

### **OCTOPLUS N.V.**

### (a limited liability company incorporated under the laws of the Netherlands, with its corporate seat in Leiden)

### Admission to listing and trading of 8,000,000 newly issued ordinary shares

On 31 October 2011, OctoPlus N.V. issued a total amount of 8,000,000 ordinary shares (the "New Shares") at a price of € 0.50 each (the "Private Placement") to new and existing investors, including CEO Jan Egberts and co-founder and former CEO Joost Holthuis.

In this Prospectus, the "Company", "OctoPlus", "we", "our", "us" and similar terms refer to OctoPlus N.V. and, where appropriate, its subsidiaries. Any reference to "Shares" shall refer to ordinary shares of the Company, including the New Shares, outstanding from time to time.

## Our business and any investments in our Shares involve significant risks. These risks are described under "Risk Factors" beginning on page 8 of this Prospectus.

The Shares outstanding immediately prior to the Private Placement are listed and traded on Euronext Amsterdam by NYSE Euronext ("Euronext Amsterdam") under the symbol "OCTO" and ISIN Code NL0000345718. We have applied for admission of 8,000,000 Shares issued pursuant to the Private Placement to listing and trading on Euronext Amsterdam and expect that trading in these Shares on Euronext Amsterdam will commence on or about 26 January 2012 (the "Listing Date").

This Prospectus does not constitute an offer to sell, or a solicitation of an offer to buy, any of the New Shares or any other securities issued by the Company.

The New Shares have not been and will not be registered under the United States Securities Act of 1933, as amended (the "Securities Act"). The New Shares were offered and sold (i) outside the United States in reliance on Regulation S under the Securities Act ("Regulation S") and (ii) inside the United States in reliance on Regulation D under the Securities Act ("Regulation D"). The New Shares may not be offered or sold within the United States or to, or for the account or benefit of, U.S. persons (as defined in Regulation S) except pursuant to an exemption from, or in a transaction not subject to, the registration requirements of the Securities Act.

The issue price for the New Shares has been determined on the basis of the quoted share price of the Shares and certain other factors.

This Prospectus constitutes a prospectus for the purposes of Article 3 of the Directive 2003/71/EC, as amended pursuant to Directive 2010/73/EU (the "Prospectus Directive") and has been prepared pursuant to Article 5:2 of the Financial Markets Supervision Act (*Wet op het financieel toezicht* (the "Financial Supervision Act")) and the rules promulgated thereunder. This Prospectus has been approved by and filed with the AFM.

Prospectus dated 24 January 2012

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### Summary

This summary provides an overview of selected information contained elsewhere in this Prospectus and should be read as an introduction to this Prospectus. Any decision to invest in our Shares should be based on consideration of this Prospectus as a whole by you. You should carefully read the Prospectus in its entirety before investing in our Shares, including the information discussed under "Risk Factors" beginning on page 7 and our consolidated financial statements and the notes thereto that are incorporated by reference in this Prospectus.

Under laws in effect in the states within the European Economic Area, no civil liability will attach to us in respect of this Summary, or any translation thereof, unless it is misleading, inaccurate or inconsistent when read together with the other parts of this Prospectus, or it does not provide, when read together with the other parts of this Prospectus, key information in order to aid investors when considering whether to invest in such securities. Where a claim relating to information contained in this Prospectus is brought before a court in a state within the European Economic Area, the plaintiff investor may, under the national legislation of the state where the claim is brought, be required to bear the costs of translating this Prospectus before the legal proceedings are initiated.

#### Summary of our Business

OctoPlus is a specialty pharmaceutical company that is globally renowned as a centre of excellence in formulation development and drug delivery. We offer clients our proven expertise in combination with our proprietary drug delivery technologies, in particular PolyActive, or other drug delivery platforms such as PLGA, for the development of controlled release formulations of injectable compounds. Such depot formulation products have demonstrated strong improvement of side effect profiles, increased patient compliance and better efficacy. In addition, we offer proven and reliable services in formulation development, process development, analytical method development and cGMP manufacturing. Our expertise is focused on formulation of biotech-derived compounds and small chemical molecules.

In October 2008, we made the strategic decision to focus exclusively on activities for which we are reimbursed. Such contracts include our well established contract formulation & manufacturing activities, but also relate to activities whereby we combine our proprietary drug delivery technologies with biopharmaceutical drugs or compounds of our partners in order to improve the properties of such product candidates. We have entered into several technology evaluation agreements to develop such controlled release formulations, whereby our clients incur the development costs and we are reimbursed for all or substantially all costs and will furthermore be entitled to potential future milestone and royalty payments. As a result of the change in strategy, existing product candidates have been and will be licensed to partners. We intend to continue to perform formulation, analytical and process development services and cGMP manufacturing for some of these product candidates, mainly Locteron<sup>®</sup>, on a fee-for-service basis. The organisational structure has been simplified and the relevant departments have been integrated. As a result, the Company has only one business unit from October 2009 onwards.

The clinically most advanced product based on our proprietary drug delivery technologies, Locteron, is being developed by Biolex, the holder of the commercial rights to Locteron. Locteron is a proprietary controlled release formulation of interferon alpha for the treatment of chronic hepatitis C infection (HCV). Hepatitis C is a common disease, affecting over 170 million people worldwide. Locteron combines interferon alpha produced by Biolex with our proprietary PolyActive microspheres. This product is designed to gradually release its active pharmaceutical ingredient over a 14-day period after a single injection. This will reduce the number of injections from 48 injections as per the current standard of care to 24 injections over the 48-week treatment period with Locteron. It is believed that an improved side effect profile, such as experienced with Locteron, will lead to enhanced patient compliance. In 2010, Biolex successfully completed a Phase IIb clinical trial with Locteron.

One of the areas in which we expect to grow is the development and production of specialty generics. Several blockbuster injectable controlled release drugs are coming off patent in the coming years, which opens the way to develop competitive products without infringing patents. It is expected that many companies would like to enter these multi-billion markets. Developing and manufacturing a generic version of an injectable controlled release drug is difficult and only a limited number of companies in the contract service business have the expertise and infrastructure to do so. We are experts in this field, as we have proven with the development and manufacturing of Locteron and many other products. During 2011, we have performed some early-stage formulation development work in this field on a fee-for-service basis and see an important and increasing role for OctoPlus in this field in the years to come. As the activities required to develop a generic drug are part of our standard service offering, significant investments are not required, certainly not in early stage development.

### Our activities

Our expertise is combining our drug delivery systems, in particular PolyActive, with our partners' biopharmaceutical drugs or other therapeutics to develop improved formulations of these biopharmaceutical drugs or other therapeutics. The improved products provide controlled and prolonged release of a drug, so as to enable reduced dosing frequency and reduced side effects with comparable or better efficacy relative to immediate release drugs. Through the use of our controlled release technologies, which are ideally suited for the controlled release of proteins, we believe that we can offer a significant differentiation to our client's product candidates and other therapeutic products. Our technologies are designed to have the advantage of efficiently and gradually delivering a drug in its native form. We believe that products based on our technologies can be applied in many therapeutic areas.

For other clients we also offer formulation services and manufacture clinical product without the use of our proprietary drug delivery technologies. Since our establishment in 1995, we have successfully provided these services on a fee-for-service basis to a diverse and international group of more than 160 pharmaceutical and biotechnology companies, focusing mainly on protein therapeutics and to a lesser extent on small molecule drugs. Typically, our clients are charged on an hourly basis for our development services, while manufacturing is charged on a lump-sum basis.

We have established a strong competitive position in the market for contract formulation and manufacturing of clinical-grade products, particularly in the area of pharmaceutical formulations for therapeutic proteins. We believe that our capabilities in the contract formulation and manufacturing business are demonstrated by our high rates of repeat business.

### **Our Strategy**

The key elements of our corporate strategy are explained in detail in the Business section, and include:

- Continue to grow our contract formulation and manufacturing business
- Build a portfolio of licensed products based on our proprietary technologies of which Locteron is an important value driver
- Build a portfolio of specialty generics products with partners

### **Risks Associated with Our Business**

Our business is subject to numerous risks, such as risks related to our business and strategy, our dependence on third parties, our intellectual property, our employees and growth, regulatory approval and other government regulations, and our financial condition. These risks are more fully described in the section entitled "Risk Factors" immediately following this Summary.

### Corporate Information

We are a public company with limited liability incorporated under the laws of the Netherlands and are registered with the Trade Register of the Chamber of Commerce of The Hague under number 28075073 and we have our corporate seat in Leiden, the Netherlands. Our business address is Zernikedreef 12, 2333 CL Leiden, the Netherlands and our website is <u>www.octoplus.nl</u>.

### Summary of Terms of the Private Placement

Issuer	OctoPlus N.V., a public company with limited liability incorporated under the laws of the Netherlands, with its corporate seat in Leiden, the Netherlands.
Private Placement	The Private Placement consisted of a private placement to qualified investors in the Netherlands and certain other jurisdictions (the "Investors") of 8,000,000 New Shares at a price of $\in$ 0.50 each (the "Issue Price"), totalling an amount of $\in$ 4.0 million. The New Shares were offered and sold outside the United States in reliance on Regulation S.
Pari Passu	The rights of holders of the New Shares and our existing Shares rank <i>pari passu</i> with each other.
Shares Outstanding	Immediately prior to the Private Placement, we had 36,778,974 Shares outstanding (see "Description of Share Capital and Corporate Governance – Share Capital – Authorized and Issued Share Capital").
	Immediately following the Private Placement, we have 44,778,974 Shares outstanding.
Share Ownership	A significant part of our outstanding share capital is owned by Life Sciences Partners, Signet Healthcare Partners, Sodoro B.V., Glaxo Smithkline plc, Innoven Partenaires S.A., and A. Strating (the "Major Shareholders").
Dilution	As a result of the issue of the 8,000,000 New Shares pursuant to the Private Placement, the number of outstanding shares increased from 36,778,974 to 44,7778,974. Any shareholder of OctoPlus who has not participated in the Private Placement suffered a dilution of 21.8% as a result of the Private Placement.
Listing Date	On or about 26 January 2012.
Use of Proceeds	We intend to use the net proceeds we have received from issuance of the New Shares, after deduction of the expenses related to the Private Placement for working capital purposes and, insofar possible within the available cash balance, to retain upside by co-investing along with clients in selected high-value projects in the area of specialty generics (see "Use of Proceeds").
Dividends	We do not anticipate paying any dividends for the foreseeable future (see "Dividend Policy").
Voting Rights	Holders of the Shares will be entitled to one vote per Share at General Meetings of Shareholders (see "Description of Share Capital and Corporate Governance – General Meetings of Shareholders and Voting Rights").
Share Trading Information	ISIN Code: NL0000345718
	Common Code: 026668441

Euronext Amsterdam Symbol: OCTO

### **Summary Consolidated Financial Information**

The summary consolidated financial information set forth below should be read in conjunction with "Operating and Financial Review", "Selected Financial Information" and our consolidated financial statements and their related notes that are incorporated by reference in this Prospectus. The year-end consolidated financial information has been extracted from our consolidated financial statements for the years ended 31 December 2008, 2009 and 2010 that have been audited by Deloitte Accountants B.V., independent auditors. The six months consolidated financial information has been extracted from our unaudited condensed consolidated interim financial statements as of and for the six months periods ended 30 June 2010 and 30 June 2011. Our results for the six months period ended 30 June 2011 are not necessarily indicative of our results for the full year.

Our consolidated financial statements, from which the summary consolidated financial information set forth below has been derived, were prepared in accordance with IFRS adopted by the European Union (EU). The summary consolidated financial information set forth below may not contain all of the information that is important to you.

	Year ended 31 December			Six months ended 30 June	
(in thousands)	2008*,**	2009**	2010	2010	2011
				UNAUDITED	UNAUDITED
Service revenues	€ 8,708	€ 18,636	€7,978	€3,808	€3,931
License and other revenues	8,160	370	86	22	9
Income from subsidies	55	40	265	192	12
Total (consolidated) revenues	16,923	19,046	8,329	4,022	3,952
Cost of materials and work					
contracted out	4,446	3,502	1,090	641	466
Wages and salaries	9,809	10,184	6,394	3,469	3,447
Depreciation and amortization	1,607	2,770	2,774	1,246	1,087
Other costs	5,384	4,390	3,292	1,712	1,381
Total operating costs	21,246	20,846	13,550	7,068	6,381
Operating loss	(4,323)	(1,800)	(5,221)	(3,046)	(2,429)
Interest	(1,886)	(1,157)	(981)	(494)	(456)
Result for the period	€ (6,209)	€ (2,957)	€(6,202)	€(3,540)	€(2,885)

#### Consolidated Statement of Comprehensive Income Information

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of comprehensive income, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

\*\* In the year 2010, the Company changed its presentation of operating costs. This change in presentation was applied retrospectively resulting in some reclassifications in the 2008 and 2009 statement of comprehensive income, as detailed on page 49 and 50 of the 2010 Annual Report, which is incorporated by reference.

### Segmented Income Statement Information

From October 2009 onwards, OctoPlus operates in one reportable segment and does not prepare and report financial statements per segment.

### **Consolidated Statement of Financial Position Information**

	Year ended 31 December			Six months ended 30 June	
(€ in thousands)	2008*	2009	2010	2010	2011
				UNAUDITED	UNAUDITED
Intangible fixed assets	€3,044	€2,374	€2,059	€2,216	€1,920
Other fixed assets	20,955	19,752	17,347	18,729	16,462
Current assets	6,139	7,615	5,941	4,822	4,677
Total assets	30,138	29,741	25,347	25,767	23,059
Equity	575	11,343	8,935	7,970	6,206
Non-current liabilities	11,191	10,316	9,296	9,815	8,758
Current liabilities	18,372	8,082	7,116	7,982	8,095
Total equity and liabilities	€30,138	€29,741	€25,347	€25,767	€23,059

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of financial position, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

### Consolidated Cash Flow Statement Information

	Year ended 31 December			Six months ended 30 June	
(€ in thousands)	2008*	2009	2010	2010	2011
				UNAUDITED	UNAUDITED
Result for the period	€ (6,209)	€ (2,957)	€(6,202)	€(3,540)	€(2,885)
Depreciation and amortization	1,607	2,770	2,774	1,246	1,087
Other non-cash transactions	565	(2,608)	468	165	850
Cash flow from operating activities	(4,037)	(2,795)	(2,960)	(2,129)	(948)
Cash flow from investing activities	(6,698)	(1,456)	(58)	(51)	(33)
Cash flow from financing activities	7,338	8,446	2,412	(694)	(527)
Net cash flow	(3,397)	4,195	(606)	(2,874)	(1,508)
Cash at beginning of period	2,515	(882)	3,313	3,313	2,707
Cash at end of period	(882)	3,313	2,707	439	1,199
Net cash flow	€(3,397)	€4,195	€(606)	€(2,874)	€(1,508)

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of financial position, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

### **Risk Factors**

Investing in the Shares involves a high degree of risk. You should carefully consider the risks described below and all of the other information set forth in this Prospectus before deciding to invest in any of our Shares. If any of the events or developments described below occurs, our business, financial condition or results of operations could be negatively affected. In that case, the trading price of the Shares could decline, and you could lose all or part of your investment in the Shares.

Although we believe that the risks and uncertainties described below are our most material risks and uncertainties, they are not the only ones we face. Additional risks and uncertainties not presently known to us or that we currently deem immaterial may also have a material adverse effect on our business, results of operations or financial condition and could negatively affect the price of the Shares.

### **Risks Related to Our Business and Strategy**

We need to enter into agreements with clients to develop product candidates based on our drug delivery technologies to further develop our business and if we fail to enter into such agreements, or if we fail to perform the obligations under these agreements, we will not generate revenues from our drug delivery technology platform and we may not be successful.

An important part of our business is to establish and maintain commercial relationships with pharmaceutical and biotechnology companies for the development and commercialization of their product candidates based on our drug delivery technologies, in particular PolyActive. In February 2005, we entered into a collaboration agreement with Biolex Therapeutics, Inc. ("Biolex") to develop and commercialize Locteron. Locteron combines our PolyActive drug delivery technology with Biolex' proprietary BLX-883 interferon alpha. In October 2008, we entered into a product rights acquisition agreement with Biolex obtained all commercialization rights of Locteron. We have successfully completed our Phase II trial for OP-145 CSOM and are looking for a commercialization partner. From 2008 onwards, we also entered into several agreements with pharmaceutical and biotechnological companies to evaluate our drug delivery platform with their compound in order to assess the feasibility for controlled release formulation of their compound. There is currently one other product based on our technologies in full development, which is a product funded by ESBATech, an Alcon/Novartis company.

The process of establishing these relationships is difficult, time-consuming and involves significant uncertainty. Moreover, even if we establish these relationships, it may be difficult to maintain or perform under such agreements, as our clients may seek to renegotiate or terminate their relationships with us due to unsatisfactory clinical results, a change in business strategy, a change of control or other reasons. If we or our clients fail to fulfil any responsibilities in a timely manner, or at all, contractual disputes may arise and the research, clinical development or commercialization efforts related to that agreement could be delayed or terminated.

Our ability to predict the success of our agreements is limited due to (amongst others) the complexity and uncertainty of these projects. These agreements may provide for milestone payments upon the achievement of specified clinical and regulatory milestones. They may also provide royalty-based revenue if product candidates of our clients are successfully commercialized. We may not be able to achieve any of the milestones provided for in our agreements or derive any royalty revenue with respect to these agreements.

