect to this grant. We may be required to repay the amount of any overpayment of the grant if the Commonwealth of Australia gives us notice that they have paid us more than we are entitled to be paid. To date, we have not been required to repay any amounts paid to us under our current two grant agreements and we are not aware of any current circumstances that would require us to repay any such amounts.

#### **D. Exchange Controls**

For a description of any governmental laws, decrees, regulations or other legislation of Australia which may affect (1) the import or export of capital, including the availability of cash and cash equivalents for use by us, or (2) the remittance of dividends, interest or other payments to nonresident holders of our securities, see “Item 10. Additional Information – B. Constitution.”

#### **E. Taxation**

The following is a summary of certain material Australian income tax and U.S. federal income tax considerations related to the ownership and disposition of our ordinary shares or ADSs that may be relevant to you if you are a U.S. Holder (as defined below). This summary is based on the Australian and U.S. tax laws currently in effect. The term “U.S. Holder” means a beneficial owner of our ordinary shares or ADSs that is, for U.S. federal income tax purposes, a citizen or individual resident of the United States, a domestic corporation, an estate whose income is subject to U.S. federal income tax regardless of its source, or a trust if a U.S. court can exercise primary supervision over the administration of the trust and one or more U.S. persons are authorized to control all substantial decisions of the trust, or a trust that has a valid election in effect under applicable U.S. Treasury regulations to be treated as a U.S. person.

Prospective purchasers of our ordinary shares or ADSs should consult their own tax advisors regarding the application of the Australian and U.S. federal income tax laws to their particular situations as well as any tax considerations under other tax laws (such as estate and gift tax laws), or the laws of any province, state or local jurisdiction.

If an entity is treated as a partnership, the tax treatment of a partner will generally depend on the status of the partner and upon the activity of the partnership. If you are a partner of a partnership that will hold our ordinary shares or ADSs, we suggest you consult your own tax advisor.

#### **Australian Taxation**

The following summary of the Australian taxation implications is based on the provisions of the Income Tax Assessment Act 1936, the Income Tax Assessment Act 1997, the International Tax Agreements Act 1953, or IntTAA with the United States Convention as amended by the United States Protocol, or USDTA, public taxation rulings and available case law current as at the date of this annual report, or collectively referred to in this section as Australian Taxation Laws. The Australian Taxation Laws and their interpretation are subject to change at any time.

#### **General Principle of Taxation in Australia**

This summary discusses only two items of income that may arise from an investment in our ordinary shares or ADSs, namely:

- capital gains realized from the sale of our ordinary shares or ADSs; and
- dividends that may be paid by us with respect to those shares and ADSs.

Please note that we have not paid any dividends to date and do not expect to pay any in the near to medium term.

## **Capital Gains on Sale of Shares or ADSs**

Under Australian law, income tax is typically not payable on the gain made on the disposal of ordinary shares or ADSs by U.S. Holders unless the profit is of income in nature and sourced in Australia or the sale is subject to tax on any net capital gains, in each case as broadly summarized below.

### **When the Profit on Sale is Income in Nature**

For U.S. Holders who are carrying on a business in Australia through a permanent establishment or that are providing personal services in Australia through a fixed base (where any profit on the sale of our ordinary shares or ADSs is effectively connected to that business or fixed place, e.g., the U.S. Holder is carrying on a business of share trading in Australia), then any profit made on the sale of our ordinary shares or ADSs may be regarded as ordinary income and may therefore be required to be included in the assessable income of the relevant U.S. Holders and taxed accordingly.

If the previous paragraph does not apply, a U.S. Holder that realizes a capital gain from the sale of our ordinary shares or ADSs may nevertheless still be subject to Australian tax on the capital gain in the circumstances set out below.

### **When the Sale is Subject to Capital Gains Tax**

A U.S. Holder will be required to include in its assessable income in Australia any “net capital gains” that it realizes which have a “necessary connection with Australia.” Net capital gains for a U.S. Holder will only arise where there are capital gains or capital losses realized with respect to assets which have a “necessary connection with Australia.” A U.S. Holder of our ordinary shares or ADSs will only have a “necessary connection with Australia” if the U.S. Holder and its associates beneficially own or have owned in the 5 years prior to a sale of our ordinary shares or ADSs, at least 10% of the value of our combined number of issued shares (except any shares that carry a right only to participating in a distribution of profits or capital to a limited extent) and ADSs, or if the US Holder uses our ordinary shares or ADSs in carrying on business through a permanent establishment in Australia.

Therefore, unless a U.S. Holder satisfies these tests there will be no tax payable on any gain on the sale of those shares or ADSs.

A U.S. Holder who satisfies these tests will be required to calculate its net capital gains for the relevant income year taking into account the capital gain (and any relevant discount thereon) or capital loss realized on the sale of our ordinary shares or ADSs. The net capital gain is then included in the U.S. Holder’s assessable income in Australia and will be taxed accordingly.

A summary of a method for calculating net capital gains is to: (1) deduct from the capital gains, all capital losses; (2) deduct from the capital gains all past unapplied net capital losses; and (3) reduce the remaining capital gains by any applicable capital gains discount. Natural persons and some trusts are entitled to a 50% capital gains discount in circumstances where our ordinary shares or ADSs have been sold after being held for more than 12 months. The 50% capital gains discount is not available to companies.

## **Dividends**

Dividends paid to U.S. Holders will be subject to the withholding tax provisions of the Australian Taxation Laws, unless the US Holders hold their shares in us through a permanent establishment in Australia.

The general withholding tax rate in Australia for dividends is 30% but under the USDTA this is reduced to 5% of the gross amount of the dividend if the person beneficially entitled to the dividend is a company which holds at least 10% of the voting power in the company or otherwise is reduced to 15%. If the U.S. Holder has held shares which hold a voting power of at least 80% for at least a 12 month period then there may be no withholding tax if the holder is a certain type of person such as a listed company.

However certain dividends paid to non-residents are exempt from withholding tax.

Australia has an imputation system which allows a company which distributes profits to its members to pass on to its members a credit for the tax already paid by the company to its members. This is known as a franking



credit. To the extent that the dividend is franked, the dividend is not subject to withholding tax. This means that a fully franked dividend is not subject to any withholding tax. To the extent that the dividend is not franked, then that part of the dividend will be subject to withholding tax but at the reduced rate referred to above.

A dividend which is unfranked is also exempt from withholding tax to the extent it is referable to certain categories of foreign income of the payer which are treated on a conduit basis in Australia.

There are also additional exemptions depending on the nature of the shareholder which are designed to ensure that an entity that is otherwise exempt from tax is not subject to withholding tax, e.g., charitable institutions.

If a US Holder holds shares in us through a permanent establishment in Australia, then dividends paid on those shares will not be subject to withholding tax but will be assessed as taxable income in Australia and taxed at the marginal tax rates. Such a US Holder may be entitled to a tax offset for any franking credit attached to such dividends.

## **U.S. Taxation**

The following is a summary of certain material U.S. federal income tax considerations related to the ownership and disposition of our ordinary shares and ADSs that may be relevant to you if you are a U.S. Holder. This summary is based on the Internal Revenue Code of 1986, as amended (the “Code”), existing and proposed Treasury regulations promulgated under the Code and administrative and judicial interpretations of the Code, all as of the date of this annual report and all of which are subject to change, possibly with retroactive effect.

This summary is also based in part upon the representations of the depository and the assumption that each obligation in the deposit agreement and any related agreement will be performed in accordance with its terms. In general, and taking into account such assumptions, a U.S. Holder of ADSs will be treated as an owner of the ordinary shares represented by those ADSs. Therefore, exchanges of ordinary shares for ADSs, and ADSs for ordinary shares, will not have federal income tax consequences.

This summary deals only with ordinary shares and ADSs held as capital assets within the meaning of Section 1221 of the Code. It does not discuss all of the tax considerations that may be relevant to U.S. Holders in light of their particular circumstances or to U.S. Holders subject to special rules, such as dealers in securities or currencies, traders in securities that elect to mark-to-market their securities, expatriates, partnerships and other pass through entities, tax-exempt organizations, insurance companies, U.S. Holders subject to the alternative minimum tax, U.S. Holders that actually or constructively own 10% or more of our ordinary shares, U.S. Holders holding our ordinary shares or ADSs as part of a hedging or constructive sale transaction, “straddle,” conversion transaction, or other integrated transaction, or U.S. Holders whose functional currency is not the U.S. dollar.

## **Ownership of Ordinary Shares and ADSs**

The gross amount of any distribution received by a U.S. Holder with respect to our ordinary shares or ADSs generally will be included in the U.S. Holder’s gross income as a dividend to the extent attributable to our current and accumulated earnings and profits (as determined under U.S. federal income tax principles). To the extent a distribution received by a U.S. Holder is not a dividend because it exceeds the U.S. Holder’s pro rata share of our current and accumulated earnings and profits, it will be treated first as a tax-free return of capital and reduce (but no below zero) the adjusted tax basis of the U.S. Holder’s shares. To the extent the distribution exceeds the adjusted tax basis of the U.S. Holder’s shares, the remainder will be taxed as capital gain (the taxation of capital gain is discussed under the heading “Sale of Ordinary Shares and ADSs” below).

For taxable years beginning after December 31, 2002 and before January 1, 2009, dividends received by non-corporate U.S. holders from a qualified foreign corporation are taxed at the same preferential rates that apply to net long-term capital gains. A foreign corporation is a “qualified foreign corporation” if it is eligible for the benefits of a comprehensive income tax treaty with the United States (the income tax treaty between Australia and the United States is such a treaty) or its shares or ADSs with respect to which such dividend is paid are readily tradable on an established securities market in the United States (such as the Nasdaq National Market on which our ADSs will be traded). Notwithstanding satisfaction of one or both of these conditions, a foreign corporation is not a qualified foreign corporation if it is a passive foreign investment company (“PFIC”) for the taxable year of the corporation in which the dividend is paid or the preceding taxable year. A foreign corporation will also not be a qualified foreign

corporation with respect to a particular holder (subject to certain limited exceptions) if it was a PFIC for any taxable year in that holder's holding period. Dividends received from a foreign corporation that is not a qualified foreign corporation will be taxed at ordinary income tax rates. As discussed in more detail below, under the section entitled "Taxation—Passive Foreign Investment Companies," there is a risk that we could be a PFIC in the future.

If a distribution is paid in Australian dollars, the U.S. dollar value of such distribution on the date of receipt is used to determine the amount of the distribution received by a U.S. Holder (and the amount of Australian tax withheld, if any). A U.S. Holder who continues to hold such Australian dollars after the date on which they are received, may recognize gain or loss upon their disposition due to exchange rate fluctuations. Generally such gains and losses will be ordinary income or loss from U.S. sources.

U.S. Holders may deduct Australian tax withheld from distributions they receive from us for the purpose of computing their U.S. federal taxable income or alternatively elect to claim a foreign tax credit against their U.S. federal income tax liability for such taxes. The foreign tax credit is subject to a number of limitations and the rules governing its determination are very complex. Prospective U.S. Holders should consult their own tax advisors to determine whether and to what extent they would be entitled to claim a foreign tax credit.

Corporate U.S. Holders generally will not be allowed a dividends received deduction with respect to dividends they receive from us.

### **Sale of Ordinary Shares and ADSs**

Subject to the passive foreign investment company rules discussed below, a U.S. Holder that sells or otherwise disposes of ordinary shares or ADSs will recognize capital gain or loss equal to the difference between the U.S. dollar value of the amount realized and its adjusted tax basis in those ordinary shares or ADSs. This gain or loss generally will be capital gain or loss from U.S. sources, and will be long-term capital gain or loss if the U.S. Holder held its shares for more than 12 months. Generally, the net long-term capital gain of a non-corporate U.S. Holder recognized before January 1, 2009 is subject to tax at a top marginal rate of 15%. Capital gain that is not long-term capital gain is taxed at ordinary income tax rates.

### **Passive Foreign Investment Companies**

We will be a PFIC if in any taxable year either: (a) 75% or more of our gross income consists of passive income; or (b) 50% or more of the value of our assets is attributable to assets that produce, or are held for the production of, passive income. Subject to certain limited exceptions, if we meet the gross income test or the asset test for a particular taxable year, ordinary shares or ADSs held by a U.S. Holder in that year will be treated as shares of a PFIC ("Pharmaxis PFIC Shares") for that year and all subsequent years in the U.S. Holder's holding period, even if we fail to meet either test in a subsequent year.

Gain realized from the sale of Pharmaxis PFIC Shares will be subject to tax under the excess distribution regime, unless the U.S. Holder makes one of the elections discussed below. Under the excess distribution regime, federal income tax on a U.S. Holder's gain from the sale of Pharmaxis PFIC Shares would be calculated by allocating the gain ratably to each day the U.S. Holder held its ordinary shares or ADSs. Gain allocated to years preceding the first year in which we were a PFIC in the U.S. Holder's holding period, if any, and gain allocated to the year of disposition would be treated as gain arising in the year of disposition and taxed as ordinary income. Gain allocated to all other years (the "Pharmaxis PFIC Years") would be taxed at the highest tax rate in effect for each of those years. Interest for the late payment of tax would be calculated and added to the tax due for each of the Pharmaxis PFIC Years, as if the tax was due and payable with the tax return filed for that year. A distribution that exceeds 125% of the average distributions received on Pharmaxis PFIC Shares by a U.S. Holder during the 3 preceding taxable years (or, if shorter, the portion of the U.S. Holder's holding period before the taxable year) would be taxed in a similar manner.

A U.S. Holder may avoid taxation under the excess distribution regime by making a qualified electing fund ("QEF") election. For each year that we would meet the PFIC gross income test or asset test, an electing U.S. Holder would be required to include in gross income, its pro rata share of our net ordinary income and net capital gains, if any. The U.S. Holder's adjusted tax basis in our shares would be increased by the amount of such income inclusions. An actual distribution to the U.S. Holder out of such income inclusions would not be treated as a dividend and would decrease the U.S. Holder's adjusted tax basis in our shares. Gain realized from the sale of our ordinary shares

or ADSs covered by a QEF election would be taxed as a capital gain. A U.S. Holder may make a QEF election, only if we agree in advance to provide the U.S. Holder the information necessary to allow the U.S. Holder to comply with the QEF rules. Due to the administrative burden associated with our providing this information to each U.S. Holder, we will not agree to provide this information to U.S. Holders. Accordingly, a U.S. Holder will not be eligible to make a QEF election.

A U.S. Holder may also avoid taxation under the excess distribution regime by timely making a mark-to-market election. An electing U.S. Holder would include in gross income the increase in the value of its Pharmaxis PFIC Shares during each of its taxable years and deduct from gross income the decrease in the value of its Pharmaxis PFIC Shares during each of its taxable years. Amounts included in gross income or deducted from gross income by an electing U.S. Holder are treated as ordinary income and ordinary deductions from U.S. sources. Deductions for any year are limited to the amount by which the income inclusions of prior years exceed the income deductions of prior years. Gain from the sale of Pharmaxis PFIC Shares covered by an election is treated as ordinary income from U.S. sources while a loss is treated as an ordinary deduction from U.S. sources only to the extent of prior income inclusions. Losses in excess of such prior income inclusions are treated as capital losses from U.S. sources. A mark-to-market election is timely if it is made by the due date of the U.S. Holder's tax return for the first taxable year in which the U.S. Holder held our ordinary shares or ADSs that includes the close of our taxable year for which we met the PFIC gross income test or asset test. A mark-to-market election is made on IRS Form 8621.

As noted above (under the heading titled "Ownership of Ordinary Shares and ADSs"), a PFIC is not a qualified foreign corporation and hence dividends received from a PFIC are not eligible for taxation at preferential net long-term capital gain tax rates. Similarly, ordinary income included in the gross income of a U.S. Holder as a result of the holder having made a QEF election or a mark-to-market election, and dividends received from corporations subject to such election, are not eligible for taxation at preferential net long-term capital gain rates.

Based on an analysis of our gross income and the value of our assets, we believe that we were a PFIC for our taxable year ended June 30, 2004, but do not believe that we were a PFIC for our taxable year ended June 30, 2005. Although we do not anticipate being a PFIC in our current taxable year or in the future, we may meet the PFIC income test and be a PFIC if we earn more interest income than anticipated or earn less operating income than anticipated.

#### **U.S. Information Reporting and Backup Withholding**

United States information reporting and backup withholding requirements may apply with respect to distributions to U.S. Holders, or the payment of proceeds from the sale of shares, unless the U.S. Holder: (a) is an exempt recipient (including a corporation); (b) complies with certain requirements, including applicable certification requirements; or (c) is described in certain other categories of persons. The backup withholding tax rate is currently 28%. Any amounts withheld from a payment to a U.S. Holder under the backup withholding rules may be credited against any U.S. federal income tax liability of the U.S. Holder and may entitle the U.S. Holder to a refund.

#### **F. Dividends and Paying Agents**

Not applicable.

#### **G. Experts**

Not applicable.

#### **H. Documents on Display**

We will file annual reports and other information with the Securities and Exchange Commission after the effectiveness of this registration statement on Form 20-F. We will file annual reports on Form 20-F and submit other information under cover of Form 6-K. As a foreign private issuer, we are exempt from the proxy requirements of Section 14 of the Exchange Act and our officers, directors and principal shareholders will be exempt from the insider short-swing disclosure and profit recovery rules of Section 16 of the Exchange Act. Annual reports and other information we file with the Commission may be inspected at the public reference facilities maintained by the Commission at Room 1024, Judiciary Plaza, 450 Fifth Street, N.W., Washington, D.C. 20549, and at its regional

offices located at 233 Broadway, New York, New York 10279 and 500 West Madison Street, Suite 1400, Chicago, Illinois 60661, and copies of all or any part thereof may be obtained from such offices upon payment of the prescribed fees. You may call the Commission at 1-800-SEC-0330 for further information on the operation of the public reference rooms and you can request copies of the documents upon payment of a duplicating fee, by writing to the Commission. In addition, the Commission maintains a web site that contains reports and other information regarding registrants (including us) that file electronically with the Commission which can be assessed at <http://www.sec.gov>.

#### **I. Subsidiary Information**

Not applicable.

#### **ENFORCEABILITY OF CIVIL LIABILITIES**

We are a public company incorporated and domiciled under the laws of Australia. A majority of our directors and executive officers are residents of countries other than the United States. Furthermore, all or a substantial portion of their assets and our assets are located outside the United States. As a result, it may not be possible for you to:

- effect service of process within the United States upon any of our directors and executive officers or on us; or
- enforce in U.S. courts judgments obtained against any of our directors and executive officers or us in the U.S. courts in any action, including actions under the civil liability provisions of U.S. securities laws;
- enforce in U.S. courts judgments obtained against any of our directors and senior management or us in courts of jurisdictions outside the United States in any action, including actions under the civil liability provisions of U.S. securities laws; or
- to bring an original action in an Australian court to enforce liabilities against any of our directors and executive officers or us based upon U.S. securities laws.

You may also have difficulties enforcing in courts outside the United States judgments obtained in the U.S. courts against any of our directors and executive officers or us, including actions under the civil liability provisions of the U.S. securities laws.

#### **ITEM 11. QUANTITATIVE AND QUALITATIVE DISCLOSURES ABOUT MARKET RISK**

See “Item 5. Operating and Financial Review and Prospects – B. Liquidity and Capital Resources – Quantitative and Qualitative Disclosures about Market Risk.”

#### **ITEM 12. DESCRIPTION OF SECURITIES OTHER THAN EQUITY SECURITIES**

Not applicable.

## PART II

### ITEM 13. DEFAULTS, DIVIDEND ARREARAGES AND DELINQUENCIES

Not applicable.

### ITEM 14. MATERIAL MODIFICATIONS TO THE RIGHTS OF SECURITY HOLDERS AND USE OF PROCEEDS

- A. Not applicable.
- B. Not applicable
- C. Not applicable
- D. Not applicable
- E. Use of Proceeds
  - 1. Our registration statement on Form F-1 was declared effective by the Securities and Exchange Commission on November 7, 2005 and is assigned No. 333-128581.
  - 2. Pursuant to a preliminary prospectus dated October 17, 2005, the Company initially offered 1,400,000 ADSs which amount was subsequently reduced to 1,300,000 in the final prospectus dated November 7, 2005. The Offering closed on November 10, 2005.
  - 3. Not applicable.
  - 4. (a) All shares offered pursuant to the final prospectus were sold and the Offering terminated.
    - (b) CIBC World Markets was lead underwriter.
    - (c) Ordinary Shares in the form of American Depositary Shares were registered and sold.
    - (d) Amount we registered for sale by us: 21,000,000 Ordinary Shares representing 1,400,000 ADSs; we sold 19,500,000 Ordinary Shares (1,300,000 ADSs) for an aggregate offering price of US\$31,408,000.  
Amount we registered for sale by selling shareholders upon exercise of overallotment option: 3,150,000 Ordinary Shares representing 210,000 ADSs; selling shareholders sold 2,635,890 Ordinary Shares (175,726 ADSs) pursuant to a partial exercise of the overallotment option.
    - (e) The underwriting discount payable to the underwriters and other expenses of the offering are estimated to be US\$ 3,062,266, all of which was paid to non-affiliates.
    - (f) The net proceeds of the offering after deducting underwriting discount and expenses were US\$28.3 million.
    - (g) All of the net proceeds of the offering are invested in short-term interest-bearing investment-grade securities of non-affiliates.
    - (h) Not applicable.

Concurrently with the offering described above, we placed 19,900,000 Ordinary Shares in an offshore transaction (“Australian Placement”) at the same price per Ordinary Share at which Ordinary Shares (in the form of ADSs) were sold pursuant to the registration statement described above. The net proceeds of the Australian Placement after underwriting fees and expenses, were US\$30.2 million.