A substantial part of our future commercial potential depends on Locteron, a product candidate which is still under development. If our licensee, Biolex, is unable to bring Locteron to market successfully, or if significant delays are experienced in doing so, our ability to generate revenue will be harmed. Locteron has completed two Phase IIa clinical trials and one Phase IIb clinical trial. Our ability to generate future revenues from Locteron will depend on the successful development and commercialization by Biolex of Locteron. Locteron could be unsuccessful if:

- it does not demonstrate acceptable safety and efficacy in future clinical trials or otherwise does not meet applicable regulatory standards for approval;
- it does not offer therapeutic or other improvements over existing or future drugs used to treat the same conditions;
- it is not capable of being produced in commercial quantities at acceptable costs;
- it is not accepted in the medical community or by third-party payors; or
- Biolex's production method for interferon alpha is not approved by the regulatory authorities;
- we are unable to manufacture such product candidate in sufficient quantities to conduct clinical testing, because our facilities and operations fail to meet the complex requirements of the European Medicines Evaluation Agency (the "EMEA"), the US Food and Drug Administration (the "FDA") and other regulatory agencies;
- Biolex fails to devote sufficient resources towards the development and approval of Locteron, either directly or indirectly through a commercial partner; or
- Biolex terminates our collaboration agreement at will upon due notice, subject to the survival of certain obligations, or upon our material breach of the product rights acquisition agreement.

Due to the significant costs associated with a clinical Phase III study with Locteron, it has always been clear that Biolex would require a commercial partner before the start of such a study. Biolex has been in negotiations with potential commercial partners for Locteron for a period of time. Finding a commercial partner is a time-consuming process because (i) the size of a transaction will be very significant, (ii) it requires in-depth due diligence and extensive discussions about the product and the futures of and developments in the market. These discussions have taken more time than initially anticipated by Biolex. This delay might well be influenced by the significant developments in the hepatitis C market as discussed in "Risk Factors - Risks Related to Our Business and Strategy – Our business faces rapid technology change". We currently expect that Locteron will not be commercially available until 2016 at the earliest, if at all. If Biolex and we are unable to make Locteron commercially available, we will not generate revenues and we may not be successful. Notwithstanding positive clinical results so far, the results of the clinical trials completed to date do not provide assurance that acceptable efficacy or safety will be adequately demonstrated in Phase III clinical trials.

## Our clients may not be able to commence or complete clinical trials or may not gain market acceptance for their product candidates. This could result in the delay or termination of the project and we may not be able to generate further revenues from that project.

To obtain the requisite regulatory approvals to market and sell any product candidates by our clients, it must be demonstrated through extensive pre-clinical and clinical trials, that such product candidates are safe and effective in humans. Pre-clinical and clinical trials are expensive, can take many years and have an uncertain outcome. A failure of one or more clinical trials could occur at any stage of testing.

Positive or timely results from pre-clinical and early clinical trials do not ensure positive or timely results in late stage clinical trials or product approval by the EMEA, the FDA or any other regulatory authority. Product candidates that show positive pre-clinical or early clinical results often fail in later stage clinical trials.

Any delay in commencing or completing clinical trials conducted by our clients could delay further development of the product candidates involved and may harm our business as such delay could have a negative impact on our revenues and cash position in the (near) future.

#### Our business faces intense competition.

We face competition in the area of the pharmaceutical contract formulation and manufacturing services we offer. There are many organizations in our sector which are actively engaged in offering similar services as we do to the same markets. These competitors may succeed in acquiring new projects more rapidly than we do and this will impact our ability to generate service revenues.

In the area of drug delivery technologies, there is a number of existing and new technologies on the market that can offer similar benefits as our drug delivery technologies. Our clients may elect to use an alternative drug delivery technology for the controlled release of their compound instead of one of our drug delivery technologies, which will impact our ability to generate service revenues and revenues in the form of milestones and royalties.

The pharmaceutical and biotechnology industries are highly competitive. Any product candidates that our clients successfully develop will compete with existing and future therapies. There are many organizations, including pharmaceutical companies, biotechnology companies, academic laboratories, research institutions, governmental agencies and public and private universities, which are actively engaged in developing products that target the same markets as product candidates, which we develop for our clients.

These competitors may have substantially greater financial, technological, manufacturing, marketing, managerial and research and development resources and experience than our clients may have. These organizations may also have much more experience than our clients have in pre-clinical and clinical trials of new drugs and in obtaining regulatory approvals. Accordingly, these competitors may succeed in developing competing technologies and products more rapidly than we and our clients do.

The product candidates of our clients if successfully developed and approved for commercial sale, will compete with a number of drugs and therapies currently manufactured and marketed by major pharmaceutical and biotechnology companies. Such product candidates may also compete with new products currently under development by others. These new products may turn out to be safer or may work better, or be as effective but cheaper, than the products of our clients. If our competitors develop and market product candidates that are safer, more effective or cheaper, or develop, obtain regulatory approval and market such product candidates earlier than we and our clients do, our commercial opportunity will be reduced or eliminated.

### Our business faces rapid technological change.

Within the pharmaceutical and biotechnology industries, major technological changes can happen quickly. The rapid technological change, or the development by competitors of technologically improved or different drug delivery systems or products, could render our platform technologies or the product candidates of our clients obsolete or non-competitive. For example, the current standard of care for HCV infection consists of a combination therapy using interferon alpha and ribavirin. Locteron is designed to be an improved interferon alpha component of this therapy. Recently, two direct acting anti-virals (Incivek from Vertex Pharmaceuticals and Victrelis from Merck) have been approved by the FDA.

Incivek reduces the necessary HCV treatment period from 48 to 24 weeks for many patients, and shows a higher cure rate than traditional therapies, both in treatment naïve as well as treatment experienced patients. Incivek has rapidly become the new standard of care regimen, combined with ribavirin and interferon alpha. Even though the therapy duration is expected to be significantly reduced in the future, the interferon alpha market is expected to expand as more patients are eligible for treatment and better therapy will attract patients that so far selected not to be treated, with interferon alpha remaining an important element in such future hepatitis C treatment.

In addition, several companies are exploring combinations of direct antiviral compounds that do not require interferon injections and can be given through pills only, so-called all-oral therapy. Clinical studies are ongoing in this area, but no products have been approved and no final proof has been established to date to show that an all-oral therapy would be technically or commercially feasible for all HCV genotypes, including the most difficult to treat genotype la.

In the event that a new standard of care emerges that does not use interferon alpha, Locteron could become obsolete for (part of) the market. In case Locteron becomes obsolete for part of the market, the likelihood of us generating income from Locteron decreases and the magnitude of this possible income is also expected to decrease. In case Locteron becomes obsolete for the entire market, it is unlikely that we would receive any income from the product in the future.

### Our clients have significant product liability exposure, which may also harm our business and our reputation.

Our clients face an inherent risk of product liability lawsuits related to the testing of product candidates, and will face an even greater risk if such product candidates are introduced commercially. An individual may bring a liability claim against a client if one of its product candidates causes, or merely appears to have caused, an injury. If our clients cannot successfully defend themselves against a product liability claim, they may incur substantial liabilities, which may also harm our business and reputation.

Our clients are highly dependent upon market perceptions, their brands and the safety and quality of their products. Our clients could be adversely affected if they, or their brands, are subject to negative publicity and this may also harm our reputation and our brands. Our clients and we could also be adversely affected if any of their products or any similar products distributed by other companies prove to be, or are asserted to be, harmful to consumers. Also, because of their and our dependence upon market perceptions, any adverse publicity associated with illness or other adverse effects resulting from consumers' use or misuse of such products or any similar products distributed by other companies could have a material adverse impact on the results of operations of our clients and on our results of operations.

We are also exposed to similar liability risks due to the provision of pharmaceutical development and manufacturing services by our contract formulation and manufacturing business.

Some of our supply agreements with clients contain liability and/or indemnification provisions under which we may claim damages from our counterparties and under which our counterparties may claim damages from us, including damages caused by product defects. In the event we need to claim damages from a counterparty, we may not receive payments covering in full our damages, either because the applicable provision limits the payment to a certain amount, is unenforceable for any reason or because the counterparty is unable to pay (due to insolvency or otherwise). Although in many cases we try to limit our liability, such limitations may not be effective in the event that we need to pay damages and we nevertheless could become liable to make substantial payments.

We have liability insurance, which we currently believe is adequate to cover liabilities we may incur. However, our current or future insurance coverage may prove insufficient to cover any liability claims brought against us. Because of the increasing costs of insurance coverage, we may not be able to maintain insurance coverage at a reasonable cost or obtain insurance coverage that will be adequate to satisfy any liability that may arise.

### We do not have large-scale manufacturing experience.

Although we have manufactured certain product candidates on a small scale for use in clinical trials, we cannot assure you that we can successfully manufacture the products of our clients under current Good Manufacturing Practices ("cGMP") and applicable laws and regulations in sufficient quantities for clinical trials, or in a timely or economical manner. We have not yet produced any biopharmaceutical products on a large scale or in commercial quantities and do not have the facilities and staff required for commercial products that require large-scale manufacturing. To obtain such resources, we may

seek to enter into collaborative arrangements with other parties. There can be no assurance that we will be able to obtain such capabilities through arrangements with others on acceptable terms, if at all.

### We are exposed to complex production issues.

The formulation and manufacturing services that we offer can be highly complex. From time to time, issues may arise in the formulation laboratory or manufacturing facility, in both cases for a variety of reasons including equipment malfunction, failure to follow specific protocols and procedures, issues with raw materials, environmental factors and systematic failure of our information technology and other internal infrastructure systems, including corporate firewalls, servers, leased lines and connection to the Internet. Such issues could affect the formulation success, the production of a particular batch or series of batches. This could, among other things, lead to increased costs, lost revenue, damage to client relations, reimbursement to clients for lost active pharmaceutical ingredients, time and expense spent investigating the cause and, depending on the cause, similar losses with respect to other batches or products.

### Expenses related to our development and manufacturing capacity are largely fixed.

The amount that the pharmaceutical and biotechnology industry spends on formulation development and manufacturing for clinical trials and commercial use and in particular how much it spends on outsourcing such activities may have a large impact on our revenues and profitability. For example consolidation in the industries in which our clients operate may have an impact on such spending as clients integrate acquired operations, including research and development departments and manufacturing. Also, additional competition may emerge and may, among other things, result in a decrease of the fees paid for services or a reduction in work for us. As a result, we may periodically experience overcapacity in terms of development resources and manufacturing resources which could affect our financial position as the costs related to these resources are largely fixed in the medium-term.

## We are vulnerable to natural and other types of disaster, which could cause damage to our facilities and equipment and require us to cease or curtail operations.

All of our facilities are located within a single facility in Leiden, the Netherlands. We are vulnerable to damages from natural and other types of disasters, in particular fires and floods. Depending on the severity and duration of such a disaster, our ability to operate our business could be seriously impaired when either any of the identified types of disaster occurs. In addition, the nature of our activities and of much of our equipment could make it difficult for us to recover from these types of disaster. We have a property damage insurance as well as back-up scenarios in place but these might still not cover all our losses resulting from these disasters. We do not plan to purchase additional insurance to cover such potential losses due to the cost of obtaining such coverage. Any significant losses that are not recoverable under our insurance policies could seriously impair our business and financial condition.

### **Risks Related to Our Intellectual Property**

## Our success is dependent on intellectual property rights held by us, and third parties, and the value of such rights is complex and uncertain.

Our success depends, in part, on our ability to obtain, maintain and enforce our intellectual property rights relating to our technologies and product candidates that are developed based on our drug delivery systems and to preserve our trade secrets and other proprietary information. Our success also depends, in part, on the ability of our licensors to obtain, maintain and enforce their intellectual property rights to the extent that we rely on those rights.

Patent positions, in particular those of pharmaceutical and biotechnology companies, including ours, involve complex legal and factual questions. Validity and enforceability of patents cannot be guaranteed. Patents may be challenged, invalidated, deemed unenforceable, or circumvented. The validity, enforceability and commercial value of patents are therefore highly uncertain.

We intend to apply for patents covering our technologies and when possible the combination of our technology and product candidates, as we deem appropriate. We may fail to apply for patents on important technologies or products in a timely fashion, or at all, and in any event, the patent applications filed by us may not result in issued patents, or their validity may be challenged. Any patents obtained by us may not be sufficiently broad to prevent others from using our technologies or from developing competing products. Furthermore, others may independently develop similar or alternative technologies or design around our patented technologies. In addition, if challenged, our patents may be declared invalid. Even if valid, our patents may fail to provide us with any competitive advantages.

Any adverse outcome we may experience while attempting to obtain, maintain or enforce our intellectual property rights may have a material adverse effect on our business and financial condition.

## Litigation or third party claims of intellectual property infringement could require substantial time and money to resolve. Unfavorable outcomes in these proceedings could limit the value of our intellectual property rights and our activities.

We may need to resort to litigation to enforce or defend our intellectual property rights, including any patents issued to us. If a competitor or client files a patent application having a scope covering our technology or any of the product candidates that are developed based on our drug delivery systems in order to protect our rights, we may have to participate in expensive and time-consuming opposition proceedings, re-examination proceedings, or nullity proceedings, whichever is applicable, before the European Patent Office, the US Patent and Trademark Office ("USPTO") or patent authorities in other jurisdictions. We cannot guarantee that there will be no claims from third parties alleging that such product candidates infringe their intellectual property rights. Third parties may assert that we or our clients are employing their proprietary technologies without authorization and they may resort to litigation to attempt to enforce their rights. Third parties may own or obtain patents relevant to our technology or the technology of our collaborators or the product candidates that are developed based on our drug delivery systems and claim that the use of this technology or any of these product candidates infringes their patents.

We may not be able to enter into collaboration agreements for the development or commercialization of our current product candidates due to third party patent protection. Our business may be harmed if we cannot negotiate a necessary or desirable license, can obtain such a license only on unattractive or unacceptable terms, or if we are unable to redesign our product candidates or processes to avoid actual or potential infringement of patents or other intellectual property. Our efforts to obtain, protect and defend our patents and other intellectual property rights, whether successfully or not, may be expensive and may require us to incur substantial costs, including the diversion of management and technical personnel. An unfavorable ruling in patent or intellectual property litigation could subject us to significant liabilities to third parties, require our clients to cease developing, manufacturing or selling the affected products or using the affected processes, require us to license the disputed rights from third parties, or result in awards of substantial damages against us. During the course of any patent litigation, there may be public announcements of the results of hearings, motions and other interim proceedings or developments in the litigation. If securities analysts or investors regard these announcements as negative, the market price of our Shares may decline. General proclamations or statements by key public figures may also have a negative impact on the perceived value of our intellectual property.

### Confidentiality agreements with employees and others may not adequately prevent disclosure of trade secrets and other proprietary information.

We rely substantially upon proprietary materials, information, trade secrets and know-how to conduct our research and development activities, and to attract and retain collaborators, licensees and clients. We take steps to protect our proprietary rights and information, including the use of confidentiality and other agreements with our employees and consultants and with academic and commercial relationships. However, these steps may be inadequate, agreements may be violated, or there may be no adequate remedy available for a violation of an agreement. We cannot assure you that our proprietary information will not be disclosed or that we can meaningfully protect our trade secrets. Our competitors may independently develop substantially equivalent proprietary information or may otherwise gain access to our trade secrets, which could adversely affect our ability to compete in the market.

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

## We rely significantly on the skills and experience of qualified management and scientific personnel and as a consequence on our ability to attract, retain and motivate qualified management and scientific personnel.

Our success depends on the skills and experience of qualified management and scientific personnel. In particular, we are dependent on Mr. Egberts, our Chief Executive Officer, and the other members of our Executive Board. We also rely significantly on the know-how, expertise and experience of our scientific personnel. If we are not able to retain these persons, we may not be able to successfully conduct our business. We do not maintain "key person" insurance in relation to any of our employees.

Recruiting and retaining qualified personnel is critical to our success. We may not be able to attract and retain qualified personnel on acceptable terms given the competition among pharmaceutical and biotechnology companies for similar personnel. We also experience competition for the hiring of scientific personnel from universities and research institutions. In addition, we rely on consultants and advisors, including scientific and clinical advisors, to assist us in formulating our strategy. Our consultants and advisors may be employed by third parties or may have commitments under consulting or advisory contracts with third parties that may limit their availability to us.

#### We may encounter difficulties in managing our growth.

We may experience further growth in the number of our employees and the scope of our operations over the next several years. To manage future growth, we must continue to implement and improve our managerial, operational and financial systems, and continue to recruit and train additional qualified personnel. Due to our limited resources, we may not be able to effectively manage the expansion of our operations or to recruit and train additional qualified personnel. The expansion of our operations may lead to significant costs and may divert our management and business development resources. Any inability to manage growth could delay the execution of our business plans, which would adversely affect our results.

### **Risks Related to Regulatory Approval and Other Government Regulations**

## The regulatory approval process is costly and lengthy and our clients may not be able to successfully obtain all required regulatory approvals.

The pre-clinical development, clinical trials, manufacturing, marketing and labelling of pharmaceuticals and medical devices are all subject to extensive regulation by numerous governmental authorities and agencies in the European Union, the United States and other jurisdictions. Our clients must obtain regulatory approval for product candidates before marketing or selling any of them. The approval process is typically lengthy and expensive, and approval is never certain. It is not possible to predict how long the approval processes of the EMEA, the FDA or any other applicable regulatory agency will take or whether any such approvals ultimately will be granted. The EMEA, the FDA and other regulatory agencies have substantial discretion in the drug and medical device approval process, and positive results in pre-clinical or early clinical trials provide no assurance of success in later phases of the approval process. Generally, pre-clinical and clinical trials of products and medical devices can take many years and require the expenditure of substantial resources, and the data obtained from these trials can be susceptible to varying interpretations that could delay, limit or prevent regulatory approval. The risks associated with the approval process include delays or rejections in the regulatory approval process based on the failure of clinical or other data to meet expectations, or the failure of the product or medical device to meet a regulatory agency's requirements for safety and efficacy. In addition, regulatory approval, if obtained, may significantly limit the indicated uses for which a product may be marketed.