### ITEM 15. CONTROLS AND PROCEDURES

#### Evaluation of Disclosure Controls and Procedures

An evaluation was performed under the supervision and with the participation of our management team, including our Chief Executive Officer and Chief Financial Officer, regarding the effectiveness of the design and operation of our disclosure controls and procedures as of the end of the June 30, 2005. Based on that evaluation, our management, including our Chief Executive Officer and Chief Financial Officer, have concluded that our disclosure controls and procedures as of June 30, 2005 provided reasonable assurance of effectiveness that information required to be disclosed by us in reports that we file or submit under the Exchange Act is recorded, processed, summarized and reported within the time periods specified in Securities and Exchange Commission rules and forms. In designing and evaluating the disclosure controls and procedures, management recognized that any controls and

procedures, no matter how well designed and operated, can provide only reasonable, rather than absolute, assurance of achieving the desired control objectives, and management necessarily was required to apply its judgement in evaluating the cost-benefit relationship of possible controls and procedures.

### Changes in Internal Controls Over Financial Reporting

There were no changes in our internal control over financial reporting (as defined in Rule 13a-15(f) of the Exchange Act) that occurred during the year ended June 30, 2005 that have materially affected, or are reasonably likely to materially affect, our internal controls over financial reporting.

### ITEM 16A. AUDIT COMMITTEE FINANCIAL EXPERT

Our board of directors adopted its audit committee charter on December 4, 2003. Our board has determined that we do not have a financial expert serving on our audit committee as defined by Item 401(h) of Regulation S-K of the Exchange Act.

We believe that the combined knowledge, skills and experience of the members of our Audit Committee enables them, as a group, to act effectively in the fulfillment of their tasks and responsibilities, including those under the Sarbanes-Oxley Act of 2002, without appointing a member who would qualify as an audit committee financial expert.

### ITEM 16B. CODE OF ETHICS

We have adopted a Code of Conduct that applies to the Chief Executive Officer and all senior financial officers, or persons performing similar functions, of the Company. A copy of the Company's Code of Conduct is attached as an exhibit to this annual report on Form 20-F. The Code of Conduct is also posted on the Company's website at [www.pharmaxis.com.au](http://www.pharmaxis.com.au).

### ITEM 16C. PRINCIPAL ACCOUNTANT FEES AND SERVICES

The following table represents aggregate fees billed to the Company for fiscal years ended June 30, 2005 and June 30, 2004 by PricewaterhouseCoopers, the Company's principal accounting firm.

	Years ended June 30,	
	2004	2005
	(in thousands)	
Audit fees	\$ 99	\$ 507
Audit-related fees	—	—
Tax	—	—
Other	7	—
	<u>\$ 106</u>	<u>\$ 507</u>

Audit fees include fees for audit or review services in accordance with generally accepted auditing standards and fees for services that generally only the auditor can reasonably provide.

The Audit Committee of our Board of Directors chooses and engages our independent auditors to audit our financial statements. In 2005, our Board of Directors also adopted a policy requiring management to obtain the audit committee's approval before engaging our independent auditors to provide any other audit or permitted non-audit services to us. This policy, which is designed to assure that such engagements do not impair the independence of our auditors, requires the audit committee to pre-approve audit and non-audit services that may be performed by our auditors.

**ITEM 16D. EXEMPTIONS FROM THE LISTING STANDARDS FOR AUDIT COMMITTEES**

Not applicable

**ITEM 16E. PURCHASES OF EQUITY SECURITIES BY THE ISSUER AND AFFILIATED PURCHASERS**

Not applicable.

## PART III

### ITEM 17. FINANCIAL STATEMENTS

Not applicable.

### ITEM 18. FINANCIAL STATEMENTS

The financial statements are included as the "F" pages attached to this report

### ITEM 19. EXHIBITS

The following exhibits are filed as part of this report

<u>EXHIBIT NO.</u>	<u>DESCRIPTION</u>
1.1	Constitution*
2.1	Form of American Depositary Receipt (included in Exhibit 2.2) *
2.2	Form of Depositary Agreement (Incorporated by reference to the Registration Statement on Form F-6 (File No. 333-120026) filed by the Company with the Commission on October 28, 2004)
4.1	Form of Deed of Access to Documents and Indemnity entered into with Company Directors and Company Secretary*
4.2	Form of Employment Agreement entered into with Company Officers*
4.3	Lease dated June 22, 2001 between Trust Company of Australia Ltd and Aortech Biomaterials Pty Ltd, and Transfer of Lease dated October 15, 2002 among Aortech Biomaterials Pty Ltd, the Company (formerly Pharmaxis Pty Ltd) and Trust Company of Australia Ltd*
4.4	Agreement dated October 14, 1999 between ANU Enterprises Pty Ltd (formerly Anutech Pty Ltd) and the Company (formerly Praxis Pharmaceuticals Australia Pty Ltd), as amended*
4.5	Patent License Agreement dated October 10, 2001 between the Company and Central Sydney Area Health Service (formerly Praxis Pharmaceuticals Australia Pty Ltd) *
4.6	Service Agreement dated December 9, 2002 between ANU Enterprises Pty Ltd (formerly Anutech Pty Ltd) and the Company, as amended*
4.7	R&D Start Grant Program Grant Agreement dated June 17, 2003 between the Commonwealth of Australia (acting through the Industry Research and Development Board) and the Company*
4.8	Pharmaceuticals Partnerships Program P <sup>3</sup> Funding Agreement dated August 12, 2004 between the Commonwealth of Australia (represented by the Department of Industry, Tourism and Resources) and the Company*
11.	Code of Conduct
12.	Certification required by Rule 13a-14(a)
13.	Certification pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002

\* Incorporated by reference to our Registration Statement on Form 20-F (File No. 000-51505)



**SIGNATURES**

The registrant hereby certifies that it meets all of the requirements for filing on Form 20-F and that it has duly caused and authorized the undersigned to sign this registration statement on its behalf.

**PHARMAXIS LTD**

By:           /s/  DAVID M. MCGARVEY            
Name: **David M. McGarvey**  
Title: **Chief Financial Officer**

Date: December 20, 2005

## **Index to Financial Statements**

<u>Report of PricewaterhouseCoopers, Independent Registered Public Accounting Firm</u>	F-2
<u>Balance Sheets as of June 30, 2004 and 2005</u>	F-3
<u>Statements of Operations for the years ended June 30, 2003, 2004 and 2005 and for the period from inception (May 29, 1998) to June 30, 2005</u>	F-4
<u>Statements of Changes in Shareholders' (Deficit) Equity for the period from inception (May 29, 1998) to June 30, 2005</u>	F-5
<u>Statements of Cash Flows for the years ended June 30, 2003, 2004 and 2005 and for the period from inception (May 29, 1998) to June 30, 2005</u>	F-6
<u>Notes to Financial Statements</u>	F-7
<u>Schedule II – Valuation and Qualifying Accounts</u>	F-20

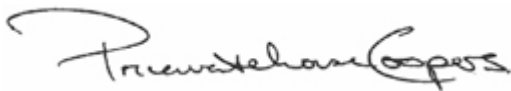
**PricewaterhouseCoopers**  
**ABN 52 780 433 757**

Darling Park Tower 2  
201 Sussex Street  
GPO BOX 2650  
SYDNEY NSW 1171  
DX 77 Sydney  
Australia  
www.pwc.com/au  
Telephone +61 2 8266 0000  
Facsimile +61 2 8266 9999

**Report of Independent Registered Public Accounting Firm**

To the board of directors and shareholders of Pharmaxis Ltd:

In our opinion, the financial statements listed in the accompanying index present fairly, in all material respects, the financial position of Pharmaxis Ltd (the "Company") (a development stage enterprise) at June 30, 2004 and 2005, and the results of its operations and its cash flows for each of the three years in the period ended June 30, 2005 and, cumulatively, for the period from May 29, 1998 (date of inception) to June 30, 2005 in conformity with accounting principles generally accepted in the United States of America. In addition, in our opinion, the financial statement schedule listed in the accompanying index presents fairly, in all material respects, the information set forth therein when read in conjunction with the related financial statements. These financial statements and the financial statement schedule are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements and the financial statement schedule based on our audits. We conducted our audits of these statements in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. An audit includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements, assessing the accounting principles used and significant estimates made by management, and evaluating the overall financial statement presentation. We believe that our audits provide a reasonable basis for our opinion.



PricewaterhouseCoopers



WHB Seaton

Sydney, Australia

19 September 2005, except for Note 19 for which the date is 10 November 2005  
Liability limited by a scheme approved under Professional Standards Legislation

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Balance Sheets**  
**(Australian dollars)**

	As of June 30,	
	2004	2005
	(in thousands, except share data)	
<b>ASSETS</b>		
Current assets:		
Cash and cash equivalents	\$ 25,101	\$ 33,268
Grant receivable	—	106
Other current assets	264	718
	25,365	34,092
Property, plant and equipment, net	1,324	2,376
Intangible assets, net	1,162	1,106
Other long-term assets	260	262
	\$ 28,111	\$ 37,836
<b>LIABILITIES AND SHAREHOLDERS' EQUITY</b>		
Current liabilities:		
Accounts payable	\$ 245	\$ 757
Accrued compensation	294	484
Accrued clinical development liabilities	806	546
Other accrued liabilities	135	582
	1,480	2,369
Commitments and Contingencies		
Shareholders' equity:		
Ordinary shares, \$Nil par value; 108,016,000 and 134,770,092 shares issued at June 30, 2004 and 2005, respectively	37,058	56,339
Deficit accumulated during the development stage	(10,427)	(20,872)
	26,631	35,467
	\$ 28,111	\$ 37,836

See accompanying notes to the financial statements.

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Statements of Operations**  
**(Australian dollars)**

	Years ended June 30,			Period from inception (May 29, 1998) to June 30, 2005
	2003	2004	2005	
	(in thousands, except per share data)			
Revenue	\$ —	\$ —	\$ —	\$ —
Operating expenses:				
Research and development	925	4,806	7,885	14,664
General and administrative	981	2,182	3,105	6,550
Commercial	—	—	807	807
Amortization of intangible assets	86	89	90	485
Fair value of stock options issued to employees related to:				
Research and development	261	253	115	753
Commercial	—	—	116	116
General and administrative	122	279	29	527
Total operating expenses	2,375	7,609	12,147	23,902
Loss from operations	(2,375)	(7,609)	(12,147)	(23,902)
Interest and other income	327	1,123	1,702	3,256
Amortization of preference share issue expenses	(65)	(161)	—	(226)
Net loss	\$(2,113)	\$(6,647)	\$(10,445)	\$(20,872)
Basic and diluted net loss per share	\$ (0.19)	\$ (0.09)	\$ (0.08)	\$ (0.61)

See accompanying notes to the financial statements.

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Statements of Changes in Shareholders' (Deficit) Equity**  
**(Australian dollars)**

	Ordinary shares		Accumulated Deficit	Total Shareholders' (Deficit) Equity
	Shares	Amount		
	(in thousands, except share data)			
Balance at inception (May 29, 1998)	—	—	—	—
Issuance of ordinary shares at \$0.125 per share for cash on incorporation of the Company in May 1998	808	—	—	—
Issuance of ordinary shares at \$0.125 per share as consideration for patent license	11,200,000	1,400	—	1,400
Fair value of stock options issued to directors and employees in year ended June 30, 2000	—	74	—	74
Fair value of stock options issued to directors and employees in year ended June 30, 2001	—	77	—	77
Fair value of stock options issued to directors and employees in year ended June 30, 2002	—	70	—	70
Net loss for period from inception to June 30, 2002	—	—	(1,667)	(1,667)
<b>Balance at June 30, 2002</b>	<b>11,200,808</b>	<b>1,621</b>	<b>(1,667)</b>	<b>(46)</b>
Cancellation of common stock	(808)	—	—	—
Fair value of stock options issued to directors and employees	—	383	—	383
Net loss	—	—	(2,113)	(2,113)
<b>Balance at June 30, 2003</b>	<b>11,200,000</b>	<b>2,004</b>	<b>(3,780)</b>	<b>(1,776)</b>
Conversion of "A" and "B" class convertible redeemable preference shares to ordinary shares	46,816,000	11,631	—	11,631
Issuance of ordinary shares at \$0.50 per share for cash in initial public offering of Company in Australia in November 2003, net of issuance costs	50,000,000	22,891	—	22,891
Fair value of stock options issued to directors and employees	—	532	—	532
Net loss	—	—	(6,647)	(6,647)
<b>Balance at June 30, 2004</b>	<b>108,016,000</b>	<b>37,058</b>	<b>(10,427)</b>	<b>26,631</b>
Issuance of ordinary shares at \$0.75 per share for cash in private placement to Australian institutional and sophisticated investors closed in December 2004, net of issuance costs	22,000,000	15,702	—	15,702
Share purchase plan closed in December 2004 at \$0.75 per share, net of issuance costs	4,362,092	3,256	—	3,256
Issuance of ordinary shares upon exercise of employee options	392,000	63	—	63
Fair value of stock options issued to directors and employees	—	260	—	260
Net loss	—	—	(10,445)	(10,445)
<b>Balance at June 30, 2005</b>	<b>134,770,092</b>	<b>\$56,339</b>	<b>\$ (20,872)</b>	<b>\$ 35,467</b>

See accompanying notes to the financial statements.

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Statements of Cash Flows**  
**(Australian dollars)**

	Years ended June 30,			Period from inception to June 30, 2005
	2003	2004	2005	
(in thousands)				
<b>Cash flows from operating activities:</b>				
Net loss	\$(2,113)	\$(6,647)	\$(10,445)	\$ (20,872)
Adjustments to reconcile net loss to net cash used in operating activities:				
Depreciation	142	353	489	1,064
Amortization of intangible assets	86	89	90	485
Amortization of preference share issue expenses	65	161	—	226
Fair value of stock options issued for service	383	532	260	1,396
Change in assets and liabilities:				
Prepaid expenses and other current assets	(15)	(180)	(454)	(718)
Grant receivable	(63)	63	(106)	(106)
Other long-term assets	(245)	(16)	(2)	(262)
Accounts payable	(8)	132	512	757
Accrued compensation	94	196	190	484
Accrued clinical development liabilities	13	793	(260)	546
Other accrued liabilities	317	(244)	446	582
Net cash used in operating activities	(1,344)	(4,768)	(9,280)	(16,418)
<b>Cash flows from investing activities:</b>				
Purchases of property, plant and equipment	(1,344)	(360)	(1,541)	(3,440)
Payment for patent applications	(84)	(46)	(33)	(191)
Net cash used in investing activities	(1,428)	(406)	(1,574)	(3,631)
<b>Cash flows from financing activities:</b>				
Proceeds from issuance of ordinary shares, net of issuance costs	—	22,891	18,958	41,849
Proceeds from issuance of ordinary shares from the exercise of employee options	—	—	63	63
Proceeds from issuance of "A" class convertible redeemable preference shares, net of issuance costs	—	—	—	2,000
Proceeds from issuance of "B" class convertible redeemable preference shares, net of issuance costs	9,405	—	—	9,405
Net cash provided by financing activities	9,405	22,891	19,021	53,317
Net increase in cash and cash equivalents	6,633	17,717	8,167	33,268
Cash and cash equivalents at beginning of period	751	7,384	25,101	—
Cash and cash equivalents at end of period	\$ 7,384	\$ 25,101	\$ 33,268	\$ 33,268
<b>Supplemental non-cash activities:</b>				
Issuance of ordinary shares on conversion of "A" and "B" class convertible redeemable preference shares	\$ —	\$ 11,631	\$ —	\$ 11,631

See accompanying notes to the financial statements.

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Notes to Financial Statements**  
**(for the years ended June 30, 2003, 2004 and 2005 and for the period from inception**  
**(May 29, 1998) to June 30, 2005)**  
**(in A\$ thousands, except share and per share amounts)**

**1. Organization of the Company**

Pharmaxis Ltd (the "Company") was incorporated on May 29, 1998 in the Australian Capital Territory, Australia. On June 6, 2002 the Company changed its name from Praxis Pharmaceuticals Australia Pty Ltd to Pharmaxis Pty Ltd. On September 5, 2003 the Company changed its name from Pharmaxis Pty Ltd to Pharmaxis Ltd. The Company was listed on the Australian Stock Exchange on November 10, 2003 following its initial public offering and on the NASDAQ National Market on August 29, 2005.

The Company's primary activities since incorporation have been the purchase of a license for our autoimmune technology from the Australian National University; conducting research in the area of autoimmune diseases; the filing of three new patent families; commencing preclinical development of one autoimmune candidate (PXS25); obtaining a license for our respiratory technology from the Central Sydney Area Health Service; establishing our own manufacturing facility for the production of Aridol and Bronchitol to standards of Good Manufacturing Practice; obtaining a license to manufacture to GMP from the Australian Therapeutics Goods Administration; commencing and successfully completing a 646 patient, 12 center Phase III clinical trial of our lung capacity test Aridol for the diagnosis of the existence and severity of asthma; commencing and successfully completing a 60 patient, 4 center Phase II clinical trial of Bronchitol for the disease bronchiectasis; commencing and successfully completing a 59 patient, eight center Phase II clinical trial of Bronchitol for the disease cystic fibrosis; establishing a management team with relevant experience in sales and marketing, manufacturing, clinical trial management, regulatory affairs and finance; and listing the Company on the Australian Stock Exchange and the NASDAQ National Market.

**2. Basis of Presentation**

Pharmaxis Ltd is an Australian company and its operations are located there. Accordingly, its books of account are maintained in Australian dollars and its annual and interim financial statements are prepared in accordance with accounting principles generally accepted in Australia ("A GAAP"). These financial statements are presented in accordance with the accounting principles generally accepted in the United States of America ("U.S. GAAP"). All amounts are expressed in Australian dollars and are rounded to the nearest thousand, except share and per share data.

The Company's financial statements have been prepared assuming the Company will continue as a going concern. The Company has sustained operating losses since inception and expects such losses to continue as it furthers its research and development programs.

**3. Summary of Significant Accounting Policies**

*Use of Estimates*

The preparation of financial statements in conformity with accounting principles generally accepted in the United States of America requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, disclosures of contingent assets and liabilities at the date of the financial statements and the reported amounts of revenues and expenses during the reporting period. Significant management judgment is required to make estimates in relation to: the carrying value and recoverability of intangible assets and property, plant and equipment; the recoverability of deferred income taxes; and the accruing of liabilities for clinical and preclinical development activities. Actual results could differ from those estimates.



### *Cash and Cash Equivalents*

The Company invests its excess cash in bank deposits, money market accounts and bank accepted commercial bills. The Company considers all highly liquid investments with an original maturity of 90 days or less at the time of purchase to be cash equivalents.

Bank accepted commercial bills are acquired at a discount to their face value. The bills are carried at cost plus a portion of the discount recognised as income on an effective yield basis. The discount brought to account each period is accounted for as interest received.

### *Concentration of Credit Risk and Other Risks and Uncertainties*

Cash and cash equivalents consist of financial instruments that potentially subject the Company to concentration of credit risk to the extent of the amount recorded on the balance sheet. The Company's cash and cash equivalents are invested with three of Australia's four largest banks. The Company is exposed to credit risk in the event of default by the banks holding the cash and cash equivalents to the extent of the amount recorded on the balance sheets. The Company has not experienced any losses on its deposits of cash and cash equivalents.

Product candidates developed by the Company may require approvals or clearances from the U.S. Food and Drug Administration ("FDA") or other international regulatory agencies prior to commercialized sales. There can be no assurance that the Company's product candidates will receive any of the required approvals or clearances. If the Company was denied approval or clearance or such approval was delayed, it may have a material adverse impact on the Company.

### *Foreign currency and currency transactions*

Monetary assets and liabilities in foreign currencies are translated into Australian dollars at the rate ruling at the date of transaction. Monetary assets and liabilities denominated in foreign currencies are retranslated into Australian dollars at the rate of exchange ruling at the balance sheet date. Transaction gains and losses are recognized in the statement of operations. The Company has no assets denominated in foreign currencies and immediately settles liabilities denominated in foreign currencies.

The Company has recorded foreign currency transaction losses of \$0, \$0, \$0 and \$0 for each of the years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005, respectively. The Company has recorded foreign currency transaction gains of \$2, \$0, \$0 and \$2 for each of the years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005, respectively.

The Company's functional and reporting currency is the Australian dollar.

### *Fair Value of Financial Instruments*

For financial instruments consisting of cash and cash equivalents, accounts payable and accrued liabilities included in the Company's financial statements, the carrying amounts are reasonable estimates of fair value due to their short maturities.

### *Property, Plant and Equipment*

Property, plant and equipment are recorded at acquisition cost, less accumulated depreciation and amortization. Depreciation and amortization are calculated on a straight-line basis over the estimated useful lives of the related assets, which are generally three to ten years. Amortization of leasehold improvements is computed using the straight-line method over the shorter of the remaining lease term or the estimated useful life of the related assets, typically three years. Maintenance and repairs are charged to operations as incurred.

The Company receives grants under certain government research agreements to purchase plant and equipment. The grants are recognized against the acquisition costs of the related plant and equipment as and when the related assets are purchased. Grants received in advance of the relevant expenditure are treated as deferred research grants and included in Other Accrued Liabilities on the balance sheet as the Company does not control the monies until the relevant expenditure has been incurred.

#### *Impairment of Long-Lived Assets*

The Company reviews its capital assets, including patents and licenses, for impairment whenever events or changes in business circumstances indicate that the carrying amount of the assets may not be fully recoverable. In performing the review, the Company estimates undiscounted cash flows from products under development that are covered by these patents and licenses. An impairment loss would be recognized when estimated undiscounted future cash flows expected to result from the use of the asset and its eventual disposition is less than the carrying amount of the asset. Impairment, if any, is measured as the amount by which the carrying amount of the asset exceeds its fair value. Impairment, if any, is assessed using discounted cash flows. Related patents are grouped in estimating future cash flows to determine whether patents are impaired and in measuring the amount of the impairment. Through June 30, 2005, there have been no such impairments.