Additional clinical trials may be required if clinical trial results are negative or inconclusive, which will require our clients to incur additional costs and will cause significant delays. If the necessary regulatory

approvals are not received, we will not be able to generate revenues and may not become profitable. Our clients may encounter significant delays in the regulatory process that could result in excessive costs that may prevent our clients from continuing to develop such product candidates. In addition, the failure to comply with applicable regulatory requirements may result in criminal prosecution, civil penalties, product recalls, withdrawal of product approval, mandatory restrictions and other actions that could impair our ability to conduct our business.

# Product candidates which are developed by our clients will remain subject to ongoing regulatory review even if they receive marketing approval. If we and our clients fail to comply with continuing regulations, these approvals could be withdrawn and the sale of the products could be suspended.

Even if regulatory approval to market a particular product candidate is received, the approval could be conditioned on conducting additional costly post-approval studies or could limit the indicated uses included in the labelling of products. Moreover, the product may later cause adverse effects that limit or prevent its widespread use, force our clients to withdraw it from the market or impede or delay the ability to obtain regulatory approvals in additional countries. In addition, the manufacturer of the product and its facilities will continue to be subject to regulatory review and periodic inspections to ensure adherence to applicable regulations. After receiving marketing approval, the manufacturing, labelling, packaging, adverse event reporting, storage, advertising, promotion and record keeping related to the product will remain subject to extensive regulatory requirements. If we fail to comply with applicable regulatory requirements, or previously unknown problems with the products, manufacturers or manufacturing processes are discovered, we could be subject to administrative or judicially imposed sanctions, including restrictions on our manufacturing processes, civil or criminal penalties or fines, injunctions, voluntary or mandatory product recalls, suspension or withdrawal of regulatory approvals, and/or total or partial suspension of production. Any such sanction could severely harm our business and financial condition.

Our contract formulation and manufacturing business involves drug development and manufacturing on behalf of third parties. Both of these activities, comprising decision-making processes, manufacturing processes, procurement, manufacture, storage, shipping and record keeping, must comply with the FDA's cGMP regulations, among other requirements. FDA oversight, including facility inspections, is ongoing, and a failure to comply with applicable requirements can result in adverse publicity, warning letters, civil and criminal liability, and restrictions on and/or prohibitions against the performance of these activities, any of which may result in a loss of business.

### **Risks Related to Our Financial Condition**

### The Company has not secured sufficient working capital for its present requirements, being the next 12 months from the date of this Prospectus.

If no additional cash is received from OctoPlus' Fee-For-Services Business, its business based upon its proprietary drug delivery technologies and OP-145 or our mitigation action plan, as defined and described in "Operating and Financial Review– Working Capital Statement - OctoPlus' mitigating action plan", OctoPlus', the available balance of net cash (cash and cash equivalents minus bank overdrafts) at the date of this Prospectus is expected to deplete in the course of the second quarter of 2012 and we would require an additional  $\in 4.5 - 5.0$  million cash until 12 months after the date of this Prospectus. Our mitigating action plan is to raise new funds through new share issuances. These additional funds may not be available when we need them on terms that are acceptable to us, or at all. In addition, market developments may impact our ability to raise additional funds. If adequate funds are not available on a timely basis, we may be required to delay, scale back or eliminate certain of our activities, which would have a material adverse affect on our financial condition or cause us to discontinue our operations. In addition, if OctoPlus' equity becomes negative, OctoPlus shall have to announce this to the market together with the planned measures to improve its solvability and liquidity position. For details, reference is made to "Operating and Financial Review – Cash Flows – Working Capital Statement".

### We have incurred losses since 2002

We are not profitable and have incurred losses in each year since 2002. Our net loss for the years ended 31 December 2010, 2009 and 2008 was  $\in$  6.2 million,  $\in$  3.0 million, and  $\in$  6.2 million, respectively, and  $\in$  2.9 million for the first six months of 2011. As of 30 June 2011, we had an accumulated deficit of  $\in$  52.0 million. Our losses, among other things, have caused our shareholders' equity and cash position to decrease. To date, we have derived a substantial portion of our revenues from our contract formulation and manufacturing business. We have also received government subsidies, a significant up-front fee from Biolex for the sale of the commercialization rights to Locteron and nominal licensing revenues through the out-licensing of certain technologies, i.e. the granting of rights to third parties to use our technologies in accordance with the scope of the license agreements entered into with such parties.

In October 2008, we received an up-front fee from Biolex for the sale of the commercialization rights to Locteron. We do not expect that we will generate royalty income from the sale of products for the foreseeable future. If Locteron or the other current product candidates that we intend to develop with current and future clients, fail in clinical trials or do not gain regulatory approval, or if such product candidates do not achieve market acceptance, we may never achieve profitability. Even if we achieve profitability in the future, we may not be able to sustain profitability in subsequent periods.

### We have significant payment obligations

Our total payment obligations under contract as of 30 June 2011, as outlined in "Operating and Financial Review – Contractual Obligations", amount to  $\in$  19.5 million and relate to the period up to the year 2028. These payment obligations mostly related to the lease of our headquarters for a period of twenty years and the lease of a significant part of the equipment used in our new manufacturing facility for a period of five years. These payment obligations could harm the Company in its freedom to operate.

## We may need additional funding in the future, which may not be available to us on acceptable terms, or at all, and which could force us to delay, reduce or eliminate product development programs or commercialization efforts or cause us to discontinue our operations.

We may need to raise additional capital to pursue our business strategy. Our future funding requirements will depend on many factors, including:

- our ability to continue to generate revenues from our contract formulation and manufacturing business;
- terms and timing of any collaborative, licensing and other arrangements that we may establish;
- timing, receipt and amount of royalties, if any, from the potential products, or any up-front or milestone payments during their development phase;
- the cost of preparing, filing, prosecuting, defending and enforcing any patent claims and other intellectual property rights; and
- the extent to which we acquire or invest in businesses, products or technologies, although we currently have no commitments or agreements relating to any of these types of transactions.

Additional funds may not be available when we need them on terms that are acceptable to us, or at all. In addition, market developments may impact our ability to raise additional funds (see also "Operating and Financial Review – Working Capital Statement – OctoPlus' mitigating action plan"). If adequate funds are not available on a timely basis, we may be required to delay, scale back or eliminate certain of our activities, which would have a material adverse affect on our financial condition or cause us to discontinue our operations.

Raising additional capital by issuing securities or through collaboration and licensing arrangements may cause dilution to existing shareholders, restrict our operations or require us to relinquish proprietary rights.

We may seek the additional capital necessary to fund our operations through public or private equity offerings, debt financings and collaboration and licensing arrangements. In this respect we note that raising new funds is part of our mitigating action plan (see "Operating and Financial Review – Working Capital Statement – OctoPlus' mitigating action plan"). To the extent that we raise additional capital through the issuance of Shares or convertible debt, your ownership interest will be diluted, and the terms of such securities may include liquidation or other preferences that adversely affect your rights as a shareholder. Debt financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions such as incurring additional debt, making capital expenditures or declaring dividends. If we raise additional funds through collaboration and licensing arrangements with third parties, we may have to relinquish valuable rights to our technologies, or grant licenses on terms that are not favorable to us.

## We face the potential loss of significant revenue from the loss, modification or delay of large contracts or the inability to secure new contracts.

To date, our principal sources of revenue have been our contract formulation and manufacturing business and, to a lesser extent, government subsidies and license revenues. Our customer base presently comprises approximately 40 clients who award us with work on a contract-by-contract basis. In 2010, our five largest clients accounted for 45% of our total revenues. Our clients may resolve, on relatively short notice, to suspend or terminate the development activities that we conduct on their behalf. We have in the past experienced such project suspensions or terminations with significant clients. We expect that a small number of clients will continue to represent a significant part of our contract formulation and manufacturing revenues. In addition, our clients and target clients, as they try to reduce costs as a result of budgetary limits or changing priorities, may proceed without or with decreased assistance of biopharmaceutical services companies like us. Some of our clients and target clients are loss making and as such will be dependent on external financing for their product development programs. If they fail to secure such external financing, they may not have the financial means to continue working with us. The loss, modification or delay of a large contract or of multiple contracts, or the inability to secure new contracts, could have a material adverse effect on our operating results.

### **Risks Related to the Shares**

## The price of our Shares may be volatile and you may not be able to sell our Shares at or above the price you pay for them.

We cannot predict the extent to which an active market for our Shares will develop or be sustained, or how the development of such a market might affect the market price for our Shares. An illiquid market for our Shares may result in lower trading prices and increased volatility, which could adversely affect the value of your investment.

The Issue Price was agreed between us and the Investors based on a number of factors, including market conditions in effect shortly prior to the Private Placement, which may not be indicative of future performance. The market price for our Shares may fall below the Issue Price. The market price of our Shares could fluctuate substantially due to factors described above and a number of other factors, including, but not limited to:

- · changes in financial estimates or recommendations by securities analysts;
- sales of large blocks of our Shares;
- sales of our Shares by our executive officers, directors and significant shareholders;
- restatements of our financial results and/or material weaknesses in our internal controls;
- publication of research reports about us or the biopharmaceutical industry by securities or industry analysts;

- failure to meet or exceed securities analysts' expectations relating to our financial results;
- speculation in the press or investment community generally;
- general economic conditions; and
- war, acts of terrorism and other man-made or natural disasters.

The stock markets in general and the markets for biopharmaceutical stocks in particular, have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. These broad market fluctuations may adversely affect the trading price of our Shares.

### The ownership of our Shares will continue to be highly concentrated and your interests may conflict with the interests of our existing shareholders.

Life Sciences Partners, Signet Healthcare Partners, Sodoro B.V., Glaxo Smithkline plc, Innoven Partenaires S.A., and A. Strating (the "Major Shareholders") own a significant part of our Shares (see chapter "Major Shareholders"). Accordingly, these shareholders may have a significant influence over the outcome of corporate actions requiring shareholder approval, including the election of members of our Supervisory Board and Executive Board, any merger, consolidation or sale of all or substantially all of our assets or any other significant corporate transaction. These shareholders could delay or prevent a change of control of our Company, even if such a change of control would benefit our other shareholders. The significant concentration of share ownership may adversely affect the trading price of our Shares due to investors' perception that conflicts of interest may exist or arise.

## Our Executive Board has broad discretion over the use of the net proceeds received by us from the Private Placement and may not apply the net proceeds effectively or in ways with which you agree.

Our Executive Board has broad discretion over the use of net proceeds from the sale of the New Shares. We intend to use the net proceeds from the Private Placement primarily for working capital purposes and to retain upside by co-investing along with clients in selected high-value projects in the area of specialty generics (see "Use of Proceeds"). You will not have an opportunity, as part of your investment decision, to assess whether the net proceeds received by us are being used appropriately. We cannot assure you that our Executive Board will apply the net proceeds effectively or that the net proceeds will be invested to yield a favorable return.

## Future sales, or the possibility of future sales, of a substantial amount of our Shares may depress the price of our Shares.

Future sales of our Shares, or the perception that such sales will occur, could cause a decline in the market price of our Shares. A significant part of our Shares is currently owned by the Major Shareholders (see chapter "Major Shareholders").

### US and other non-Dutch holders of our Shares may not be able to exercise pre-emption rights.

In the event of an increase in our share capital, holders of our Shares are generally entitled to certain pre-emption rights, unless these rights are excluded by a resolution of the General Meeting of Shareholders or of the Executive Board, if so designated by the General Meeting of Shareholders or pursuant to our Articles of Association.

In particular, US holders of our Shares may not be able to exercise pre-emption rights unless a registration statement under the Securities Act is declared effective with respect to the Shares issuable upon exercise of such rights or an exemption from the registration requirements is available. We intend to evaluate at the time of any rights issue the cost and potential liabilities associated with any such registration statement, as well as the indirect benefits and costs to us of enabling the exercise by US holders of their pre-emption rights for our Shares and any other factors considered appropriate at the

time. We will then make a decision as to whether to file such a registration statement, although currently we do not expect that any registration statement would be filed. No assurance can be given that any registration statement would be filed or that any exemption from registration would be available to enable the exercise of a US holder's pre-emption rights.

### If securities or industry analysts do not publish research or reports about our business, or if they change their recommendations regarding our Shares adversely, the price and trading volume of our Shares could decline.

The trading market for our Shares will be influenced by the research and reports that industry or securities analysts publish about us or our business. We are followed by analysts. If one or more of the analysts who cover us or our industry downgrade our Shares, the market price of our Shares would likely decline. If one or more of these analysts ceases coverage of us or fails to regularly publish reports on us, we could lose visibility in the financial markets, which could cause the market price of our Shares or trading volume to decline.

### We do not intend to pay dividends for the foreseeable future.

The Company has never paid any dividends to its Shareholders. We also do not intend to pay any dividends for the foreseeable future. Payment of future dividends to shareholders will effectively be at the discretion of the Executive Board, subject to the approval of the Supervisory Board after taking into account various factors including our business prospects, cash requirements, financial performance and new product development. In addition, payment of future dividends may be made only if our shareholders' equity exceeds the sum of our called up and paid-in share capital plus the reserves required to be maintained by law and by our Articles of Association. Accordingly, investors cannot rely on dividend income from our Shares and any returns on an investment in our Shares will likely depend entirely upon any future appreciation in the price of our Shares.

### Important Information

No person is or has been authorized to give any information or to make any representation in connection with the New Shares, other than as contained in this Prospectus, and, if given or made, any other information or representation must not be relied upon as having been authorized by us. The delivery of this Prospectus at any time after the date hereof will not, under any circumstances, create any implication that there has been no change in our affairs since the date hereof or that the information set forth in this Prospectus is correct as of any time since its date.

OctoPlus N.V. accepts responsibility for the information contained in this Prospectus. Having taken all reasonable care to ensure that such is the case, OctoPlus N.V. further declares that the information contained in this Prospectus is, to the best of its knowledge, in accordance with the facts and contains no omission likely to affect its import.

### Notice to Investors

The distribution of this Prospectus may be restricted by law in certain jurisdictions. Persons in possession of this Prospectus are required to inform themselves about and to observe any such restrictions.

This Prospectus may not be used for, or in connection with, and does not constitute, any offer to sell, or a solicitation of an offer to buy, any of the New Shares or any other securities issued by the Company.

The New Shares have not been approved or disapproved by the US Securities and Exchange Commission, any State securities commission in the United States or any other US regulatory authority, nor have any of the foregoing passed upon or endorsed the merits of the Private Placement or the accuracy or adequacy of this Prospectus. Any representation to the contrary is a criminal offence in the United States.

#### **Presentation of Financial and Other Information**

Certain figures contained in this Prospectus have been subject to rounding adjustments. Accordingly, in certain instances the sum of the numbers in a column or a row in tables contained in this Prospectus may not conform exactly to the total figure given for that column or row.

All references in this Prospectus to "euros" or "€" are to the currency introduced at the start of the third stage of the Economic and Monetary Union, pursuant to the Treaty establishing the European Economic Community, as amended by the Treaty on the European Union. All references to "US dollars", "US\$" or "\$" are to the lawful currency of the United States.

Any financial information in this Prospectus that has not been extracted from our audited consolidated financial statements for the years 2008, 2009 and 2010 is unaudited.

### **Exchange Rates**

We publish our consolidated financial statements in euros. The exchange rates below are provided solely for information and convenience. No representation is made that the euro could have been, or could be, converted into US dollars at these rates.

The table below shows the high, low, average and end of period exchange rates expressed in US dollars per € 1.00 for the years given, using the noon buying rate in New York City for cable transfers in foreign currencies as certified for customs purposes by the Federal Reserve Bank of New York (the "Noon Buying Rate") for the periods indicated.

				End of
Year ended 31 December	High	Low	Average	Period
		(US Dollar	s per Euro)	
2005	1.3476	1.1667	1.2444	1.1842
2006	1.3327	1.1860	1.2563	1.3197
2007	1.4862	1.2904	1.3708	1.4603
2008	1.6010	1.2446	1.4710	1.3919
2009	1.5100	1.2547	1.3935	1.4332
2010	1.4536	1.1959	1.3261	1.3269
2011	1.4875	1.2926	1.3821	1.2973

On 20 January 2012, the Noon Buying Rate for the euro was € 1.00 = \$ 1.2948

### Market Data and Other Information from Third Parties

All references to market data, industry statistics and industry forecasts in this Prospectus consist of estimates compiled by the World Health Organization, the US Centers for Disease Control and Prevention, the Public Health Agency of Canada, the US National Institutes of Health, Datamonitor, Decision Resources, Reuters, Intercontinental Marketing Services, equity research analysts, other biopharmaceutical companies and ourselves.

In this Prospectus, we make certain statements regarding our competitive position, the expected size of relevant markets and the side effects or efficacy of current treatments for the relevant diseases. We believe these statements to be true based on market data and industry statistics which are in the public domain.

The information in this Prospectus that has been sourced from third parties has been accurately reproduced and, as far as we are aware and able to ascertain from the information published by that third party, no facts have been omitted that would render the reproduced information inaccurate or misleading.

### **Documents Incorporated by Reference**

Our Articles of Association (*statuten*) (in the Dutch and English language) and certain parts of our Annual Reports for the years ended 31 December 2008, 2009 and 2010, our Condensed Consolidated Interim Financial Statements for the first six months ended 30 June 2010 and our Condensed Consolidated Interim Financial Statements for the first six months ended 30 June 2011, listed below, are incorporated by reference into this Prospectus. Our six months consolidated financial information is unaudited. The information in these documents that is not incorporated, is either not relevant for investors or is covered elsewhere in this Prospectus. No other documents or information form part of, or are incorporated by reference into, this Prospectus. Copies of our Annual Reports for the years ended 31 December 2008, 2009 and 2010, our Condensed Consolidated Interim Financial Statements for the first six months ended 30 June 2011 and our Condensed Consolidated Interim Financial Statements for the six months ended 30 June 2011 may be obtained free of charge for the life of this Prospectus by sending a request in writing to us at our business address: Zernikedreef 12, 2333 CL Leiden, the Netherlands. All Annual Reports are also available via <u>www.octoplus.nl</u>.

Annual Report for the year ended 31 December 2008	Incorporated by reference
Consolidated balance sheet	page 28
Consolidated income statement	page 29
<ul> <li>Consolidated statement of changes in equity</li> </ul>	page 30

Consolidated cash flow statement	page 31
Notes to the consolidated financial statements	page 32-60
Independent auditors' report	page 68-69
Annual Report for the year ended 31 December 2009	Incorporated by reference
Consolidated statement of financial position	page 42
Consolidated statement of comprehensive income	page 43
Consolidated statement of changes in equity	page 44
Consolidated statement of cash flows	page 45
Notes to the consolidated financial statements	page 46-72
Independent auditors' report	page 80-81
Annual Report for the year ended 31 December 2010	Incorporated by reference
Consolidated statement of financial position	page 44
Consolidated statement of comprehensive income	page 45
Consolidated statement of changes in equity	page 46
Consolidated statement of cash flows	page 47
Notes to the consolidated financial statements	page 48-77
Independent auditors' report	page 86-87
Condensed consolidated Interim Financial Statements 30 June 2010	Incorporated by reference
Consolidated statement of financial position	page 3

Condensed consolidated statement of changes in equity page 5

• Condensed consolidated statement of comprehensive income

Condensed consolidated statement of cash flows page 6

page 4

• Notes to the condensed consolidated interim financial statements page 7-9

Condensed consolidated Interim Financial Statements 30 June 2011	Incorporated by reference
Consolidated statement of financial position	page 3
Condensed consolidated statement of comprehensive income	page 4
Condensed consolidated statement of changes in equity	page 5
Condensed consolidated statement of cash flows	page 6
Notes to the condensed consolidated interim financial statements	page 7-9

### **Forward-Looking Statements**

This Prospectus contains forward-looking statements, including statements about our beliefs and expectations. These statements are based on our current plans, estimates and projections, as well as our expectations of external conditions and events. In particular the words "expect", "anticipate", "predict", "estimate", "project", "may", "could", "should", "would", "will", "intend", "believe" and similar expressions are intended to identify forward-looking statements. Forward-looking statements involve inherent risks and uncertainties and speak only as of the date they are made. We undertake no duty to and will not necessarily update any of them in light of new information or future events, except to the extent required by applicable law. We caution investors that a number of important factors could cause actual results or outcomes to differ materially from those expressed in any forward-looking statements. These factors include, but are not limited to those discussed under "Risk Factors".

### **Dividend Policy**

We currently intend to retain future earnings, if any, to finance the growth and development of our business. As a result, we do not anticipate paying any dividends for the foreseeable future.

Our dividend policy will, however, be reviewed from time to time and payment of any future dividends will be effectively at the discretion of the Executive Board, subject to approval of the Supervisory Board, after taking into account various factors including our business prospects, cash requirements, financial performance, in accordance with our Articles of Association (see "Description of the Share Capital and Corporate Governance – Dividends and Other Distributions") and the requirements of Dutch law. Under Dutch law, payment of dividends may be made only if our shareholders' equity exceeds the sum of our called up and paid-in share capital plus the reserves required to be maintained by law and by our Articles of Association.

### **Use of Proceeds**

We have raised  $\in$  4.0 million in gross proceeds from the issue of the New Shares. The net proceeds we have received from the Private Placement are estimated to be approximately  $\in$  3.6 million, after deducting the estimated commissions and expenses of  $\in$  0.4 million payable by us. We intend to use the net proceeds we have received from the Private Placement primarily for working capital purposes and, insofar possible within the available cash balance, to retain upside by co-investing along with clients in selected high-value projects in the area of specialty generics.

This expected use of net proceeds represents our current intentions based upon our present plans and business conditions. The amounts and timing of our actual expenditures depend on numerous factors. As a result, the Executive Board will retain broad discretion over the allocation of the net proceeds from the Private Placement. Pending use of the net proceeds of the Private Placement, we intend to invest the net proceeds in cash equivalents, government bonds, high grade and corporate notes and commercial paper.

The Company is of the opinion that the net proceeds of the Private Placement do not result in sufficient cash for its present requirements, being the next 12 months from the date of this Prospectus. Details of the cash potentially needed by the Company until December 2012 and possible sources to obtain this cash are outlined in Chapter "Operating and Financial Review – Cash Flows – Working Capital Statement").

### **Capitalization and Indebtedness**

The table below sets forth our unaudited consolidated cash and cash equivalents, capitalization and indebtedness as of 31 October 2011, on an actual basis including the net proceeds of the Private Placement.

The financial information in the table below has been extracted or derived from our 31 October 2011 management accounts which have not been audited or reviewed and will not be published. You should read this table together with our consolidated financial statements incorporated by reference in this Prospectus, as well as the information under "Operating and Financial Review" appearing elsewhere in this Prospectus. The table below is prepared for illustrative purposes only and, because of its nature, may not give a true picture of our financial condition following 31 October 2011.

(€ in thousands)	31 October 2011 UNAUDITED
<i>Capitalization</i> Guaranteed current liabilities Secured current liabilities Unguaranteed / unsecured current liabilities <b>Total current liabilities</b>	- 7,655 <b>7,655</b>
Guaranteed non-current liabilities Secured -noncurrent liabilities Unguaranteed / unsecured non-current liabilities Total non-current liabilities	- 8,436 <b>8,436</b>
Share capital – ordinary shares Share premium Retained earnings (accumulated deficit) Legal reserves Other reserves <b>Total equity</b>	5,373 55,580 (54,454) - 1,007 <b>7,506</b>
Total capitalisation	23,597
Indebtedness Cash Cash equivalents Trading securities Liquidity	2,798 - - <b>2,798</b>
Current financial receivable	340
Current bank debt Current portion of non-current liabilities Other current financial debt <b>Current financial debt</b>	136 1,104 - <b>1,240</b>
Current financial indebtedness	(1,898)

Non-current Bank loans

-

Bonds issued	-
Finance lease liabilities	8,436
Other non-current loans	-
Non-current financial indebtedness	8,436
Net financial indebtedness	6,538

No guarantees have been issued in respect of the current and non-current financial indebtedness as at 31 October 2011. An amount of  $\in$  772,000 of the non-current financial indebtedness and an amount of  $\in$  795,000 of the current financial indebtedness as at 31 October 2011 is secured by means of a right of pledge on the shares of OctoPlus Development B.V. pursuant to the lease facility with ABN Amro Lease Maatschappij N.V. An amount of  $\in$  136,000 of the current financial indebtedness as at 31 October 2011 is secured by means of a right of pledge over equipment, inventories, receivables and patents (with the exception of patents owned by OctoPlus PolyActive Sciences B.V.) of OctoPlus N.V. and its subsidiaries pursuant to the credit line facility with ABN AMRO Bank N.V. (see also Chapters "Operating and Financial Review – Cash Flows").

As at 31 October 2011, our net asset value per share was € 0.17 (unaudited).

As at 31 October 2011, our authorized capital amounted to  $\notin$  9,600,000 and consisted of 80,000,000 ordinary shares, all with a nominal value of  $\notin$  0.12.

As at 31 October 2011, 44,778,974 ordinary shares were outstanding. After having received the last payment of the Private Placement on 2 November 2011, all outstanding ordinary shares were fully paid up.

See "Operating and Financial Review – Contractual Obligations" for information about certain contingent obligations of ours. Please also refer to "General Information – Financial and Trading Position" for significant changes in our financial and trading position since 30 June 2011.

### **Selected Financial Information**

The selected consolidated financial information set forth below should be read in conjunction with "Operating and Financial Review", and our consolidated financial statements and their related notes that are incorporated by reference in this Prospectus. The year-end consolidated financial information has been extracted from our consolidated financial statements for the years ended 31 December 2008, 2009 and 2010 that have been audited by Deloitte Accountants B.V., independent auditors. The six months consolidated financial information has been extracted from our unaudited condensed consolidated interim financial statements as of and for the six months periods ended 30 June 2010 and 30 June 2011. Our results for the six months period ended 30 June 2011 are not necessarily indicative of our results for the full year.

Our consolidated financial statements, from which the selected consolidated financial information set forth below has been derived, were prepared in accordance with IFRS adopted by the European Union (EU). The selected consolidated financial information set forth below may not contain all of the information that is important to you.

	Year ended 31 December			Six months ended 30 Jur	
(€ in thousands)	2008*,**	2009**	2010	2010	2011
				UNAUDITED	UNAUDITED
Service revenues	€ 8,708	€ 18,636	€7,978	€3,808	€3,931
License and other revenues	8,160	370	86	22	9
Income from subsidies	55	40	265	192	12
Total (consolidated) revenues	16,923	19,046	8,329	4,022	3,952
Cost of materials and work					
contracted out	4,446	3,502	1,090	641	466
Wages and salaries	9,809	10,184	6,394	3,469	3,447
Depreciation and amortization	1,607	2,770	2,774	1,246	1,087
Other costs	5,384	4,390	3,292	1,712	1,381
Total operating costs	21,246	20,846	13,550	7,068	6,381
Operating loss	(4,323)	(1,800)	(5,221)	(3,046)	(2,429)
Interest	(1,886)	(1,157)	(981)	(494)	(456)
Result for the period	€ (6,209)	€ (2,957)	€(6,202)	€(3,540)	€(2,885)

### Consolidated Statement of Comprehensive Income Information

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of comprehensive income, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

\*\* In the year 2010, the Company changed its presentation of operating costs. This change in presentation was applied retrospectively resulting in some reclassifications in the 2008 and 2009 statement of comprehensive income, as detailed on page 49 and 50 of the 2010 Annual Report, which is incorporated by reference.

### Segmented Income Statement Information

From October 2009 onwards, OctoPlus operates in one reportable segment and does not prepare and report financial statements per segment.

### **Consolidated Statement of Financial Position Information**

	Year ended 31 December			Six months ended 30 June	
(€ in thousands)	2008*	2009	2010	2010	2011
				UNAUDITED	UNAUDITED
Intangible fixed assets	€3,044	€2,374	€2,059	€2,216	€1,920
Other fixed assets	20,955	19,752	17,347	18,729	16,462
Current assets	6,139	7,615	5,941	4,822	4,677
Total assets	30,138	29,741	25,347	25,767	23,059
Equity	575	11,343	8,935	7,970	6,206
Non-current liabilities	11,191	10,316	9,296	9,815	8,758
Current liabilities	18,372	8,082	7,116	7,982	8,095
Total equity and liabilities	€30,138	€29,741	€25,347	€25,767	€23,059

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of financial position, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

### **Consolidated Cash Flow Statement Information**

	Year ended 31 December			Six months ended 30 June		
(€ in thousands)	2008*	2009	2010	2010	2011	
				UNAUDITED	UNAUDITED	
Result for the period	€ (6,209)	€ (2,957)	€(6,202)	€(3,540)	€(2,885)	
Depreciation and amortization	1,607	2,770	2,774	1,246	1,087	
Other non-cash transactions	565	(2,608)	468	165	850	
Cash flow from operating activities	(4,037)	(2,795)	(2,960)	(2,129)	(948)	
Cash flow from investing activities	(6,698)	(1,456)	(58)	(51)	(33)	
Cash flow from financing activities	7,338	8,446	2,412	(694)	(527)	
Net cash flow	(3,397)	4,195	(606)	(2,874)	(1,508)	
Cash at beginning of period	2,515	(882)	3,313	3,313	2,707	
Cash at end of period	(882)	3,313	2,707	439	1,199	
Net cash flow	€(3,397)	€4,195	€(606)	€(2,874)	€(1,508)	

\* In the year 2009, the Company changed its accounting policies with regard to the treatment of land. This change in accounting policies was applied retrospectively resulting in some reclassifications in the 2008 statement of financial position, as detailed on page 47 of the 2009 Annual Report, which is incorporated by reference.

### **Operating and Financial Review**

You should read the following in conjunction with the "Selected Financial Information" and our consolidated financial statements and their related notes that are incorporated by reference in this Prospectus. Our consolidated financial statements have been prepared in accordance with IFRS adopted by the European Union (EU). Our six months consolidated financial information has been extracted from our unaudited condensed consolidated interim financial statements as of and for the six months periods ended 30 June 2010 and 30 June 2011. Our results for the six months period ended 30 June 2011 are not necessarily indicative of our results for the full year.

In addition to historical information, the following review includes forward-looking information that involves risks, uncertainties and assumptions. Our actual results and the timing of events could differ materially from those anticipated by these forward-looking statements as a result of many factors, including those discussed below and elsewhere in this Prospectus, particularly under "Risk Factors" and "Forward-Looking Statements".

### Overview

OctoPlus is a specialty pharmaceutical company that is globally renowned as a centre of excellence in formulation development and drug delivery. Since our establishment in 1995, we have provided advanced drug formulation and clinical scale manufacturing services to biotechnology and pharmaceutical companies worldwide. We have provided our services to more than 160 clients that have progressed more than 45 products into clinical studies and six products on to the market. In 2010 approximately 46% of our revenues from this business originated in the European Union and approximately 27% of our revenues originated from clients in North America, while the remainder was sourced from other countries. Since our establishment in 1995 through 30 June 2011, we have achieved € 95 million of consolidated revenues.

The Company is not profitable and has incurred losses in each year since 2002. Our net loss for the years ended 31 December 2010, 2009 and 2008 was  $\in$  6.2 million,  $\in$  3.0 million, and  $\in$  6.2 million, respectively, and  $\in$  2.9 million for the first six months of 2011. Through the funds raised with different private placements (see "Operating and Financial Review – Liquidity) the Company was able to continue as a going concern.

The Company has not secured sufficient working capital for its present requirements, being the next 12 months from the date of this Prospectus. Under prudent assumptions, the available net cash (cash and cash equivalents minus bank overdrafts) could deplete in the course of the second quarter of 2012 and we would require an additional  $\in 4.5 - 5.0$  million cash until 12 months after the date of this Prospectus. For details, reference is made to the section "Cash Flows – Working Capital Statement" within this Chapter.

Since October 2008, we have focussed exclusively on formulation, development and manufacturing activities for which we are reimbursed. Such contracts can further strengthen our well established contract formulation and manufacturing activities, but shall also relate to activities whereby we combine our proprietary drug delivery technologies with biopharmaceutical drugs or compounds of our clients in order to improve the properties of such biopharmaceutical drugs. We have entered into several drug delivery technology evaluation projects to develop controlled release formulations for biotech and pharmaceutical companies, whereby our clients incur the development costs and we are reimbursed for all or substantially all of our costs and will furthermore be entitled to potential future milestone and royalty payments.

We no longer invest in own product development, all of our projects are customer-funded. Until 2008, we were funding development of Locteron and OP-145, a novel proprietary peptide therapeutic for the treatment of chronic otitis media, also known as chronic middle ear infection. In October 2008, we signed a product rights acquisition agreement with Biolex for the sale and out-licensing of our commercial rights to Locteron to Biolex in exchange of up-front and milestone payments up to \$ 149 million as the

development of Locteron progresses plus further royalties on sales revenues. To date, we have received a \$ 11 million up-front payment in 2008. At the same time, we signed a product development and supply agreement with Biolex for the further development and manufacturing of Locteron. This contract had a very significant contribution to our revenues, particularly in the year 2009. OP-145 has successfully completed a Phase II clinical study. We are currently in negotiations with several parties who would like to further develop OP-145 for different indications to out-license or sell product rights and intend to bring these discussions to a close in the near future.

### Material Factors Affecting our Results of Operations and Financial Condition

We believe that the factors described in the following paragraphs have had and are expected to continue to have a material effect on our operational results and financial condition.

### Revenues

Since our establishment in 1995, we have provided advanced drug formulation and clinical scale manufacturing services to biotechnology and pharmaceutical companies worldwide. We also generated some revenues based on certain development and licensing agreements concerning our (former) product candidates and technologies. In total, we generated € 95 million of consolidated revenues since our establishment through 30 June 2011.

Over the last 16 years, we believe that we have become a leading European provider of advanced drug formulation and clinical scale manufacturing services to the pharmaceutical and biotechnology industries, with a focus on difficult-to-formulate active pharmaceutical ingredients. All activities are performed in our dedicated facilities in Leiden, the Netherlands, where we operate a cGMP manufacturing facility for final product for pre-clinical and clinical trials, and for small-scale commercial use. We offer a comprehensive range of services, including formulation, process development, analytical method development and production of both biological products, such as peptides, proteins and DNA, as well as conventional synthetic small molecules. We have strong expertise in parenteral formulations and especially biotech-derived and low-soluble compounds. Our cGMP manufacturing facility allows us to offer our clients a full range of sterile pharmaceutical production options, including freeze-drying. Typically, our clients are charged on an hourly basis for our development services, while manufacturing is charged on a lump-sum basis.