#### *Interest Income*

Interest revenue is recognised as it accrues, taking into account the effective yield on the financial instruments.

#### *Research and Development*

Research and development expenses consist of costs incurred to further the Company's research and development activities and include salaries and related employee benefits, costs associated with clinical trials and preclinical development, regulatory activities, research-related overhead expenses, costs associated with the manufacture of clinical trial material, costs associated with developing a commercial manufacturing process, costs for consultants and related contract research, facility costs and depreciation. Research and development costs are expensed as incurred.

The Company receives grants under certain government research agreements that partially fund eligible research expenditure on certain of the Company's research projects. The grants are recognized against the related research and development expenses as and when the relevant research expenditure is incurred. Grants received in advance of incurring the relevant expenditure are treated as deferred research grants and included in Other Accrued Liabilities on the balance sheet as the Company does not control the monies until the relevant expenditure has been incurred. Grants due to the Company under research agreements are recorded as receivables and included on the balance sheet.

#### *Patent and License Costs*

Costs to purchase patent licenses and application costs for new patents are capitalised and amortised over the life of the patent. All of the Company's patents and licenses have finite lives with the remaining lives ranging from 11 to 20 years at June 30, 2005.

#### *Clinical Trial Expenses*

Clinical trial costs are a component of research and development expense. These expenses include fees paid to participating hospitals and other service providers, which conduct certain product development activities on behalf of the Company. Depending on the timing of payments to the service providers and the level of service provided, the Company records prepaid or accrued expenses relating to these costs.

These prepaid or accrued expenses are based on estimates of the work performed under service agreements, milestones achieved, patient enrollment and experience with similar contracts. Changes in these estimates are recognized in income in the period of change.

#### *Leased Assets*

All of the Company's leases for the years ended June 30, 2003, 2004 and 2005 are considered operating leases. The costs of operating leases are charged to the statement of operations on a straight-line basis over the lease term.

#### *Pension Costs*

As required by Australian law, the Company contributes to standard defined contribution superannuation funds on behalf of all employees at an amount up to nine percent of each such employee's salary. Superannuation is a

compulsory savings program whereby employers are required to pay a portion of an employee's remuneration to an approved superannuation fund that the employee is typically not able to access until they are retired. The Company permits employees to choose an approved and registered superannuation fund into which the contributions are paid. Contributions are charged to the statement of operations as they become payable.

#### *Income Taxes*

The Company applies Statement of Financial Accounting Standards No. 109 – Accounting for Income Taxes (SFAS 109) which establishes financial accounting and reporting standards for the effects of income taxes that result from a company's activities during the current and preceding years. Deferred tax assets and liabilities are recognized for the future tax consequences attributable to temporary differences between the financial statement carrying amounts of assets and liabilities and their respective tax bases. Deferred tax assets also are recognized for credit carryforwards. Deferred tax assets and liabilities are measured using the enacted rates applicable to taxable income in the years in which the temporary differences are expected to reverse and the credits are expected to be used. The effect on deferred tax assets and liabilities of a change in tax rates is recognized in income in the period that includes the enactment date. An assessment is made as to whether or not it is more likely than not that deferred tax assets are recoverable. This assessment includes anticipating future taxable income and the Company's tax planning strategies and is made on an ongoing basis. Consequently, future material changes in the valuation allowance are possible.

#### *Net Loss Per Share and Anti-dilutive Securities*

Basic and diluted net loss per share are presented in conformity with Statement of Financial Accounting Standards No. 128 – Earnings Per Share (SFAS 128). Basic and diluted net loss per share have been computed using the weighted-average number of ordinary shares outstanding during the period. All the share information presented for the fiscal years 2003 and 2004 and for the period from inception to June 30, 2005 have been retroactively adjusted to give effect to the 8 for 1 share split in November 2003. The potentially dilutive options issued under the Pharmaxis Employee Option Plan and the "A" and "B" class convertible redeemable preference shares were not considered in the computation of diluted net loss per share because they would be anti-dilutive.

#### *Stock-based Compensation*

The Company accounts for stock-based employee compensation arrangements using the fair value based method as prescribed in accordance with the provisions of Statement of Financial Accounting Standards No. 123 – *Accounting for Stock-Based Compensation* (SFAS 123).

#### *Recent Accounting Pronouncements*

In December 2004, the Financial Accounting Standards Board issued Statement of Financial Accounting Standards No. 123 revised – Share-Based Payment (SFAS 123R), an amendment of Statements No. 123 (SFAS 123) and 95 (SFAS 95) that addressed the accounting for share-based payment transactions in which an enterprise receives employee services in exchange for either equity instruments of the enterprise or liabilities that are based on the fair value of the enterprise's equity instruments or that may be settled by the issuance of such equity instruments. This statement eliminates the ability to account for share-based compensation transactions using the intrinsic value method as prescribed by Accounting Principles Board Opinion No. 25 – Accounting for Stock Issued to Employees (APB 25), and requires that such transactions be accounted for using a fair-value-based method and recognized as expenses in the statement of operations. As of the required effective date, the standard requires that the modified prospective method be used, which requires that the fair value of new awards granted on or following the effective date (plus unvested awards as of the effective date) be expensed over the vesting period. In addition, the statement encourages the use of the "binomial" approach to value stock options, which differs from the Black-Scholes option pricing model that the Company currently uses. Further, in March 2005, SEC issued Staff Accounting Bulletin 107 (SAB 107) providing guidance on the application of SFAS 123R. Further, as per a new rule approved by SEC in April 2005, SFAS 123R will be effective for public companies' annual, rather than interim, periods that begin after June 15, 2005. The adoption of SFAS 123R and SAB 107 is not expected to have a significant impact on its statement of operations as the Company currently expenses the fair value of its stock option grants.

In May 2005, the Financial Accounting Standards Board issued Statement of Financial Accounting Standards No. 154 – Accounting Changes and Error Corrections (SFAS 154), a replacement of Accounting Principles Board Opinion No. 20 (APB 20) and Statement No. 3 (SFAS 3), which previously addressed accounting changes. SFAS

154 establishes, unless impracticable, retrospective application as the required method for reporting a change in accounting principle in the absence of explicit transition requirements specific to the newly adopted accounting principle. SFAS 154 also provides guidance for determining whether retrospective application of a change in accounting principle is impracticable and for reporting a change when retrospective application is impracticable. SFAS 154 carries forward without change the guidance in APB 20 for reporting the correction of an error in previously issued financial statements. SFAS 154 will be effective for accounting changes and corrections of errors made in fiscal years beginning after December 15, 2005. The adoption of this standard will not have a material impact on the Company's financial statements.

#### 4. Related Party Transactions

In November 2003, The Principals Funds Management Pty Ltd, an entity in which the Company's directors DM Hanley and CPH Kiefel hold shares, was paid a fee of \$45 by Wilson HTM Corporate Finance Ltd, the underwriter and lead manager of the Company's Australian initial public offering, as consideration for a firm commitment by Principals Funds Management Pty Ltd to subscribe for shares in the Australian initial public offering.

The Principals Funds Management Pty Ltd was paid a consulting fee of \$108 during 2003, for services provided by Mr CPH Kiefel in relation to the "B" class private share issue to venture capital funds. Mr Kiefel was not a director of the Company at the time.

B.H. Smith is associated with GBS Venture Partners Ltd, The Australian Bioscience Trust and Bioscience Ventures II. Perpetual Trustees Nominees, as trustee of The Australian Bioscience Trust, received 45,000 shares from Wilson HTM Corporate Finance Ltd as consideration for a firm commitment to subscribe for shares in the Australian initial public offering in November 2003. GBS Venture Partners Ltd, as trustee and manager of Bioscience Ventures II, received 180,000 shares from Wilson HTM Corporate Finance Ltd as consideration for a firm commitment to subscribe for shares in the Australian initial public offering in November 2003.

C.J. Hillyard is associated with CM Capital Investments Pty Ltd, CM Capital Venture Trust No. 3, CIBC Australia Fund LLC and the Australia Venture Capital Fund L.P. CM Capital Investments Pty Ltd, as trustee of the CM Capital Venture Trust No. 3, received 171,777 shares from Wilson HTM Corporate Finance Ltd as consideration for a firm commitment to subscribe for shares in the Australian initial public offering in November 2003. CIBC Australia Fund LLC, as general partner of the Australia Venture Capital Fund L.P., received 53,223 shares from Wilson HTM Corporate Finance Ltd as consideration for a firm commitment to subscribe for shares in the Australian initial public offering in November 2003.

#### 5. Property, Plant and Equipment

Property, plant and equipment consist of the following:

	As of June 30,	
	2004	2005
	(in thousands)	
Plant and equipment	\$1,700	\$ 2,966
Leasehold improvements	152	166
Motor vehicles	47	92
Capital work in process	—	216
	<u>1,899</u>	<u>3,440</u>
Accumulated depreciation and amortization	(575)	(1,064)
Property, plant and equipment, net	<u>\$1,324</u>	<u>\$ 2,376</u>

Depreciation and amortization expense was \$142, \$353, \$489 and \$1,064 for the fiscal years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005, respectively.

## 6. Intangible Assets

Intangible assets consist of the following:

	As of June 30,	
	2004	2005
	(in thousands)	
Patent license	\$1,400	\$1,400
Patent applications	158	191
	1,558	1,591
Accumulated amortization	(396)	(485)
Intangible assets, net	\$1,162	\$1,106

The patent license was acquired in October 1999 from Anutech Pty Limited, an agent for the Australian National University. A license to the acquired technology was previously held by Praxis Pharmaceuticals Inc, a related party. The license was granted to Pharmaxis Ltd by Anutech at the request of Praxis Pharmaceuticals Inc, in consideration for which Praxis Pharmaceuticals Inc was issued 1.4 million ordinary shares in the Company. The license was included in long-term assets on the balance sheet at its estimated fair value of \$1,400 at the time of the purchase. The patents underlying the license relate to potential treatments for autoimmune diseases. The license is an exclusive, worldwide sub-licensable license within the field of phosphosugars as ethical therapeutics. A royalty is payable to Anutech, calculated based on revenue, net of expenses, received by the Company in connection with the Company's use of the related intellectual property. In addition the Company must reimburse Anutech for one third of any costs incurred in filing, maintaining and renewing the patents underlying the license.

## 7. Other Accrued Liabilities

Other accrued liabilities consist of the following:

	As of June 30,	
	2004	2005
	(in thousands)	
Deferred research grants	\$ 23	\$ 55
Legal and accounting fees	—	106
Manufacturing consultants	—	230
Other	112	191
	<u>\$ 135</u>	<u>\$ 582</u>

## 8. Central Sydney Area Health Service (“CSAHS”) License

In October 2001 the Company obtained an exclusive, worldwide sub-licensable license from CSAHS in relation to certain patents in the area of respiratory disease. The term of the CSAHS license is country specific and is for the longer of 10 years from the first commercial sale of products which exploits the CSAHS intellectual property in that country until the expiry of the last registered patent in that country. The Company must bear the cost of maintaining the registered CSAHS intellectual property and must use its reasonable commercial endeavors to exploit and undertake research and development of the CSAHS intellectual property. Royalties are payable by the Company to CSAHS for the term of the CSAHS license on net sales of products and services which exploit the CSAHS intellectual property. No amounts were paid or are payable under the license.

## 9. Ordinary Shares

In fiscal 2004, as part of the transition to an Australian listed public company, shareholders approved an 8 for 1 share split of all “A” and “B” class convertible redeemable preference shares, ordinary shares and other securities; a change in the Company’s status from a proprietary to a public company; the adoption of a new constitution; and, following the share split all “A” and “B” class convertible redeemable preference shares were converted to ordinary shares. Before the 8 for 1 share split the Company had on issue 1,400,000 ordinary shares, 2,000,000 “A” class convertible redeemable preference shares and 3,852,000 “B” class convertible redeemable preference shares. After the 8 for 1 share split there were 11,200,000 ordinary shares, 16,000,000 “A” class convertible redeemable preference shares and 30,816,000 “B” class convertible redeemable preference shares. After the conversion the Company had 50,816,000 ordinary shares. All share and per share amounts for the fiscal years 2003 and 2004 and the period from inception to June 30, 2005 presented in the accompanying financial statements have been retroactively adjusted to give effect to the share split.

The Company completed its initial public offering and listed on the Australian Stock Exchange on November 10, 2003, issuing 50 million shares and raising \$25,000 before costs associated with the issue.

In fiscal 2005 the Company issued 22 million ordinary shares in a private placement to institutional and sophisticated investors, 4,362,092 ordinary shares in a share purchase plan offered to existing shareholders, and 372,000 ordinary shares upon the exercise of employee options. At June 30, 2005 the Company had 134,770,092 ordinary shares on issue.

Ordinary shares entitle the holder to participate in dividends and the proceeds on winding up of the Company in proportion to the number of and amounts paid on the shares held. At a general meeting every shareholder present (in person or by proxy, attorney or representative) has one vote on a show of hands. Every shareholder present (in person or by proxy, attorney or representative) has one vote per fully paid share on a poll.

## 10. Convertible Redeemable Preference Shares

Up until the time of the Company’s Australian initial public offering, the Company had on issue 16,000,000 “A” and 30,816,000 “B” class convertible redeemable preference shares.

The Company issued 29,920,000 “B” class convertible redeemable preference shares on August 28, 2002 and 896,000 “B” class convertible redeemable preference shares on May 2, 2003, raising a total of \$9,600 in 2003 before costs associated with the issues.

The “A” and “B” class converting preference shares were convertible by the holders into ordinary shares at any time or could be compulsorily converted at the time of an initial public offering, subject to certain conditions. The conversion ratio was one ordinary share per convertible redeemable preference share, subject to variation for capital reconstructions and share dilutions.

The holders of the “A” and “B” class convertible redeemable preference shares could redeem their shares after October 15, 2005 if the Company had not listed on a securities exchange, on the winding up of the Company, or the disposal of its business or on a voluntary merger of the Company. The redemption price payable to the holders was their original issue price adjusted for any capital reconstructions plus any unpaid accrued cumulative dividends and a prorata share of surplus assets and profits at redemption date. On redemption “B” class convertible redeemable preference shares had priority over “A” class shares.

Both “A” and “B” class convertible redeemable preference shares were entitled to a 10% per annum cumulative dividend (based on original issue prices), “B” class having priority over “A” class. The dividends were only payable on the occurrence of a defined liquidity event or a voluntary merger of the Company. The dividend was not accrued as the events requiring the dividend to be paid were considered uncertain to occur and not probable. All rights to preference dividends lapsed upon the Company’s Australian listing in November 2003.

### **11. Stock Option Plan**

The Pharmaxis Employee Option Plan (“EOP”) was approved by shareholders in 1999 and amended by shareholders in June 2003. The maximum number of options available to be issued under the EOP is 15% of total issued shares including the EOP. All employees and directors are eligible to participate in the EOP, but do so at the invitation of the Board. The terms of option issues are determined by the Board. Options are generally granted for no consideration, have a life of ten years and generally vest equally over a four year period. For options granted after January 1, 2003 the annual vesting is subject to approval by the Remuneration and Nomination Committee of the Board. The Committee gives its approval for vesting based on the achievement of individual employee’s personal annual objectives. Upon a Liquidity Event, all unvested awards will become immediately exercisable. A Liquidity Event is defined as a response issued by the Company in respect of a takeover offer for all the shares of the Company.

Options granted under the EOP carry no dividend or voting rights. When exercisable, each option is convertible into one ordinary share.

The exercise price is set by the Board. Before the Company listed on the Australian Stock Exchange (“ASX”) in November 2003, the Board set the exercise price based on its assessment of the market value of the underlying shares at the time of grant. Since listing the exercise price is set as the average closing price of Pharmaxis Ltd shares on the Australian Stock Exchange on the five business days prior to the grant of the options.

A total of 392,000 options have been exercised to June 30, 2005, all in the year ended June 30, 2005.

There were 8,792,250 vested options at June 30, 2005 (7,206,500 at June 30, 2004) with a weighted average exercise price of \$0.308 (\$0.235 at June 30, 2004). A total of 6,720,000 options are subject to escrow by the ASX and cannot be exercised until November 10, 2005 (of which 5,940,000 are vested at June 30, 2005). The average remaining life of options outstanding at June 30, 2005 is 6.72 years (7.5 years at June 30, 2004). All share and option amounts for the fiscal years 2003 and 2004 and for the period from inception to June 30, 2005 have been retroactively adjusted to give effect to the share split described in note 9 above.

The following table summarizes stock option activity under the EOP:

	<u>Options Outstanding</u>	<u>Weighted-Average Exercise Price</u>	<u>Weighted-Average Fair Value</u>
Options granted	2,400,000	\$ 0.125	\$ 0.070
Balance at June 30, 2000	2,400,000	\$ 0.125	\$ 0.070
Options granted	640,000	\$ 0.125	\$ 0.069
Balance at June 30, 2001	3,040,000	\$ 0.125	\$ 0.070
Options granted	800,000	\$ 0.275	\$ 0.044
Balance at June 30, 2002	3,840,000	\$ 0.156	\$ 0.065
Options granted	5,344,000	\$ 0.313	\$ 0.168
Options cancelled/lapsed	(160,000)	\$ 0.125	\$ 0.070
Balance at June 30, 2003	9,024,000	\$ 0.249	\$ 0.126
Options granted	1,750,000	\$ 0.340	\$ 0.187
Options cancelled/lapsed	(23,000)	\$ 0.440	\$ 0.234
Balance at June 30, 2004	10,751,000	\$ 0.264	\$ 0.136
Options granted	605,000	\$ 1.005	\$ 0.548
Options cancelled/lapsed	(50,000)	\$ 0.43	\$ 0.228
Options exercised	(392,000)	\$ 0.159	\$ 0.102
Balance at June 30, 2005	10,914,000	\$ 0.308	\$ 0.160

The Company recorded stock compensation expense of \$383, \$532 and \$260, and \$1,397 in the fiscal years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005.

In accordance with SFAS 123, the fair values of the option grants were estimated on the date of each grant using the Black-Scholes option pricing model. The assumptions for these grants were:

<u>Grant Date</u>	<u>Exercise Price</u>	<u>Share Price at Grant Date</u>	<u>Volatility</u>	<u>Expected Life</u>	<u>Risk Free Interest Rate</u>
December 1, 1999	\$ 0.125	\$ 0.125	50%	6 years	6.45%
July 1, 2000	\$ 0.125	\$ 0.125	50%	6 years	6.03%
January 1, 2001	\$ 0.125	\$ 0.125	50%	6 years	5.31%
September 1, 2001	\$ 0.3125	\$ 0.125	50%	6 years	5.29%
December 2, 2001	\$ 0.125	\$ 0.125	50%	6 years	5.26%
May 12, 2003	\$ 0.3125	\$ 0.3125	50%	6 years	5.26%
July 1, 2003	\$ 0.3125	\$ 0.3125	50%	6 years	4.85%
July 4, 2003	\$ 0.3125	\$ 0.3125	50%	6 years	4.85%
December 9, 2003	\$ 0.376	\$ 0.376	50%	6 years	5.68%
April 25, 2004	\$ 0.508	\$ 0.508	50%	6 years	5.62%
June 4, 2004	\$ 0.426	\$ 0.426	50%	6 years	5.56%
February 2, 2005	\$ 0.834	\$ 0.834	50%	6 years	5.57%
May 12, 2005	\$ 1.147	\$ 1.147	50%	6 years	5.15%



The following table presents information relating to stock options outstanding under the plans as of June 30, 2005:

Exercise Price	Options Outstanding		Options Exercisable(1)
	Shares	Weighted Average Remaining Life in Years	Shares
\$ 0.125	2,720,000	4.6	2,720,000
\$ 0.3125	7,044,000	7.1	5,686,000
\$ 0.376	500,000	8.4	375,000
\$ 0.508	30,000	8.8	7,500
\$ 0.426	15,000	8.9	3,750
\$ 0.834	275,000	9.6	—
\$ 1.147	330,000	9.9	—

(1) Options Exercisable includes a total of 5,940,000 options that are vested at June 30, 2005, but are locked up and cannot be exercised until November 10, 2005.

The following table presents information relating to stock options outstanding under the plans as of June 30, 2004:

Exercise Price	Options Outstanding		Options Exercisable
	Shares	Weighted Average Remaining Life in Years	Shares
\$ 0.125	3,040,000	5.6	3,000,000
\$ 0.3125	7,136,000	8.1	4,144,000
\$ 0.376	500,000	9.4	62,500
\$ 0.508	60,000	9.7	—
\$ 0.426	15,000	9.9	—

## 12. Retirement Benefits

As required by Australian law, the Company contributes to standard defined contribution superannuation funds on behalf of all employees at an amount up to nine percent of employee salary. The Company permits employees to choose the superannuation fund into which the contributions are paid, provided the fund is appropriately registered.

The Company contributed \$69, \$125 and \$196, and \$391 for the fiscal years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005, respectively.

## 13. Income Taxes

The Company has made a taxable loss in each of the operating periods since incorporation.