Since our establishment, we have provided services to a diverse and international group of more than 160 pharmaceutical and biotechnology companies in a total of approximately 175 projects. We have contributed to over 45 different products that have been progressed by our clients into clinical development, and to six products that have received regulatory approval. Currently, our active customer base comprises around 40 clients, which are located worldwide and include small, medium and large biotechnology as well as pharmaceutical companies. Contracts with our customers vary in size and duration, ranging from projects lasting only a few months to several years. Historically, revenues generated from individual contracts have generally been in the range of  $\leq 25,000$  to  $\leq 1.5$  million. In 2010, our five largest clients represented 45% of total revenues. We have a global client base, with 46% of revenues being generated by customers in the European Union, 27% from customers in North-America and 27% from customers in other areas of the world.

Until 2008, we invested in own product development, with our main product candidates being Locteron and OP-145, a novel proprietary peptide therapeutic for the treatment of chronic otitis media, also known as chronic middle ear infection. With our change in strategy from October 2008, we have focussed exclusively on formulation, development and manufacturing activities for which we are reimbursed.

As part of the revised strategy, we signed a product rights acquisition agreement with Biolex in October 2008 for the sale and out-licensing of our commercial rights to Locteron to Biolex in exchange of up-front and milestone payments up to \$ 149 million as the development of Locteron progresses plus further royalties on sales revenues. In 2008 and 2009, Biolex contributed  $\in$  9.9 million (58%; including an \$ 11 million upfront payment) and  $\in$  11.1 million (59%) to our consolidated revenues. 2010 and 2011 revenues from Locteron were between  $\in$  0.5 million and  $\in$  1.0 million. Biolex has indicated that it would like to start

a clinical Phase III study in the year 2012. Our services, mainly through manufacturing of GMP batches, are required during a period of at least six months before the actual start of the clinical study. If Biolex indeed commences with this study in the year 2012, or in the first part of 2013, this would require us to manufacture a large number of GMP batches which would have a significant positive impact on our 2012 and future revenues as well as our cash position.

OP-145 has successfully completed a Phase II clinical study. We are currently in advanced negotiations with several parties who would like to further develop OP-145 for different indications to outlicense or sell product rights. We expect to bring these discussions to a close in the near future. Outlicensing or sale of the product is expected to result in only small (license) revenues in the near future but might result in significant license fees and royalties in the years after in case product development progresses successfully. For the growth of our business, we are dependent on the overall development activity in the pharmaceutical and biotechnology industries. In particular, in times when financing conditions for biotechnology companies are less favorable, our business typically experiences slower growth. Conversely, when financing is more readily available to fund development projects, our business grows more rapidly. The growth of our business is furthermore related to our ability to increase our capacity in terms of human and physical resources.

Our revenues tend to be concentrated, with a limited number of large customer assignments generating a significant percentage of revenues in any given year. Our largest customers do, however, tend to change over time, as our large projects tend to last from a few months to several years on average, and are often then replaced by projects for different customers. Nevertheless, the loss of a large project can have a significant impact on our results, particularly if not replaced quickly by other projects. In 2010, our five largest clients represented 45% of our total revenues. In 2009, 59% of our total revenues (€ 11.2 million) related to the product development and supply agreement with Biolex for the further development and manufacturing of Locteron.

### **Operating Costs**

Our operating costs consist of four categories: Cost of materials and work contracted out (all costs directly related to the Company's (future) revenues; such as consumables and chemicals used in our manufacturing facility and our laboratory facilities and costs related to retaining the cGMP status for the Company's manufacturing facilities); Wages and salaries (all personnel related costs; both employees and temporary personnel); Depreciation and amortization; and Other costs (such as recruitment fees and training expenditures, marketing expenditures, professional advisors and general overhead, such as premises costs and office expenses).

Our direct and indirect costs related to research and development amounted to  $\in$  12.0 million in 2008. In October 2008, we made the strategic decision to focus exclusively on activities for which we are reimbursed. As a result, direct and indirect research and development costs decreased significantly from that day onwards and are no longer reported separately.

Approximately 8% of our total 2010 operating costs were related to Cost of materials and work contracted out. Approximately 47% of our total 2010 operating costs were related to Wages and salaries. Approximately 20% of our total 2010 operating costs were related to Depreciation and amortization. Approximately 25% of our total 2010 operating costs were related to Other costs.

## Results of Operations for the Years Ended 31 December 2008, 2009, 2010 and the Six Months Ended 30 June 2010 and 2011

### Six Months Ended 30 June 2010 and 2011

Revenues (in thousands)	Six months end	Change		
	2010 2011		€	%
	(UNAUDITED)	(UNAUDITED)		
Service revenues	€ 3,808	€ 3,931	€ 123	3%
License and other revenues	22	9	(13)	(59)
Income from subsidies	192	12	(180)	(94)
Total (consolidated) revenues	€ 4,022	€ 3,952	€ (70)	(2%)

Service revenues increased by 3% as a result of revenues from projects that are based on our proprietary drug delivery technologies.

Income from subsidies decreased by 94% because the research project for a novel approach to treat myocardial regeneration the Company participates in is ending (see also "Business – Collaborations – Erasmus University").

Operating Costs (in thousands)	Six months end	Change		
	2010	2011	€	%
	(UNAUDITED)	(UNAUDITED)		
Cost of materials and work contracted out	€ 641	€ 466	€ (175)	(27%)
Wages and salaries	3,469	3,447	(22)	(1)
Depreciation and amortization	1,246	1,087	(159)	(13)
Other costs	1,712	1,381	(331)	(19)
Total operating costs	€ 7,068	€ 6,381	€ (687)	(13%)

Cost of contracted work and other external charges decreased by 27% mainly because there were no costs incurred in the first part of 2011 for the ending subsidized project.

Depreciation and amortization decreased with 13% mainly as a result of assets becoming fully depreciated, with only a limited number of capital expenditures made over the last few years.

Other costs decreased by 19%, mainly as a result of a tight cost control.

### Years Ended 31 December 2009 and 2010

Revenues (€ in thousands)	Year ended 31 December		Change	
	2009	2010	€	%
Service revenues	€ 18,636	€ 7,978	€ (10,658)	(57%)
License and other revenues	370	86	(284)	(77)
Income from subsidies	40	265	225	563
Total (consolidated) revenues	€ 19,046	€ 8,329	€ (10,717)	(56%)

Service revenues decreased by 57%. In October 2008, we sold the worldwide commercial rights for the further exclusive development and commercialization of Locteron to Biolex. In 2009, the Company generated € 11.1 million service revenues related to this sale through process development, manufacturing of batches and release and stability testing of batches manufactured for Biolex. In 2010, only release and stability testing activities of Locteron batches manufactured were performed by

OctoPlus, which generated  $\in$  0.8 million service revenues. Non-Locteron service revenues decreased slightly from  $\in$  7.5 million to  $\in$  7.1 million.

License and other revenues decreased by 77% due to the cancellation by US-based company SurModics Inc of its license contract with OctoPlus in July 2009.

Income from subsidies increased by 563% due to increased activities on a research project for a novel approach to treat myocardial regeneration.

Operating Costs (€ in thousands)	Year ended 31 December		Change	
	2009	2010	€	%
Cost of materials and work contracted out	€ 3,502	€ 1,090	€ (2,412)	(69%)
Wages and salaries	10,184	6,394	(3,790)	(37)
Depreciation and amortization	2,770	2,774	4	-
Other costs	4,390	3,292	(1,098)	(25)
Total operating costs	€ 20,846	€ 13,550	€ (7,296)	(35%)

Cost of materials and work contracted out decreased mainly as a result of manufacturing fewer batches for Locteron.

Wages and salaries decreased due to a lower headcount (2009: 148 average FTE's and 2010: 108 average FTE's).

Other costs decreased due to minimizing the use of consultants and a tight cost control.

### Years Ended 31 December 2008 and 2009

Revenues (€ in thousands)	Year ended 31 December		Change	
	2008	2009	€	%
Service revenues	€ 8,708	€ 18,636	€ 9,928	53%
License and other revenues	8,160	370	(7,790)	(2,105)
Income from subsidies	55	40	(15)	(38)
Total (consolidated) revenues	€ 16,923	€ 19,046	€ 2,123	11%

In accordance with the agreement to sell the worldwide commercial rights for the further exclusive development and commercialization of Locteron to Biolex, we received an up-front fee of \$ 11 million (€ 7.9 million) in 2008. In addition, OctoPlus is compensated on a fee-for-service basis for all formulation and development activities plus an agreed price per batch manufactured from October 2008 onwards.

Service revenues increased mainly as a result of significantly more formulation, development and manufacturing activities for Locteron in 2009. In 2008, we were compensated on a fee-for-service basis for a three month period only (October 2008 (date of the sale) until December 2008). In 2009, we are compensated on a fee-for-service basis for all activities performed during the year 2009; generating significantly more revenues.

License and other revenues decreased as a result of the up-front fee received from Biolex in 2008.

Operating Costs (€ in thousands)	Year ended 3	1 December	Change	
	2008	2009	€	%
Cost of materials and work contracted out	€ 4,446	€ 3,502	€ (944)	(27%)
Wages and salaries	9,809	10,184	375	4
Depreciation and amortization	1,607	2,770	1,163	42
Other costs	5,384	4,390	(994)	(23)
Total operating costs	€ 21,246	€ 20,846	€ (400)	(2%)

Cost of materials and work contracted out decreased as a result of lower costs for clinical studies, toxicology studies and other research and development activities.

Depreciation and amortization increased as a result of the Company's new manufacturing facilities become operational in 2009.

Other costs decreased mainly as a result of less use of outside consultants.

### Interest

### Interest Costs

In 2008, interest costs amounted to  $\in$  1,893,000 and comprises (i)  $\in$  738,000 related to the finance lease contracts for our property, plant and equipment (including the additional facilities that came available in the beginning of 2008) and some of the equipment used in the new facilities, (ii)  $\in$  330,000 through utilization of the  $\in$  2.0 million credit facility, which we obtained from ABN AMRO Bank N.V. in 2004 and other bank charges, (iii)  $\in$  458,000 related to interest charges on the convertible bridge loans provided by Life Sciences Partners, S.R. One and Biolex and (iv) an exchange loss of  $\in$  367,000 on the settlement of the convertible bridge loan with Biolex, for which we bore the currency exchange risk.

In 2009, interest costs amounted to  $\in$  1,175,000 and comprised (i)  $\in$  1,006,000 related to the finance lease contracts for our property, plant and equipment and some of the equipment used in the new facilities, (ii)  $\in$  102,000 related to interest charges on the convertible bridge loans provided by Life Sciences Partners and S.R. One and (iii)  $\in$  67,000 through utilization of the  $\in$  2.0 million credit facility and other interest charges.

In 2010, interest costs amounted to  $\in$  993,000 and comprises (i)  $\in$  958,000 related to the finance lease contracts for our property, plant and equipment and some of the equipment used in the new facilities and (iii)  $\in$  35,000 through utilization of the  $\in$  2.0 million credit facility and other interest charges.

For the six months ended 30 June 2011, the interest costs of  $\in$  466,000 mainly related to the finance lease contracts for our property, plant and equipment and some of the equipment used in the new facilities.

### Interest Income

Interest income reflects interest earned on our deposits of cash in interest bearing accounts. Interest income for the years 2010, 2009 and 2008 amounted to  $\in$  12,000,  $\in$  18,000 and  $\in$  7,000, respectively. Interest income for the six months period ended 30 June 2011 amounted to  $\in$  10,000.

### Liquidity

Prudent liquidity risk management implies maintaining sufficient cash and marketable securities plus the availability of funding through an adequate amount of committed credit facilities. To retain a positive equity and cash balance, we strive for a positive net income, and will try to obtain additional funding through raising capital if needed. Available amounts are mainly invested in fixed-term deposits with

reputable institutions / banks and we aim to remain flexible in funding by keeping committed lending facilities available. Our primary sources of liquidity have been our funds generated from our advanced drug formulation and clinical scale manufacturing services, plus equity and debt financing. We have also received some government subsidies. With our change of focus from contract formulation and manufacturing to the development of proprietary product candidates using our drug delivery technologies from the year 2000 onwards, our capital requirements have increased and equity fund raising has been an increasingly important source of our liquidity. However, following the product rights acquisition agreement and the product development and supply agreement with Biolex, entered into in October 2008, our need for further capital was reduced considerably.

Following our incorporation in 1995, we completed our first equity financing round in December 2001, when two Dutch based venture capital investors, NPM Capital N.V. and 7X Life Sciences B.V., purchased ordinary shares for € 3.0 million. The proceeds of this financing round were utilized to develop our proprietary technology platforms, OctoDEX and PolyActive.

Our acquisition of Chienna B.V. in March 2003 had a significant impact on our capital resources, due to the up-front costs of the acquisition as well as costs due to Chienna B.V.'s ongoing operating expenses.

Since September 2004, we have a credit facility up to  $\in$  2.0 million with ABN AMRO Bank N.V., initially through our subsidiary OctoPlus Development B.V. and from January 2010 onwards through OctoPlus N.V. The actual amount available in a certain month is calculated as a percentage of the eligible receivables at the end of the previous month. As collateral, OctoPlus N.V. and its subsidiaries provide a pledge over their equipment, inventories, receivables and patents (with the exception of patents owned by PolyActive Sciences B.V.). In addition, the facility agreement contains a covenant that requires OctoPlus N.V.'s balance sheet total (adjusted for certain items) to equal at least 25% of the adjusted balance sheet total. The credit facility with ABN AMRO Bank N.V. is currently our only loan facility.

In January 2005, we issued 6,861,500 preference shares (which were converted into ordinary shares) and a subordinated convertible bond, raising a total of € 18.4 million. Investors in this round of financing included Life Sciences Partners, S.R. One, Innoven Partenaires S.A., Fagus N.V., SurModics, and several members of our management team. The stated use of proceeds of the 2005 financing included the further development of our proprietary technologies and our portfolio of product candidates.

In June 2006, we issued a further 278,600 preference shares (which were converted into ordinary shares) and a subordinated convertible bond as part of a second closing related to the January 2005 financing round. As a result, we raised an additional € 0.7 million, which was received in July 2006.

On 9 October 2006, we issued 4,301,076 ordinary shares pursuant to our Initial Public Offering ("IPO") and raised  $\in$  20 million in gross proceeds.

In March 2008, we entered into a convertible bridge loan facility with Life Sciences Partners and S.R. One for a cumulative amount of  $\in$  4.0 million. Monies drawn under this facility carried 15% interest. This facility including the interest thereon was redeemed through the issuance of 5,996,249 ordinary shares on 25 February 2009.

In April 2008, through our subsidiary OctoPlus Development B.V., we entered into a finance lease agreement with ABN Amro Lease N.V. (formerly known as Amstel Lease Maatschappij N.V.), pursuant to which ABN Amro Lease N.V. provides financing to OctoPlus Development B.V. by way of a sale and lease back of certain equipment with an investment value of  $\in$  3.7 million. The sale and lease back was executed in December 2008. As collateral for this lease facility, OctoPlus Development B.V. provided a pledge over its shares. In addition, the finance lease agreement contains a covenant that requires OctoPlus Development B.V.'s shareholders' equity to be at least equal to 30% of its adjusted balance sheet total.

In July 2008 we entered into a convertible bridge loan facility with Biolex for an amount of € 2.0 million which amount was increased by € 235,000 on 22 August 2008, by € 115,000 on 25 August 2008 and by €

400,000 on 8 September 2008. This bridge loan including accumulated interest was set off against the up-front fee payable by Biolex under the product rights acquisition agreement of October 2008.

In October 2008, we signed an exclusive contract (the 'product rights acquisition agreement') to sell our share of the commercial rights to Locteron to our former partner Biolex. As consideration, we received an \$ 11.0 million non-refundable, non-creditable up-front payment and a small equity interest in Biolex in October 2008. We are also eligible for additional milestone payments up to \$ 138 million and some additional Biolex non-voting common shares as well as royalties on future Locteron sales, all dependent upon certain pre-defined criteria with regard to the development, sale and/or out-licensing of Locteron being met.

On 25 February 2009, we issued 13,996,250 ordinary shares pursuant to a private placement and raised  $\in$  10.5 million in gross proceeds. Part of this private placement related to the conversion of the bridge loan facility entered into with Life Sciences Partners and S.R. One in March 2008 for a total amount of  $\in$  4.5 million (including accumulated interest).

In December 2009, we completed a private placement issuing 3,232,106 ordinary shares and raised € 4.0 million in gross proceeds.

In December 2010, we completed a private placement issuing 3,343,542 ordinary shares and raised € 3.9 million in gross proceeds.

In October 2011, we completed the Private Placement issuing 8,000,000 New Shares and raising € 4.0 million in gross proceeds.

The Company has never paid any dividends to its Shareholders.

### **Cash Flows**

In 2008, we reported a negative cash flow from operating activities of  $\in$  4.0 million. Investments in property and plant and equipment and intangible assets amounted to  $\in$  6.7 million. The positive net cash flow from financing activities amounted to  $\in$  7.3 million, and is related to the convertible bridge loans provided by Life Sciences Partners and S.R. One for  $\in$  4.0 million in total, a finance lease agreement with Amstel Lease Maatschappij N.V. for  $\in$  3.7 million in total and repayments on finance lease liabilities of  $\in$  0.4 million. As a result, the negative net cash flow from all operating, investing and financing activities in 2008 was  $\in$  3.4 million. Cash, cash equivalents, deposits and bank overdrafts reduced from  $\in$  2.5 million to  $\in$  0.9 million regative during the year in 2008, thereby utilizing some of the ABN AMRO Bank N.V.  $\in$  2.0 million credit facility.

In 2009, we reported a negative cash flow from operating activities of  $\in$  2.8 million. Investments in property and plant and equipment and intangible assets amounted to  $\in$  1.5 million. The positive net cash flow from financing activities amounted to  $\in$  8.4 million, and is related to our private placement of February 2009 ( $\in$  5.3 million positive cash flow) and our private placement of December 2009 ( $\in$  4.1 million positive cash flow) and repayments on finance lease liabilities of  $\in$  0.9 million. As a result, the positive net cash flow from all operating, investing and financing activities in 2009 was  $\in$  4.2 million. Cash, cash equivalents and bank overdrafts increased from  $\in$  0.9 million negative to  $\in$  3.3 million positive during the year in 2009.

In 2010, we reported a negative cash flow from operating activities of  $\in$  3.0 million. Investments in property and plant and equipment and intangible assets amounted to  $\in$  0.1 million. The positive net cash flow from financing activities amounted to  $\in$  2.4 million, and is related to our private placement of December 2010 ( $\in$  3.4 million positive cash flow) and repayments on finance lease liabilities of  $\in$  1.0 million. As a result, the negative net cash flow from all operating, investing and financing activities in 2010 was  $\in$  0.6 million. Cash, cash equivalents and bank overdrafts decreased from  $\in$  3.3 million positive to  $\in$  2.7 million positive during the year in 2010.

In the first half of 2011, we reported a negative net cash flow from operating activities of  $\in$  0.9 million. Investments in intangible and tangible assets were minimal. The negative net cash flow from financing activities amounted to  $\in$  0.5 million, and is related to repayments on finance lease liabilities. The total negative net cash flow from all operating, investing and financing activities in the first half of 2011 was  $\in$  1.5 million. Cash, cash equivalents and bank overdrafts decreased from  $\in$  2.7 million to  $\in$  1.2 million during the first half of 2011.

Following our strategic focus to use our technology in the context of a commercial agreement, whereby our clients incur the development costs and we will be reimbursed for all or substantially all costs and will furthermore be entitled to future milestone and royalty payments, compared to significant investments in product development at our own expense in the past, we have reduced our dependence on external funding.

For details on our current assets and current liabilities, reference is made to page 28 ("Consolidated balance sheet") of our 2008 Annual Report, which is incorporated by reference, page 42 ("Consolidated statement of financial position") of our 2009 Annual Report, which is incorporated by reference and page 44 ("Consolidated statement of financial position") of our 2010 Annual Report, which is incorporated by reference by reference.

#### Principal Investments

In 2008, our cash investments in plant and equipment and intangible assets amounted to  $\in$  6.7 million in total, nearly all of which is related to the design of our new manufacturing facility.

In 2009, our cash investments in plant and equipment and intangible assets amounted to  $\in$  1.5 million in total, nearly all of which is related to the design of our new manufacturing facility which became fully operational in June 2009.

In 2010, our cash investments in plant and equipment and intangible assets amounted to € 58,000 only. With our new manufacturing facility fully operational, only minimal investments in plant and equipment and intangible assets were required.

In the first half of 2011, our cash investments in plant and equipment and intangible assets were also minimal and amounted to € 33,000.

#### Working Capital Statement

OctoPlus has not secured sufficient working capital for its present requirements, being the next 12 months from the date of this Prospectus.

The Company is not profitable and has incurred losses in each year since 2002. Our net loss for the years ended 31 December 2010, 2009 and 2008 was  $\in$  6.2 million,  $\notin$  3.0 million, and  $\notin$  6.2 million, respectively, and  $\notin$  2.9 million for the first six months of 2011.

Between 2004 and 2008, the Company has made some long-term finance lease arrangements for its manufacturing, laboratory and office facilities as well as for part of the equipment used in its manufacturing facilities. This results in payment obligations of  $\in$  2.1 million for the year 2012 and  $\in$  16.3 million for the period between 2013 and 2028.

Including all net proceeds of the Private Placement, the Company's available net cash and cash equivalents balance at 31 October 2011 amounted to  $\in$  2.8 million.

Under prudent assumptions, as discussed in detail in the section "Operating and Financial Review – Working Capital Statement – Assumptions", the available net cash (cash and cash equivalents minus bank overdrafts) could deplete in the course of the second quarter of 2012 and would require an additional  $\in$  4.5 – 5.0 million cash until 12 months from the date of this Prospectus. The following assumptions have been used in these calculations:

- average historical revenues from our Fee-For-Service Business of € 7.0 million (for details see "Operating and Financial Review – Working Capital Statement – Assumptions" below);
- average non-Locteron revenues from our proprietary drug delivery technology platform of € 1.0 million (for details see "Operating and Financial Review – Working Capital Statement – Assumptions" below);
- Biolex will not start a Phase III clinical study in 2012 and will also not ask us to prepare for such a study through manufacturing of a significant number of batches on a fee-for-service basis during 2012 (for details see "Operating and Financial Review – Revenues");
- no revenue upside from our proprietary drug delivery technologies;
- no cash income from OP-145; and
- no net cash income from our mitigating action plan.

The development of our cash position until 31 December 2012 is negatively impacted by an estimated  $\notin$  1.0 – 1.5 million cash outflow mainly relating to a scheduled repayment of a subsidy advance and some advance payments received from customers in 2011 for which activities need to be performed in 2012.

However, the Company is confident that it will be able to generate sufficient funds to continue its business for the next 12 months from the date of this Prospectus by its mitigating action plan, as further detailed below. If the Company would not be successful in generating sufficient funds, it may ultimately enter into bankruptcy.

### OctoPlus' mitigating action plan

In addition to a continued focus on cost control, a further optimisation of our business development activities and to any additional funds we may generate through revenue upside from our Fee-For-Service Business and our business based upon our proprietary drug delivery technologies and OP-145 (see "Operating and Financial Review" – Revenues"), which have not been taken into account for the purpose of this working capital statement, we have the following mitigating action plan to generate net cash income to enable continued operations for the next 12 months from the date of this Prospectus.

• Raise new funds through new share issuances. OctoPlus has long-term relationships with most of its investors and is in contact with these and other investors on a continuous basis to view opportunities to raise additional funds, if needed, at favourable conditions to us. A significant part of these investors has participated in either one or more of the private placements that took place over the last few years. Although no commitments have been made by any of these investors to participate in a new financing round, we are confident that we are able to raise additional funds from these or other new investors based upon (the development of) our current product portfolio and the discussions we have with these (potential) investors to date; and,

We are dependent on market developments as these have an impact on both our revenue generating potential and our ability to raise additional funds. We are confident, however, that we will be able to generate sufficient funds through our normal operations and (if needed) the mitigating action plan identified above to continue our business for the next 12 months from the date of this Prospectus.

In the unlikely event that normal operations and our mitigating action plan (as set out above) do not generate sufficient cash to continue our business for the next 12 months from the date of this Prospectus, we will try to obtain additional funding through alternative manners, by:

• The sale / out-licensing of part of the Company's proprietary drug delivery technology platform such as our PolyActive drug delivery platform. As an example: our PolyActive drug delivery

platform is a vital part of both Locteron, the drug currently developed for ESBATech and part of our other feasibility studies. It is the only controlled release technology for injectable products in the world that has advanced to successful Phase IIb studies. Our PolyActive drug delivery platform is thereby a significant part of the Company's upside over its regular Fee-For-Service Business and could be very valuable to other parties.

 The sale of part of the Company assets such as (part of) the manufacturing facilities owned by the Company.

In case these initiatives do not provide for sufficient funding, the Company may ultimately enter into bankruptcy.

### Assumptions

In this section, we provide details on the assumptions used which would result in a cash shortage in the course of the second quarter of 2012 and a total cash need of  $\in 4.5 - 5.0$  million cash until 12 months from the date of this Prospectus. We discuss our two main sources of revenues; (i) revenues from our Fee-For-Service Business, which we estimate at  $\in 7.0$  million for the purpose of this working capital statement and (ii) revenues based on our proprietary drug delivery technologies and OP-145 which we estimate at  $\in 1.0$  million for the purpose of this working capital statement.

#### 1. Fee-For-Service Business

Contracts with our customers vary in size and duration. Historically, revenues generated from individual contracts have been in the range of  $\in$  25,000 to  $\in$  1.5 million. The duration of a project varies between a few months to several years. As a result, a significant part of the revenues generated in a year is signed during that year. Although we have long-term relationships with a significant part of our customers, the revenues generated from a customer can vary significantly from year-to-year depending on whether development of a product progresses successfully and the stage of development of the product.

The structure of our business as outlined above makes it difficult for us to estimate the revenues to be generated from our Fee-For-Service Business. Since the year 2008, revenues from our Fee-For-Service Business are around € 7 million per year. This estimate has been used in calculating the cash needs until 12 months after the publication of this Prospectus.

#### 2. Proprietary drug delivery technologies and OP-145

In addition to our Fee-For-Service Business, we generate revenues through development of products for customers that are based on our proprietary drug delivery technology platform. Non-Locteron revenues from this business have shown a strong growth since our change in strategy in the second part of 2008. Average annual revenues over the period 2009 –2011 from this business were around € 1.0 million and this number has been included in the calculation of the cash need until 12 months after the date of this Prospectus.

#### **Contractual Obligations**

It is our policy to lease our property, plant and equipment in order to reduce the substantial cash outlay associated with investments in property, plant and equipment. We have sold and leased back our headquarters for a period of twenty years and a significant part of the equipment used in our new manufacturing facility for a period of five years. In addition, we have some other less significant finance lease obligations. The sale and lease back transaction related to the Company's headquarters only relates to the building itself. The manufacturing and laboratory facilities within the building are still owned by OctoPlus. The sale and lease back transactions together resulted in total finance lease obligations of €

19.9 million as of 31 December 2008, € 18.4 million as of 31 December 2009 and € 17.2 million as of 31 December 2010. In 2010, our total finance lease and operating lease payments approximated € 2.1 million. Approximately € 1.2 million of this amount was related to the lease of our headquarters, and € 0.9 million was related to finance lease payments for other property, plant and equipment, mainly laboratory and production equipment.

€	July – Dec 2011	2012	2013 and 2014	2015 and later
Finance leases	€ 963,000	€ 1,925,000	€ 2,718,000	€ 10,712,000
Royalty obligations	-	-	-	-
Other operating commitments	-	-	-	-
Operating leases	112,000	224,000	448,000	2,442,000
Total	€1,075,000	€2,149,000	€3,166,000	€13,154,000

The following table sets out our payment obligations under contracts as of 30 June 2011 that provide for fixed and determinable payments over the periods indicated.

In addition, other amounts that we may be obligated to pay under our existing contractual relationships that are not currently fixed and determinable include royalty and milestone payments under our agreements with Leiden University Medical Center ("LUMC"), Utrecht University and IsoTis. Pursuant to our license agreement with LUMC for OP-145 CSOM we may be obligated to pay royalties on future sales. Upon entering into the license agreement regarding the OctoDEX technology with Utrecht University, we agreed that we would pay royalties on either net sales of products in the event that we undertake commercialization of a final product, or on royalties received from third parties. Pursuant to our agreement with IsoTis, IsoTis is entitled to royalty payments on our revenues from the sale of products using the PolyActive technology acquired from IsoTis and milestone payment as received from Biolex in October 2008 did not result in any royalty payment to be made to IsoTis. Payments under this agreement continue until the end of the life of the patents currently in the PolyActive patent portfolio. On 29 October 2007, US based company Integra Life Sciences Holdings Corporation ("Integra") acquired all issued and outstanding shares of IsoTis and any potential royalties will therefore need to be paid to Integra.

Although most of the subsidies we have received from the Dutch Ministry of Economic Affairs and the European Union are provided without repayment obligations, the subsidy related to the hGH-OctoDEX project was provided on the basis of a "soft loan", pursuant to which monies borrowed are repaid on the basis of revenues generated from this project. As we have discontinued this project, we anticipate that we will not have to repay the "soft loan".

We expect to be able to meet these contractual obligations from our anticipated revenues, as well as from the proceeds of the Private Placement. A significant part of our contractual obligations are contingent upon reaching various milestones, including commercialization milestones.

#### Outlook

Our prime focus is to build our business into a cash flow balanced organisation in the medium-term. We believe that this can be achieved by successfully fulfilling our contractual obligations and other service activities currently anticipated under the product rights acquisition agreement and the development and supply agreement with Biolex and by realizing further growth of our contract formulation and manufacturing activities.

If the activities and the resulting cash flows associated with the product rights acquisition agreement or the product development and supply agreement are substantially less than currently expected due to changes in the scope or the progress of product development or uncured material breach by OctoPlus, this could have a significant negative impact on our cash flow from operations.

Based on the current contract formulation and manufacturing order book, together with the ongoing discussions with our existing clients and potential new clients, we believe that we will continue to witness further growth for our non-Locteron contract formulation and manufacturing activities in the medium term. We will be able to meet increased demand from our customers by utilizing our expanded facilities in Leiden and making some further modest investments, predominantly in equipment.

In addition to our contract formulation and manufacturing activities, we have started a number of technology evaluation projects for the development and formulation of controlled release pharmaceuticals, which combine the proprietary product of our partners with our proprietary drug delivery technologies. We are reimbursed for our activities under these projects. Revenues from these projects are relatively modest. However, depending on the outcome of these technology evaluation projects (which on average run up to 6 - 24 months), our partners may decide to continue developing their product candidate, requiring our ongoing involvement in further formulation and manufacturing activities generating revenues as well as a potential license to our proprietary drug delivery technologies. We have, save for our agreement with Biolex, not yet entered into such a license agreement with a partner. Our goal is that on average one out of five of these technology evaluation projects will result in a license agreement, generating ongoing development and manufacturing revenues as well as license revenues and potentially royalty income at a later stage.

One of the areas in which we expect to grow is the development and production of specialty generics. Several blockbuster injectable controlled release drugs are coming off patent in the coming years, which opens the way to develop competitive products without infringing patents. It is expected that many companies would like to enter these multi-billion markets. Developing and manufacturing a generic version of an injectable controlled release drug is difficult and only a limited number of companies in the contract service business have the expertise and infrastructure to do so. We are experts in this field, as we have proven with the development and manufacturing of Locteron and many other products. During 2011, we have performed some early-stage formulation development work in this field on a fee-for-service basis and see an important and increasing role for OctoPlus in this field in the years to come. As the activities required to develop a generic drug are part of our standard service offering, significant investments are not required, certainly not in early stage development.

#### **Off Balance Sheet Arrangements**

We have no off balance sheet arrangements other than those presented above.

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

Our discussion and analysis of our financial position and results of operations are based on our consolidated financial statements, which have been prepared in accordance with IFRS adopted by the European Union (EU). The preparation of these financial statements requires us to make estimates and judgments that affect the reported amounts of assets, liabilities, revenues and expenses and related disclosure of contingent assets and liabilities. On an ongoing basis, we evaluate our estimates based on historical experience and make various assumptions that we believe to be reasonable under the circumstances, the results of which form the basis for making judgments about the carrying values of assets and liabilities that are not readily apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

A summary of our significant accounting policies is contained in our consolidated financial statements, which are incorporated by reference in this Prospectus. We consider the following accounting policies to be critical to the understanding of the results of our operations.

#### Impairment Test of Goodwill and Patents, Financial assets and Tangible Fixed Assets

In performing impairment testing of goodwill and patents, financial assets and tangible fixed assets, we make significant judgments and estimates to determine whether the cash flows we believe can be generated on the basis of these patents exceed the carrying value of our patents and goodwill. Determining cash flows requires the use of judgments and estimates that have been included in our

strategic plans and long-term forecasts. The data necessary for performing the impairment tests are based on our estimates of future cash flows. The discount rates used are estimated pre-tax rates that reflect specific risks relating to the relevant activities.

### **Convertible Loans**

In 2008, the Company obtained convertible bridge loans up to  $\in$  4.0 million (excluding accumulated interest) from Life Sciences Partners and S.R. One. This bridge financing is classified as a financial liability as it will or may be settled in the entity's own equity instruments and is a non-derivative for which the entity is or may be obliged to deliver a variable number of the entity's own equity instruments. The loan part and the option to convert are treated as two separate transactions. Both parts are initially booked at fair value and classified within short-term financial liabilities, as the loan might be repaid or converted within 12 months. The estimated future cash outflows for us in case of actual conversion are expected to be close or equal to zero. The value of the conversion option is therefore also close or equal to zero and the conversion option does not have impact on the 2008 financial statements.

On 25 February 2009, we issued 13,996,250 ordinary shares pursuant to a private placement and raised  $\in$  10.5 million in gross proceeds. Part of this private placement related to the conversion of the bridge loan facility entered into with Life Sciences Partners and S.R. One for a total amount of  $\in$  4.5 million (including accumulated interest).

#### **Revenue Recognition**

### Service Revenues

Sales of services are recognized in the accounting period in which the services are rendered, by reference to the stage of completion of the specific transaction, when the outcome of a transaction can be estimated reliably. The stage of completion is assessed on the basis of the actual service provided as a proportion of the total services to be provided.