A reconciliation of the (benefit) provision for income taxes with the amount computed by applying the statutory company tax rate of 30% to the loss before income taxes is as follows:

	Years ended June 30,						Period from inception to June 30, 2005	
	2003		2004		2005		\$	%
	\$	%	\$	%	\$	%		
	(in thousands)							
Loss before income taxes	(2,113)		(6,647)		(10,445)		(20,872)	
Income tax expense computed at statutory corporation tax rate	(634)	30	(1,994)	30	(3,134)	30	(6,262)	30
Research & development incentive	—	—	—	—	(819)	8	(819)	4
Disallowed expenses								
Stock option expense	115	(5)	160	(2)	78	(1)	419	(2)
Amortization of intangible assets	25	(1)	25	—	25	—	140	(1)
Other	—	—	1	—	3	—	5	—
Change in valuation allowance	494	(24)	1,808	(28)	3,847	(37)	6,517	(31)
<b>Income tax expense</b>	<b>\$ —</b>	<b>—</b>	<b>\$ —</b>	<b>—</b>	<b>\$ —</b>	<b>—</b>	<b>\$ —</b>	<b>—</b>

Significant components of the Company's deferred tax assets are shown below:

	As of June 30,	
	2004	2005
	(in thousands)	
Deferred tax assets (liabilities):		
Net operating loss carryforwards	2,673	6,491
Accrued compensation costs	33	68
Research grants offset to property, plant and equipment	45	30
Share issue expenses	548	801
<b>Total deferred tax assets</b>	<b>3,299</b>	<b>7,390</b>
Valuation allowance for deferred tax assets	(3,294)	(7,385)
	5	5
Patent cost amortization	(5)	(5)
<b>Net deferred taxes</b>	<b>—</b>	<b>—</b>

Deferred income taxes reflect the net tax effect of temporary differences between the carrying amounts of assets and liabilities for financial reporting and tax purposes. A valuation allowance has been established, as realization of such assets is not more likely than not.

At June 30, 2005 the Company has \$21.6 million (\$8.9 million at June 30, 2004) of accumulated tax losses available for carry forward against future earnings, which under Australian tax laws do not expire but may not be available under certain circumstances.

#### 14. Commitments and Contingent Liabilities

##### *Operating leases*

In October 2002, the Company entered into an operating lease for its headquarters and manufacturing facilities in Frenchs Forest, Sydney, Australia and moved into the facility in November 2002. The operating lease expires in June 2006, with an option to renew for a further five years thereafter. The Company's bankers have issued a bank guarantee of \$169 in relation to a rental bond to secure the payments under the lease. This bank guarantee is secured by a security deposit held at the bank. In 2003 and 2004 the Company leased additional office space in Canberra, Australia and the Company also leases equipment from time to time as required.

The Company recognized rent expense of \$238, \$346 and \$327, and \$972 for the fiscal years ended June 30, 2003, 2004 and 2005, and the period from inception to June 30, 2005, respectively.

Future minimum lease payments under all non-cancelable operating leases are as follows:

Years ending June 30,	
2006	\$322
2007 and thereafter	—
<b>Total minimum lease payments required</b>	<b>\$322</b>

#### *Government research grants*

The Company has received three separate Australian Government research grants under the R&D START Program; two of these grant programs were completed by June 30, 2003. The Government may require the Company to repay all or some of the amount of a particular grant together with interest in either of the following circumstances:

- the Company fails to use its best endeavors to commercialise the relevant grant project within a reasonable time of completion of the project; or
- upon termination of a grant due to breach of agreement or insolvency.

Technical failure of the grant funded research project does not of itself constitute failure to use best efforts to commercialise the relevant grant project. The grants have funded certain aspects of the Company's development projects for the Aridol asthma product, multiple sclerosis and cystic fibrosis. The Company continues the development of all three projects funded by the START Program and has commenced commercialisation of its asthma project as evidenced by the filing of marketing approval applications for Aridol with Australian and European Union regulatory agencies. The Company believes that the likelihood of being required to repay grant funding is remote while the Company continues to act in good faith with respect to the grants. The total amount received under the START Program at June 30, 2005 was \$4,200.

The Company has been awarded a research grant under the Australian Government's Pharmaceuticals Partnerships Program ("P3"). The Government may require the Company to repay all or some of the amount of the grant together with interest in any of the following circumstances:

- a) The Government determines that expenditures claimed on research projects do not meet the P3 guidelines; or
- b) upon termination of the grant due to breach of agreement, change in control of the Company or insolvency.

The total amount received under the P3 Program at June 30, 2005 was \$55 which has been booked as deferred government research grants.

#### **15. Research Grants**

Research grants received have been accounted for as follows:

	Years ended June 30,			Period from inception to June 30, 2005
	2003	2004	2005	
Recognized against related research and development expenses	\$751	\$1,105	\$1,132	\$ 4,551
Recognized against the acquisition cost of related plant and equipment	225	—	—	225
	<b>\$976</b>	<b>\$1,105</b>	<b>\$1,132</b>	<b>\$ 4,776</b>

Grants received in advance of incurring the relevant expenditure are treated as deferred research grants and included in Other Accrued Liabilities on the balance sheet as the Company does not control the monies until the relevant expenditure has been incurred. Grants due to the Company under research agreements are recorded as receivables and included on the balance sheet.

## 16. Guarantees and Indemnifications

Pharmaxis Ltd has entered into Deeds of Access to Documents and Indemnity agreements with certain executive officers and each of its directors. Each deed provides each respective officer with the following:

- a right to access certain board papers of the Company during the period of their tenure and for a period of seven years after that tenure ends;
- subject to the Corporations Act, an indemnity in respect of liability to persons other than the Company and its related bodies corporate that they may incur while acting in their capacity as an officer of the Company or a related body corporate, except where that liability involves a lack of good faith and for defending certain legal proceedings; and
- the requirement that the Company maintain appropriate directors' and officers' insurance for the officer.

No liability has arisen under these indemnities as at June 30, 2005.

## 17. Net Loss per Share

Basic net loss per ordinary share was computed by dividing the net loss applicable to ordinary shares by the weighted-average number of ordinary shares and contingently issuable shares outstanding during the period. Diluted net loss per ordinary share was computed by dividing the net loss applicable to ordinary shares by the weighted-average number of ordinary shares, contingently issuable shares and convertible redeemable preferred shares outstanding during the period. All periods presented in the financial statements have been retroactively adjusted to give effect to the 8 for 1 share split that occurred prior to the Company's Australian initial public offering in November 2003. Options granted to employees under the Pharmaxis Employee Option Plan are considered to be potential ordinary shares for the purpose of calculating diluted net loss per share. However, all such issued options outstanding were not included in the calculation of diluted net loss per share as the effect of including such options is anti-dilutive.

	Years ended June 30,			Period from inception to June 30, 2005
	2003	2004	2005	
Weighted average number of ordinary shares used as the denominator in calculating basic and diluted net loss per share.	11,200,000	75,744,000	123,933,133	34,068,056

## 18. Segments

The Company operates in one segment. The principal activities of the Company are the research, development and commercialisation of pharmaceutical products.

The Company operates predominantly in one geographical area, being Australia.

## 19. Subsequent Events

On August 5, 2005 the Company announced that, subsequent to a review of employee and director performance for the year ended June 30, 2005, the directors proposed to grant 954,500 options under the Pharmaxis Employee Option Plan. The terms of the options granted under the Plan are set out in note 11. The exercise price was calculated as \$1.79. The grant of options to directors requires shareholder approval and therefore 335,000 of the proposed option grants is contingent upon a favorable vote by shareholders at the Company's annual meeting on November 15, 2005.

On November 10, 2005 the Company completed a public offering of 19,500,000 ordinary shares in the form of 1,300,000 American Depositary Shares in the United States and received net proceeds of \$28,300. Concurrently the Company placed 19,900,000 ordinary shares in Australia and received net proceeds of \$30,200. Both offerings were approved by the Company's shareholders at the general meeting held on October 28, 2005.

**Pharmaxis Ltd**  
**(A Development Stage Enterprise)**

**Schedule II – Valuation and Qualifying Accounts**  
**(for the years ended June 30, 2003, 2004 and 2005 and for the period**  
**from inception (May 29, 1998) to June 30, 2005)**  
**(in A\$ thousands, except share and per share amounts)**

<u>Description at</u>	<u>Balance at Beginning of Period</u>	<u>Additions</u>		<u>Deductions</u>	<u>Balance at End of Period</u>
		<u>Charged to Costs and Expenses</u>	<u>Charged to Other Accounts(1)</u>		
<b>Year ended June 30, 2003</b>					
Deferred income tax valuation allowance	358	494	2	—	854
<b>Year ended June 30, 2004</b>					
Deferred income tax valuation allowance	854	1,808	632	—	3,294
<b>Year ended June 30, 2005</b>					
Deferred income tax valuation allowance	3,294	3,847	244	—	7,385
<b>Period from inception to June 30, 2005</b>					
Deferred income tax valuation allowance	—	6,517	868	—	7,385

(1) Recognized through equity as a result of share issue costs.

## **EXHIBITS**

11. Code of Conduct
12. Certification required by Rule 13a-14(a)
13. Certification pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002

## **EXHIBIT 11**

### **Pharmaxis Ltd Code of Conduct**

#### **1. Introduction**

Pharmaxis Ltd is committed to:

- conducting its business in a way that is accountable to shareholders and the wider financial marketplace
- being a company respected for integrity in all its dealings
- being an employer that provides opportunity and reward in an environment of trust and respect between all employees regardless of position or function
- earning the trust of shareholders
- being respected and welcomed in the communities where it does business

The reputation and integrity of Pharmaxis relies on employees observing the highest standards of behaviour when engaging in corporate activity.

#### **2. Purpose of the Code**

This Code of Conduct (“Code”) provides a common behavioural framework for all employees in the conduct of Pharmaxis business, irrespective of their specific job, direct employer or location in the world. It is a key part of the Pharmaxis culture. It sets out the standards for our relationships with our customers, employees, shareholders, suppliers, competitors, clinical trial participants, research collaborators and the community. All employees are bound by the Code and are expected to comply with the Code when representing the company.

The objective of the Code is to ensure that:

- high standards of corporate and individual behaviour are observed by all Pharmaxis employees in the context of their employment with Pharmaxis
- employees are aware of their responsibilities to Pharmaxis under their contract of employment
- all persons dealing with Pharmaxis whether it be employees, shareholders, suppliers, customers or competitors can be guided by the stated values and policies of Pharmaxis

For the purpose of this document “employees” includes Pharmaxis contractors and directors.

#### **3. General principles**

- Pharmaxis employees will conduct themselves with integrity in all areas and at all times.
- Pharmaxis will apply appropriate industry best practices to the health, safety and wellbeing of its employees, customers, suppliers, clinical trial participants and the communities in which it operates.
- Pharmaxis employees will aim to bring long-term benefits to Pharmaxis rather than short-term advantage for individuals at the expense of the company’s long-term interests.
- Pharmaxis employees will recognize that effective teamwork requires cooperation with our colleagues across the Company to achieve our corporate goals.
- Pharmaxis employees will strive to ensure their words and actions show respect for the diverse range of people and cultures with whom they work and for their human rights.
- Pharmaxis and its employees will exercise due care and diligence in the performance of their duties and responsibilities.
- Assets and confidential information will be appropriately protected and must not be used by employees for personal gain.
- Employees who raise their concerns with management when they have genuine suspicions of wrongdoing anywhere in the organisation will be taken seriously and will not be victimised.

- Pharmaxis does not make political donations.

#### 4. Obligation to comply with code

Employees will adhere to the Code both in letter and in spirit. Violation of the Code by any employee, or unethical behaviour which may affect the reputation of Pharmaxis, may be subject to disciplinary action including termination of employment.