### License and Other Revenues

License and other revenues include amounts earned from third parties with licenses and/or options to our intellectual property. License and other revenues are recognized when earned in accordance with the substance and under the terms of the related agreements and when it is probable that the economic benefits associated with the transaction will flow to the entity and the amount of the revenue can be measured reliably. In situations where we have continuing performance obligations, revenues related to license fee payments are deferred and the related revenue is recognized in the period of expected performance.

#### **Multiple Element Arrangements**

In certain circumstances, it is necessary to apply the recognition criteria to the separately identifiable components of a single transaction in order to reflect the substance of the transaction. Conversely, the recognition criteria are applied to two or more transactions together when they are linked in such a way that the commercial effect cannot be understood without reference to the series of transactions as a whole.

We offer arrangements whereby a customer licenses the right to use our intellectual property and purchases research and development services under one arrangement. When such multiple element arrangements exist, an element is accounted for as a separable element if it has value to the customer on a stand-alone basis and the fair value can be determined objectively and reliably.

When license revenues and service revenues are identified as separable elements in a multiple element transaction, the license revenue recognized is determined based on the fair value of the license in relation to the fair value of the arrangement taken as a whole and is recognized in accordance with the

accounting policy for license and other revenues as discussed above. The revenue relating to the service element, which represents the fair value of the servicing arrangement in relation to the fair value of the arrangement, is recognized over the service period. The fair values of each element are determined based on the current market price of each of the elements when sold separately.

### **Corporate Income Taxes**

We have a history of tax losses and recognize deferred tax assets arising from unused tax losses or tax credits only to the extent that the relevant fiscal unity has sufficient taxable temporary differences or there is convincing other evidence that sufficient taxable profit will be available against which the unused tax losses or unused tax credits can be utilized by the fiscal unity. Our management's judgment is that sufficient convincing other evidence is not currently available and a deferred tax asset is therefore only recognized to the extent that a fiscal unity has sufficient taxable temporary differences.

### **Share-Based Payments**

Share options granted to our employees are measured at the fair value of the equity instruments granted (indirect method of measurement). Fair value is determined through the use of an option-pricing model, known as the Binomial method, which considers, among others, the following variables:

- The exercise price of the option;
- The expected life of the option;
- The current value of the underlying Shares;
- The expected volatility of the share price, calculated considering the effect of dividends on the share price;
- The dividends expected on the Shares; and
- The risk-free interest rate for the life of the option.

Conditional option grants might include market conditions. These market conditions are included in the calculation of the fair value of the option.

For our share option plans, we believe that the Binomial method is most appropriate for determining fair values as this method allows accounting for non-transferability, vesting conditions and early exercise.

The result of the share option valuations and the related compensation expense is dependent on the model and input parameters used. Even though we consider the fair values reasonable and defensible based on the methodologies applied and the information available, others might derive at a different fair value for each of our share option plans.

### Claims

Third parties might claim amounts from us. These claims are considered by Management on a case by case basis. A provision is recorded in case there is a present obligation as a result of a past event, an outflow of resources is probable and a reliable estimate of the obligation can be made. In other cases, a contingent liability is recorded.

## Business

#### Overview

OctoPlus is a specialty pharmaceutical company that is globally renowned as a centre of excellence in formulation development and drug delivery. We offer clients our expertise in combination with our proprietary drug delivery technologies, in particular PolyActive, or other drug delivery platforms such as PLGA, for the development of controlled release formulations of injectable compounds. Such depot formulation products have demonstrated strong improvement of side effect profiles, increased patient compliance and better efficacy. In addition, we offer proven and reliable services in formulation development, process development, analytical method development and cGMP manufacturing. Our expertise is focused on formulation of biotech-derived compounds and small chemical molecules. One of our focus areas for the future is the development of specialty generics, which are remakes of medicines for which the patents have expired. Developing (including manufacturing) such a generic drug requires our proven and reliable services in formulation development, process development, analytical method development, process development is provided. We see promising opportunities to co-invest with partners in the specialty generics area. By co-investing we can increase our participation in the long-term financial upside of these specialty generics products.

Since our establishment in 1995, we have provided advanced drug formulation and clinical scale manufacturing services to biotechnology and pharmaceutical companies worldwide. We have provided our services to more than 160 clients that have progressed more than 45 products into clinical studies and six products onto the market. In 2010 approximately 46% of our revenues from this business originated in the European Union and approximately 27% of our revenues originated from clients in North America, while the remainder was sourced from other countries. Since our establishment in 1995 through 30 June 2011, we have achieved € 95 million of consolidated revenues.

In contrast to certain other approaches to improve the clinical benefit of injectable therapeutics, such as chemical modification (including pegylation and polymer conjugates) or protein engineering (including changes in primary structure and fusion), our technologies have the advantage of efficiently and gradually delivering a drug in its native form. We believe that products based on our technologies can be applied in many therapeutic areas. The clinically most advanced product based on our technologies is Locteron, a controlled release formulation of alpha interferon for the treatment of hepatitis C. Locteron was outlicensed to Biolex Therapeutics in October 2008, who has completed a Phase IIb study with the product. There is currently one other product based on our technologies in full development, which is a product funded by ESBATech, an Alcon/Novartis company. In addition, we have entered into several technology evaluation agreements to develop controlled release formulations for biotech and pharmaceutical companies. Depending on the outcome of these evaluation projects our clients may decide to continue developing such product candidates, requiring our ongoing involvement for the further formulation and manufacturing services as well as a license to our proprietary drug delivery technologies.

Our Executive Board has many years of experience in the areas of drug delivery, pharmaceutical formulation and manufacturing, business development and finance. Our facilities are located in Leiden, the Netherlands, and include a cGMP manufacturing facility, laboratories and offices, which we all lease under long-term contracts. As of 31 October 2011, we employed 112 people (excluding temporary staff), all of which are located in Leiden.

#### Founding and financing history and strategic development

OctoPlus was founded in 1995 by Joost Holthuis, Ph.D., who served on the Executive Board until August 2009, and Prof. Daan Crommelin, Ph.D., who served as the chairperson of our Scientific Advisory Board until mid 2009. Initially, we focused on providing pharmaceutical and biotechnology companies with advanced drug formulation and clinical scale manufacturing services.

Starting in 1996, we entered into a number of agreements with Utrecht University by which we received a worldwide exclusive license to the OctoDEX technology. While we continued to successfully grow our

contract formulation and manufacturing business, we made the strategic decision in early 2000 to start to exploit our expertise in the drug delivery area and the versatility of our OctoDEX platform by expanding into the development of proprietary products.

In 2001, we raised € 3 million from NPM Capital N.V. and 7X Life Sciences B.V. in order to progress OctoDEX-based product candidates into clinical trials. In 2003, we were able to broaden our technology base by gaining access to a second drug delivery system, PolyActive, through the acquisition of Chienna B.V. from IsoTis. Through collaboration with InnoCore Technologies, we also gained access to another controlled release technology being SynBiosys.

In 2005, Life Sciences Partners, S.R. One, Innoven Partenaires, Fagus, SurModics and certain other parties invested € 18.4 million in OctoPlus, which allowed us to fund further development of our proprietary technologies and the portfolio of product candidates we were developing.

In October 2006, we completed our IPO on Euronext Amsterdam raising € 20 million in gross proceeds, which enabled us to build a product pipeline based on our drug delivery technologies.

In October 2008, we signed a product rights acquisition agreement with Biolex for the sale and outlicensing of our commercial rights to Locteron to Biolex in exchange of \$ 149 million in up-front and milestone payments plus further royalties on sales revenues; we already received \$ 11 million as an upfront payment in that year. This agreement marked the start of OctoPlus' new strategy, moving away from developing product candidates at our own expense towards out-licensing our proprietary drug delivery technologies and co-developing products with partners.

In 2009 and 2010, we completed three private placements of shares raising  $\in$  13.9 million in gross proceeds. In addition and as part of the February 2009 private placement, a bridge loan facility entered into with Life Sciences Partners and S.R. One in March 2008 was converted into ordinary shares for a total amount of  $\in$  4.5 million (including accumulated interest). The proceeds of these private placements allowed us to make the final investments in OctoPlus' new production facility and continue to implement our new strategy. In October 2011, we completed the Private Placement raising  $\in$  4.0 million in gross proceeds, which we will use to strengthen our balance sheet and which will enable us to co-invest with partners in selected high-value projects in the area of specialty generics.

We remain focused on commercialising our controlled release expertise and these technologies on a product-by-product basis, starting with Locteron and further building a pipeline of products, based on our technologies and funded by clients. In addition to our activities in the development of controlled release products, our drug formulation and manufacturing services that started the business have remained an important part of our activities.

### Strategy

Our corporate strategy comprises three key elements;

- Continue to grow our contract formulation and manufacturing business. We intend to further grow our globally renowned activities in the field of drug delivery, formulation development, process development services, analytical method development and cGMP manufacturing for our clients.
- Build a portfolio of licensed products based on our proprietary technologies such as Locteron and ESBATech by winning technology evaluation projects, executing them successfully and converting them into licensing agreements. We intend to expand the number of technology evaluation projects in which we combine our drug delivery technologies with our clients' biopharmaceutical drugs or other therapeutics in need of clinical improvement. We intend to primarily apply our proprietary drug delivery technologies to those product opportunities where our partners can capitalise on the known safety, efficacy and often established drug development history of existing drugs. We believe that our technologies may allow us to improve the performance of many of these known biopharmaceutical drugs, of which drawbacks may include frequent or inconvenient dosing schedules, strong side effects and/or limited efficacy. Depending on the

outcome of these evaluation projects our clients may decide to continue developing such product candidates, requiring our ongoing involvement for the further formulation and manufacturing services as well as a license to our proprietary drug delivery technologies.

• Build a portfolio of specialty generics products with partners. A number of long-acting injectables are coming off patent in the next few years, enabling us to develop a generic version of those formulations. The required development process (including the manufacturing process) is complex and specialised and fits well with our expertise in difficult-to-formulate injectables. We are uniquely positioned to develop such products. We see promising opportunities to co-invest with partners in the specialty generics area. By co-investing we can increase our participation in the long-term financial upside of these specialty generics products.

### Description of our strategic activities

#### Drug product development and manufacturing

Over the last 16 years, we have become a worldwide provider of advanced drug formulation and clinical scale manufacturing services to the pharmaceutical and biotechnology industries, with a focus on difficult-to-formulate active pharmaceutical ingredients. All activities are performed in our dedicated facilities in Leiden, the Netherlands, where we operate different laboratories and a cGMP manufacturing facility for final product. We offer services from early stage development of a drug (including specialty generics) in our laboratory facilities until manufacturing final product (including specialty generics) for preclinical and clinical trials, and for small-scale commercial use in our cGMP manufacturing facility. We offer a comprehensive range of services, including formulation, process development, analytical method development and production of both biological products, such as peptides, proteins and DNA, as well as conventional synthetic small molecules. We have strong expertise in parenteral formulations and especially biotech-derived and low-soluble compounds. We have a lot of experience in developing longeracting formulations, using both proprietary and non-proprietary technologies in controlled release drug delivery. We use this experience in our technology evaluation projects and in our specialty generics focus area. Our cGMP manufacturing facility allows us to offer our clients a full range of sterile pharmaceutical production options, including freeze-drving, Typically, our clients are charged on an hourly basis for our development services, while manufacturing is charged on a lump-sum basis.

The size of the market, which is in this regard the number of injections, determines whether we have sufficient capacity in our cGMP manufacturing facility to manufacture at (early stage) commercial scale. More specifically with regard to specialty generics; with the specialty generics we target at difficult-to-formulate injectables. The development process of these specialty generics is complex and specialised and fits well within our expertise. As it is such a complex formulation, the price per injection that can be charged is relatively high. The actual market (in this case the number of injections) is not that large but annual global sales are very significant as a result of the high price charged per injection. Due to the relatively low number of injections, as outlined above, we might well be able to manufacture these specialty generics at commercial scale in our facilities. If we are not able to manufacturing (which will still take a few years) and would need to go to a third party for commercial manufacturing.

Our goal is to deliver successful outcomes to our customers by providing the highest quality of service, flexibility and expertise. We have well-established capacities and expertise in the fields of product manufacturing and state-of-the-art formulation techniques. For example, we have developed various protein formulations, including freeze-dried proteins, liposomal and other lipid-based products, micro/nanoparticles, immunotherapeutic formulations, as well as DNA-containing formulations for our clients. Primarily, we formulate products that are intended for parenteral administration. In addition, we have successfully developed other formulations, such as a pulmonary liposome formulations a protein product for colon targeting, protein-containing gels as well as pulmonary and topical formulations of oligonucleotides. We believe our achievements in the contract formulation and manufacturing business have made us well-known to the pharmaceutical and biotechnology industries as a quality service provider for the development of critical and complex formulations.

The full spectrum of our capacities in drug product development and manufacturing is set out below:

- **Characterisation.** We have an analytical group capable of characterizing the therapeutic substance provided by our clients and analysing the formulated products. Our characterization capacities include the analysis of biopharmaceuticals for which we use state-of-the-art technologies;
- **Pre-formulation and formulation.** We have developed specialised formulation technologies based on liposomes, lipid complexes, emulsions and microspheres. Our core competencies in this area include the formulation of proteins, formulations using liposomes, and the solubilisation of low soluble compounds. Our capabilities include the ability to freeze-dry clinical batches and small-scale commercial batches of therapeutic products. We can formulate high-potency compounds, such as cytostatics and hormones, on a laboratory scale;
- Drug delivery systems. We use our expertise in various drug delivery systems as an integral part
  of our formulation services. These systems include off-patent technologies based on liposomes,
  micelles and dispersions;
- Analytical method development and validation. We apply commonly used procedures in the biopharmaceutical industry for analytical method development and validation. In order to maintain and strengthen our position we also develop and evaluate novel analytical techniques;
- **Clinical manufacturing.** We have the capacity to manufacture or have manufactured a wide range of liquid and semi-solid biopharmaceutical products, including injectable, oral, dermal and pulmonary formulations, for use in clinical trials. Our facilities comprise a fully equipped production area for aseptic filling of liquid formulations, freeze-drying and colloidal preparations. With the opening of our expanded manufacturing facility in June 2009, our manufacturing capacity increased from 3,000 units per batch to 10,000 units per batch. Therefore, we cannot only produce sufficient materials for toxicology, Phase I, Phase II and Phase III clinical trials, but we can also manufacture small-scale commercial products;
- **Stability studies.** We perform stability studies in accordance with the International Conference on Harmonization guidelines.

### Market Developments

We see an ongoing trend within the pharmaceutical industry where large pharmaceutical companies reduce the size of their research & development departments, effectively using more services from contract development, manufacturing and research organisations. This has a potential positive impact on the amount of business that we can win. Since the economic downturn in 2008, capital markets have been tight impacting the ability of biotechnology companies to raise sufficient funds to continue their drug development efforts. This has a potential negative impact on the speed and size of the business that we can win.

#### Controlled release drug delivery

Our expertise in the area of controlled release is to combine our proprietary drug delivery technologies, in particular PolyActive, with our partners' biopharmaceutical drugs or other therapeutics that are in need of clinical improvement. It is by using this technology that we aim to leverage our broad expertise in formulation development and drug delivery in order to develop pharmaceutical products that improve existing approaches to the treatment of serious illnesses. We do not expect, anymore, to invest significantly in product development at our own expense but to use our technology in the context of a commercial agreement, whereby our clients incur the development costs and we will be reimbursed for all or substantially all costs and will furthermore be entitled to future milestone and royalty payments.

PolyActive is our drug delivery system based on a series of biodegradable polyethers and polyesters. We acquired the PolyActive drug delivery technology as part of our acquisition of Chienna B.V. from IsoTis in 2003. PolyActive was first introduced as a biomaterial over 15 years ago. Since then, it has been used in medical devices for orthopedic and wound repair applications in over 5,000 patients, and two medical devices based on PolyActive polymers have received FDA approval to date. We have access to a detailed safety and toxicity file concerning the PolyActive polymer that was prepared by IsoTis, the former owner of the PolyActive technology, and was filed with the FDA for the use of PolyActive in medical devices.

We focus on the use of PolyActive to create injectable and biodegradable microspheres that enable the controlled release of biopharmaceuticals. Attractive candidates that could potentially be delivered by PolyActive microspheres include therapeutic proteins, peptides and small lipophilic molecules. PolyActive refers to polyether esters comprising blocks of polybutylene terephthalate and polyethylene glycol. By varying the length and composition of these PolyActive polymers, we can tailor the drug release profile of our product candidates. We have tested various compositions of PolyActive in order to establish different in vitro release profiles of our PolyActive technology. Depending on the design and composition of the PolyActive-based microspheres, we have obtained release profiles ranging from one week to several months.

We have successfully used PolyActive for controlled release of several therapeutic biopharmaceuticals in a number of in vitro and in vivo studies. We are capable of manufacturing PolyActive microspheres under cGMP conditions in our manufacturing facility, using a well-established procedure based on a double-emulsion protocol.

We are continuously looking to further improve our proprietary drug delivery technology platform.

### **Our Products**

The following table summarises key information about the proprietary product candidates we have developed, based on our proprietary technology or active pharmaceutical ingredient. We no longer invest in own product development and have out-licensed or are in the process of out-licensing the products we've developed.

Product Candidate	Indication	Stage of Development	Partner Status	Next Expected Development Milestone
Locteron	Chronic HCV Infection	Phase IIb completed	Out-licensed and transferred to Biolex <sup>1</sup>	Biolex to start Phase III studies in 2012
OP-145 CSOM	Chronic Middle Ear Infection	Phase II completed	Co-development with Green Cross Corporation <sup>2</sup>	Discussions to out- license or sell product rights ongoing which we intend to finalize in the near future <sup>3</sup>

1 In October 2008, we concluded a product rights acquisition agreement with Biolex for the development and exclusive commercialisation of Locteron by Biolex.