#### 5. Integrity

All Pharmaxis' employees are expected to conduct business in a manner that enhances the Company's reputation and integrity. The overriding principles which govern behaviour and business dealings are:

- All business should be conducted in accordance with the laws and regulations of the countries in which Pharmaxis operates.
- Pharmaxis will compete fairly in the markets in which it operates.
- Corrupt practices are unacceptable. No bribes or similar payments will be made to, or accepted from, any party. Business entertainment must be in compliance with the Pharmaxis general business expense policy and in the case of dealings with government agencies, within the rules and regulations of the agency concerned. Third parties must not be used to circumvent this policy.
- All commercial transactions will be properly and accurately recorded.
- Sales agents, consultants and similar advisers will be appointed in accordance with these principles and paid at a rate consistent with the value of their services.
- Conflicts of interest: Employees will disclose to the company any activities that involve, or could appear to involve, a conflict between their personal interests and the interests of Pharmaxis. Areas where conflicts might arise include: ownership interests in businesses with which Pharmaxis deals, share ownership, direct or indirect personal interest in contracts, seeking or accepting gifts or entertainment beyond levels considered reasonable in the business environment, employment outside of Pharmaxis or use of confidential information. Where a conflict of interest arises or may arise, an employee must disclose the conflict to his or her supervisor. The supervisor will ensure that measures are put in place to protect parties affected by the conflict of interests from that conflict. Depending upon the nature and extent of the conflict, the measures taken to protect against conflict of interest may include ensuring that the conflicted or potentially conflicted employee does not participate in the relevant decision or activity, or disclosing the conflict to third parties affected by the conflict and ensuring that the conflicted or potentially conflicted employee is not the sole decision maker on the matter. Employees who are directors of Pharmaxis or its subsidiaries will be guided by the constitution of Pharmaxis or the subsidiary and will ensure that they abide by the provisions in the Corporations Act or equivalent local legislation governing conflicts of interest. Where there is doubt the matter should be raised with the Chairman of Pharmaxis.
- Corporate opportunities: Employees will not take advantage of property, information or position, or opportunities arising from these, for personal gain or to compete with Pharmaxis.
- Confidentiality: Employees will not breach Pharmaxis's confidentiality or make use of confidential information obtained from Pharmaxis for personal gain or in a manner which would be detrimental to Pharmaxis. Confidential information of Pharmaxis is not to be used except in ways which are authorised by Pharmaxis or legally mandated. The same applies to confidential information belonging to third parties which an employee may obtain in the course of their duties. Pharmaxis will respect the confidentiality of its employees.
- Conduct of Clinical Trials: The well being of clinical trial participants is to be placed first. Company sponsored clinical trials will adhere to the principles of Good Clinical Practice.
- Use of company assets: Pharmaxis assets must be acquired, maintained and used in an efficient manner and for legitimate Pharmaxis business purposes. Pharmaxis assets also include patents, trade secrets and other proprietary and confidential information.
- Share trading policy: Employees will comply with the Pharmaxis Share Trading Policy.
- Continuous disclosure and shareholder communications: Employees will comply with the Pharmaxis Continuous Disclosure and Shareholder Communications Policy.



- Email: Employees are responsible for using the company's email and internet resources responsibly and for company use. All communications and information transmitted by, received from or created or stored in the company's information systems are the company's records and property.
- Improper Influence on Conduct of Audits: It is inappropriate behavior to attempt to improperly influence the Company's auditor.

## 6. Employment practices

Employees should be treated justly and fairly and rewarded for their achievements. Employees should feel that they can discuss, on a confidential basis, any problem associated with their employment with the Company. Pharmaxis employment policies include:

- Occupational health and safety: Pharmaxis is committed to providing a safe and healthy working environment and to working with contractors and suppliers that also operate a safe and healthy work environment. Pharmaxis can only provide a healthy and safe working environment with the cooperation of every employee. As part of that cooperation employees must:
  - Acquaint themselves with the requirements of all applicable occupational health and safety laws.
  - Apply best operating practice in our industry with regard to safety and health.
  - Regularly review risk management systems to properly identify, control and prevent accidents in the workplace. When accidents happen, employees should be treated fairly during the subsequent investigation.
  - Safety procedures should never be compromised to meet operational targets.
- Non-discrimination and sexual harassment: Pharmaxis is an equal opportunities employer. The company is committed to developing an appropriately qualified and experienced workforce, and providing a work environment in which everyone is treated fairly and with respect, irrespective of gender, race, sexual orientation, age, disability, religion or ethnic origin. Employment and advancement within Pharmaxis will be based on merit.
- Creating an environment where everyone is encouraged to give their best and realise their full potential.
- Freedom of association: Pharmaxis respects the individual's right to freedom of association.
- Creating an environment where employees can discuss any problems connected with their work confident that they will receive a fair, impartial and confidential review of the issue.
- Except in the case of summary dismissal for serious misconduct, giving employees a reasonable chance to remedy any inappropriate conduct or performance.

## 7. Responsibilities to the community:

- Environment: Pharmaxis values the natural environment and is committed to sound environmental practices in our operations. Pharmaxis employees are asked to take a thoughtful approach to environmental risk management. The company is required to comply with relevant environmental laws and regulations and adopts the following principles:
  - Thorough understanding of all relevant environmental laws, regulations and industry best practice relevant to the company's areas of operation.
  - Improving efficiency of resource usage.
  - Minimising emissions and waste.
  - Responding to community concerns with integrity, honesty and respect.
- Human rights: Pharmaxis supports and upholds the principles contained in the Universal Declaration on Human Rights. In particular Pharmaxis will not tolerate child labour or forced labour in the company's operations or those of our suppliers.
- Animal Welfare: The development of new pharmaceutical products is a highly regulated process, controlled on an individual country by country basis. Government agencies enforce stringent efficacy and safety testing standards that must be attained before pharmaceutical products can be administered to humans. These government mandated safety and efficacy standards include testing in non-human species.  
Pharmaxis seeks to minimize the number of tests it conducts involving animals.

The company's tests for efficacy and safety are conducted in a number of third party facilities situated in Australia, the USA and the EEC. The operations of these facilities are regulated by the legislation of Australia, the USA and the EEC respectively. Such regulation typically includes oversight of tests by an ethics committee responsible to ensure the tests are justified on a technical and ethical basis, and that the facility fully complies with local standards for the conduct of such tests.

#### **8. Reporting violations of the code of ethics or other unethical conduct**

Trust and integrity are vital to Pharmaxis. Misconduct and malpractice breach trust and endanger our reputation and business. The best way of protecting trust is for employees who have genuine suspicions about wrongdoing to know that they have a safe environment in which to speak up without fear of reprisal or victimisation. Employees who raise their concerns with management when they have genuine suspicions of wrongdoing will be taken seriously and will not be victimised. As Pharmaxis is currently a small company, employees should contact the chief executive officer or the chairman of the board with respect to ethical and conduct issues. Their concerns will be properly investigated and any necessary action taken.

Each employee shall receive a copy of this Code of Conduct. Management shall monitor compliance and implement special training and/or additional monitoring as appropriate.

## EXHIBIT 12

### CHIEF EXECUTIVE OFFICER CERTIFICATION

I, Alan D. Robertson, Chief Executive Officer of Pharmaxis Ltd (the "Registrant"), certify that:

1. I have reviewed this annual report on Form 20-F of the Registrant;

2. Based on my knowledge, this annual report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this annual report;

3. Based on my knowledge, the financial statements, and other financial information included in this annual report, fairly present in all material respects the financial condition, results of operations and cash flows of the Registrant as of, and for, the periods presented in this annual report;

4. The Registrant's other certifying officer and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) for the Registrant and have:

- a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the Registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this annual report is being prepared;
- b) Evaluated the effectiveness of the Registrant's disclosure controls and procedures and presented in this annual report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this annual report based on such evaluation; and
- c) Disclosed in this annual report any change in the Registrant's internal control over financial reporting that occurred during the period covered by this annual report that has materially affected, or is reasonably likely to materially affect, the Registrant's internal control over financial reporting; and

5. The Registrant's other certifying officer and I have disclosed, based on our most recent evaluation of internal controls over financial reporting, to the Registrant's auditors and the audit committee of the Registrant's board of directors:

- a) All significant deficiencies and material weaknesses in the design or operation of internal controls over financial reporting which are reasonably likely to adversely affect the Registrant's ability to record, process, summarize and report financial information; and
- b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the Registrant's internal controls over financial reporting.

**Pharmaxis Ltd**

Date: December 20, 2005

/s/ ALAN D. ROBERTSON  
Alan D. Robertson  
Chief Executive Officer

## CHIEF FINANCIAL OFFICER CERTIFICATION

I, David M. McGarvey, Chief Financial Officer of Pharmaxis Ltd (the "Registrant"), certify that:

1. I have reviewed this annual report on Form 20-F of the Registrant;
2. Based on my knowledge, this annual report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this annual report;
3. Based on my knowledge, the financial statements, and other financial information included in this annual report, fairly present in all material respects the financial condition, results of operations and cash flows of the Registrant as of, and for, the periods presented in this annual report;
4. The Registrant's other certifying officer and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) for the Registrant and have:
  - a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the Registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this annual report is being prepared;
  - b) Evaluated the effectiveness of the Registrant's disclosure controls and procedures and presented in this annual report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this annual report based on such evaluation; and
  - c) Disclosed in this annual report any change in the Registrant's internal control over financial reporting that occurred during the period covered by this annual report that has materially affected, or is reasonably likely to materially affect, the Registrant's internal control over financial reporting; and
5. The Registrant's other certifying officer and I have disclosed, based on our most recent evaluation of internal controls over financial reporting, to the Registrant's auditors and the audit committee of the Registrant's board of directors:
  - a) All significant deficiencies and material weaknesses in the design or operation of internal controls over financial reporting which are reasonably likely to adversely affect the Registrant's ability to record, process, summarize and report financial information; and
  - b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the Registrant's internal controls over financial reporting.

**Pharmaxis Ltd**

Date: December 20, 2005

/s/ DAVID M. MCGARVEY  
David M. McGarvey  
Chief Financial Officer

**EXHIBIT 13**

**CERTIFICATION PURSUANT TO  
18 U.S.C. SECTION 1350,  
AS ADOPTED PURSUANT TO  
SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002**

In connection with the Annual Report of Pharmaxis Ltd (the "Company") on Form 20-F for the year ended June 30, 2005 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I, Alan D. Robertson, the Chief Executive Officer of the Company, certify, pursuant to and for purposes of 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002, that, to my knowledge:

- (1) The Report fully complies with the requirements of Section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and results of operations of the Company.

Date: December 20, 2005

/s/ ALAN D. ROBERTSON  
**Alan D. Robertson**  
**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

In connection with the Annual Report of Pharmaxis Ltd (the "Company") on Form 20-F for the year ended June 30, 2005 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I, David M. McGarvey, the Chief Financial Officer of the Company, certify, pursuant to and for purposes of 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002, that, to my knowledge:

- (1) The Report fully complies with the requirements of Section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and results of operations of the Company.

Date: December 20, 2005

/s/ DAVID M. MCGARVEY  
David M. McGarvey  
Chief Financial Officer