2 In November 2006, we concluded an out-licensing agreement with Green Cross Corporation of South Korea for the co-

development of OP-145 CSOM and the exclusive commercialisation of OP-145 CSOM in the South Korean market. 3 See also "Operating and Financial Review – Proprietary drug delivery technologies and OP-145".

#### Locteron for Chronic Hepatitis C

Locteron is a proprietary controlled release formulation of interferon alpha for the treatment of chronic hepatitis C infection (HCV). Hepatitis C is a common disease, affecting over 170 million people worldwide. Infection by HCV is a very serious medical condition that can lead to cirrhosis of the liver, a condition where healthy liver cells are killed by infection with the hepatitis C virus and are replaced by scar tissue. Resulting damage to the liver can lead to impaired liver function, liver cancer and ultimately death.

Locteron combines interferon alpha produced by Biolex with our proprietary PolyActive microspheres. This product is designed to gradually release its active pharmaceutical ingredient over a 14-day period after a single injection. Currently marketed peoplated interferons for the treatment of HCV are dosed

once-every-week. By releasing interferon alpha in the body in a gradual manner, our PolyActive microspheres avoid the high initial blood levels of the active drug that characterise the current standard of care. These high initial levels of interferon alpha contribute to the acute flu-like symptoms commonly associated with the current standard of care for HCV. Locteron aims to provide at least the same therapeutic benefit to HCV patients with fewer, less severe and less frequent side effects and a more convenient dosing schedule. It is believed that an improved side effect profile will lead to enhanced patient compliance. The expected intended superior side effect profile of Locteron may attract and maintain patients on therapy who currently delay or refuse treatment, in particular in light of the current 48-week treatment period for HCV genotype 1, the HCV variant most prevalent in Western countries.

In 2006, OctoPlus and Biolex completed a Phase I clinical study in which the safety, pharmacokinetics and pharmacodynamics of Locteron were investigated in healthy volunteers. Results of this Phase I clinical trial showed that a single dose of Locteron was safe and well tolerated. In particular, groups receiving Locteron reported fewer, less severe and shorter lasting flu-like symptoms than those subjects receiving PEG-Intron, a commonly prescribed pegylated interferon.

In 2007, OctoPlus and Biolex completed a 12-week Phase IIa clinical trial to evaluate the safety, tolerability and preliminary efficacy of Locteron administered once-every-two-weeks in combination with ribavirin in treatment-naïve hepatitis C patients. Results of the study showed that Locteron was generally safe and well tolerated. In addition, the results showed that 12 weeks of treatment with Locteron in combination with ribavirin was effective in reducing hepatitis C virus levels.

In 2009, OctoPlus and Biolex completed a separate Phase IIa clinical study in order to generate clinical and regulatory experience with Locteron in the United States and to make a direct comparison with the current standard of care in HCV.

In 2010, Biolex completed a 48-week Phase IIb trial with Locteron, and Biolex has presented final results after 72 weeks of treatment from the Phase IIb study for Locteron versus PEG-Intron® at the International Liver Congress in April 2011, organised by the European Association for the Study of the Liver (EASL). The results confirm Locteron's improved tolerability profile, showing a statistically significant reduction in flu-like adverse events for patients treated with a Locteron injection once-every-two-weeks, compared to weekly PEG-Intron dosing. In addition, the final data also showed lower rates of depression for Locteron compared to patients receiving PEG-Intron.

BLX-883, the interferon alpha ingredient in Locteron, is produced for Biolex by Cook Pharmica in accordance with cGMP guidelines at Cook Pharmica's facilities in Indianapolis. We have been manufacturing Locteron by combining our PolyActive microspheres with BLX-883 in our licensed manufacturing facility in Leiden, the Netherlands.

### **OP-145 CSOM for Chronic Middle Ear Infection**

OP-145 CSOM is our novel, proprietary peptide therapeutic for the treatment of chronic middle ear infection (also referred to as chronic otitis media), administered locally through an eardrop formulation. Chronic middle ear infection is one of the most common childhood chronic infectious diseases worldwide. We believe that worldwide approximately 200 million people have a chronic middle ear infection. Approximately 60% of those suffering from chronic middle ear infection experience some degree of hearing impairment as a result of infection.

Acute middle ear infections are caused by a range of microbes, including bacteria and various fungi. These acute infections are usually eradicated easily using common antibiotics. However, a more serious condition may develop when certain toxic microbial degradation products linger in the mucus of the middle ear after the eradication of the microbes associated with an acute infection. In these circumstances, we believe the presence of these toxins contributes to chronic middle ear infection. The toxins associated with chronic middle ear infection are unresponsive to traditional antibiotics. Often, surgical procedures are required to treat these chronic infections and clear the degradation products from the site of infection. OP-145 CSOM is designed to neutralize these toxins at the site of infection and to locally restore the body's natural clearance mechanism. We have demonstrated in pre-clinical studies that OP-145 CSOM binds to and clears two important bacterial toxins, known as lipopolysaccharide ("LPS") and lipoteichoic acid ("LTA"). Both LPS and LTA are common constituents in the cell walls of bacteria and are released into the middle ear when the various bacteria that cause middle ear infection are destroyed by the immune system or by antibiotics.

In addition, we have demonstrated in pre-clinical studies that OP-145 CSOM interacts with lipid components on the surface of bacteria typically associated with chronic middle ear infection. By binding to these surface components, OP-145 CSOM destabilizes the cell membrane leading to bacterial cell death. In light of these two characteristics, we believe that OP-145 CSOM may offer benefits for patients with chronic middle ear infection that cannot be treated adequately with current therapies.

Based on the anticipated advantages of OP-145 CSOM, we intend initially to focus on chronic middle ear infection patients who do not respond to current treatment approaches, including patients who have recurrent chronic middle ear infection despite having undergone surgery.

In April 2006, we completed a Phase I/IIa safety and tolerability study comprising a dose-ranging assessment in 16 therapy-resistant patients who had an average disease duration of 17 years. Results from this study showed that OP-145 CSOM was safe and well tolerated, and gave an early indication of efficacy.

In a double-blind, placebo-controlled Phase II study completed in 2008, we have successfully demonstrated the safety and efficacy of OP-145 CSOM.

We are currently in advanced negotiations with several parties who would like to further develop OP-145 for different indications to out-license or sell product rights and intend to bring these discussions to a close in the near future. Out-licensing or sale of the product is expected to result in only small (license) revenues in the near future but might result in significant license fees and royalties in the years after in case product development progresses successfully.

### **Our Client base**

Since our establishment, we have provided services to a diverse and international group of more than 160 pharmaceutical and biotechnology companies, focusing mainly on protein therapeutics and to a lesser extent on small molecule drugs. We have contributed to over 45 different products that have been progressed by our clients into clinical development, and to six products that have received regulatory approval. Currently, our active customer base comprises around 40 clients, which are located worldwide and include small, medium and large biotechnology as well as pharmaceutical companies. Contracts with our customers vary in size and duration, ranging from projects lasting only a few months to several years. Historically, revenues generated from individual contracts have generally been in the range of  $\in$  25,000 to  $\in$  1.5 million. In 2010, our five largest clients represented 45% of our total revenues. We have a global client base, with 46% of our 2010 revenues being generated by customers in the European Union, 27% from customers in North-America and 27% from customers in other areas of the world.

#### Biolex Therapeutics, Inc.

In October 2008, we entered into a product rights acquisition agreement and a product development and supply agreement with Biolex following earlier agreements of 2005 and 2003 for the co-development of Locteron. Pursuant to these agreements, Biolex has obtained all commercialization rights, while we continue to provide development services and remain involved as manufacturer of Locteron. Biolex may choose to find another manufacturer of the drug product; in that event Biolex will pay OctoPlus a fee to buy the manufacturing rights. We will participate in any revenues from the commercialization of Locteron through up-front and milestone payments up to \$ 149 million as the development of Locteron progresses plus royalties on sales revenues. To date, we have received a \$ 11 million up-front payment in 2008. Biolex can terminate the product rights acquisition agreement for cause or at will by providing written notice. In the event of a termination at will by Biolex: (i) all the licenses granted to Biolex (other than cross-licenses to certain jointly owned intellectual property) will terminate, (ii) the non-compete obligations applicable to OctoPlus will terminate, (iii) Biolex will assign to OctoPlus its interest in certain intellectual property assigned to Biolex in connection with the execution of the 2008 agreement and (iv) certain license granted from Biolex to OctoPlus with respect to intellectual property retained by Biolex will become effective upon termination. In the event of a termination by Biolex (other than cross-licenses to certain jointly owned intellectual property) will terminate, (ii) the non-compete obligations applicable to OctoPlus: (i) all the licenses granted to Biolex (other than cross-licenses to certain jointly owned intellectual property) will terminate, (ii) the non-compete obligations applicable to OctoPlus will terminate and (iii) Biolex will assign to OctoPlus its interest in certain intellectual property assigned to Biolex in connection with the execution of the 2008 agreement. The scope of the intellectual property that will be assigned back to us in the event of a termination by Biolex for cause is more limited than in the event of a termination by Biolex.

In the event of the termination of the product rights acquisition agreement by either OctoPlus or Biolex, either party will have the right to also terminate the product development and supply agreement, subject to certain phase out arrangements and, in certain circumstances, transition arrangements for the transfer of manufacturing to Biolex.

Under the product development and supply agreement, OctoPlus may be subject to penalties of up to US\$ 4 million in the event OctoPlus fails to meet certain performance standards with respect to the supply of products to Biolex.

In conjunction with the above agreements, Biolex and we entered into an agreement pursuant to which intellectual property rights used in our collaboration have been transferred to OctoPlus PolyActive Sciences B.V., a 100% subsidiary of the Company. We have to refrain from certain actions which may affect the rights of Biolex under these agreements. In case of a material breach we will be subject to a penalty of  $\in$  8 million.

#### Collaborations

We have entered into several collaborative arrangements with other biopharmaceutical companies and certain academic institutions. Our most important collaborations are summarized below.

#### Erasmus University

In September 2004, we entered into a research and development agreement with the Thorax Center of Erasmus University Medical Center (the "Thorax Center") in Rotterdam, the Netherlands, for a novel treatment of myocardial (cardiac muscle) regeneration. This project is in the pre-clinical proof-of-concept phase. Under the agreement, we will own all intellectual property resulting from the study and the Thorax Center will receive a non-exclusive, royalty-free license to this intellectual property for internal research purposes. We will also pay a royalty to the Thorax Center based on our total revenues in respect of licenses we grant to third parties, subject to an overall cap. Either party may, with due notice, terminate this research and development agreement at will. This collaboration was subsidized by the Dutch Ministry of Economic Affairs from the start of the project until 30 June 2011.

#### **Green Cross Corporation**

In November 2006, we licensed OP-145 CSOM for the treatment of chronic middle ear infection to Green Cross Corporation for the exclusive sale and distribution in South Korea of products based thereon. In addition, we have agreed to perform development work for Green Cross Corporation with respect to the formulation of an OP-145 CSOM product and provide the active ingredients necessary for Green Cross Corporation to perform the studies for registration of the OP-145 CSOM product in South Korea free of charge. After we have found a commercial partner for this product, an international multicenter clinical Phase III program is scheduled to commence in which Green Cross Corporation will be responsible for conducting the clinical trials in South Korea.

Green Cross Corporation is required to pay us a number of nominal milestone payments. Future milestone payments are due upon the grant of a certain patent, upon submission for approval of the clinical study in South Korea and upon the commercial launch of the product based upon OP-145 CSOM. Green Cross Corporation is required to pay us royalties based on gross receipts for sales of OP-145 CSOM by Green Cross Corporation. Furthermore, we are entitled to 50% of any amounts that Green Cross Corporation receives from its sublicensees.

The agreement shall remain in effect until the expiration of the last valid patent relating to the OP-145 CSOM technology licensed or, if later, 15 years after the launch of the product in South Korea. The agreement may be terminated by either party upon a specified notice period or by either party at any time for unremedied default.

#### IsoTis N.V.

We have had a partnership with IsoTis (currently part of Integra) since 1999. Through our acquisition in 2003 of Chienna B.V., a subsidiary of IsoTis, we acquired from IsoTis patents and licenses to patents relating to the PolyActive technology. The license agreement with IsoTis was terminated in April 2007 and replaced by a new agreement. Under the 2007 agreement, we acquired the patents not then owned by us relating to the PolyActive technology, except for the patent family covering IsoTis' SynPlug<sup>™</sup> product. In connection with the acquisition of the patents relating to the PolyActive technology, except for the 2007 agreement provided for the transfer to us of the PolyActive name and trademarks. IsoTis retains an exclusive, world-wide, royalty-free license with a right to grant sub-licenses to make, use, sell, market and develop any PolyActive rights relating to soseouschondral and/or osteochondral plugs, fillers, cement restrictors (including IsoTis' product SynPlug) and an exclusive, world-wide, royalty-free licenses to make, use, sell, market and develop any PolyActive rights to grant sub-licenses to make, use, sell, royalty-free license with a right to grant sub-licenses to make, use, sell, market and develop any PolyActive rights relating to estimate the patent of the patent provided to the use of the PolyActive technology in a medical device to be used to measure and/or monitor blood glucose.

Under the 2007 agreement, IsoTis is entitled to a low single-digit royalty on our revenues of PolyActivebased products. In addition, we will owe Technologiestichting STW, a non-profit organization associated with the Netherlands Ministry of Economic Affairs and the Ministry of Education and Science providing research grants, a low single-digit royalty on the revenues of PolyActive-based products for the duration of the base patent, which expires in 2016. The term of the 2007 agreement is until the expiration of the last valid patent rights licensed to IsoTis.

Pursuant to our supply agreement with IsoTis, we have long-term access to a continued supply of PolyActive polymers. We also have access to updates of the Device Master File provided to the FDA and as maintained by IsoTis. IsoTis transferred the production equipment of PolyActive polymers to PolyVation B.V. In the event IsoTis wishes to sell the production equipment, we have a first right of refusal to purchase the production equipment at a predetermined price. In the event we do not purchase this equipment, IsoTis is obligated to ensure uninterrupted supply of PolyActive to us by the third party supplier under the same terms and conditions as the current arrangement.

#### Leiden University Medical Center

In 2002, we entered into a co-operation and license agreement with LUMC for OP-145 CSOM. Pursuant to that agreement, we were granted rights to develop OP-145 CSOM for the treatment of chronic middle ear infection and chronic infections of the upper respiratory tract. In March 2006, we amended our agreement with LUMC, so that we now have rights to develop OP-145 CSOM for all indications.

We have paid both a nominal up-front license fee and a nominal minimum royalty for the first and second year of our development program to LUMC. We will owe LUMC royalties based on net sales of OP-145 CSOM. We have also agreed to pay LUMC a certain percentage of any sub-license revenues we receive in relation to OP-145 CSOM.

The license agreement provides LUMC with certain termination rights if we do not pursue the development of OP-145 CSOM for a specified period of time. Furthermore, if we are responsible for a delay in the commercialization of OP-145 CSOM in a specific territory and for a specific indication, LUMC may terminate our rights to OP-145 CSOM as they relate to that territory and indication.

#### Utrecht University

Starting in 1996, we executed several agreements with Utrecht University, whereby Utrecht University granted us an exclusive license to all patents related to the OctoDEX technology. Pursuant to the relevant agreements, Utrecht University is entitled to receive royalties from us based either on our own sales of products that use the OctoDEX technology or on royalties that we receive from third parties, based on the sale of products using the OctoDEX technology.

### **Intellectual Property**

We consider patents and other intellectual property rights to be vital to the success of our business. It is our policy to actively seek patent protection for our know-how, technologies, products, and their uses. Therefore, we analyze the results of our research and development activities to identify patentable subject-matter and file new patent applications as appropriate. In addition, we will continue to evaluate and potentially acquire externally generated intellectual property to further improve our competitive position.

Patents are the cornerstone of our proprietary protection, but we may also seek other registered intellectual property rights such as trademarks or utility models. In addition, we make use of trade secrets whenever indicated. In an effort to maintain the ownership of our proprietary information, we require our consultants, advisors and collaborators to execute confidentiality and invention assignment agreements. With respect to our employees, under Dutch law, employers own the intellectual property rights of inventions made by their employees during the course of their employment.

It is our intention to seek protection for any of our inventions that hold future value for out-licensing. We file and maintain our patents in countries and jurisdictions which we consider most important to our business. These typically include the major European jurisdictions, the United States, often also Japan and, depending on the invention, other attractive markets or countries playing an important role in the pharmaceutical industries.

According to our patent strategy, we aim at building strong patent portfolios to protect our know-how broadly on the drug delivery technology level as well as, together with our collaboration partners, on the more specific product level in order to maximize the strength and duration of protection. Consequently, we expect that every major product based on one of our proprietary delivery technologies will eventually be covered by at least one delivery platform patent and at least one product-specific patent.

OctoPlus' current patent portfolio consists of 215 granted patents and 42 patent applications, which are divided into 20 patent families, of which 5 relate to OctoDEX and OctoVAX, 11 to PolyActive and the remainder to other technologies and products, including OP-145 CSOM. OctoPlus has recently disbanded, or plans to disband in the near future, a number of patents and patent applications lacking commercial prospects resulting in an easier to manage patent portfolio and lower maintenance costs. Many of these patents have been filed in our own name, while for certain others we hold exclusive licenses. Certain of our patents and patent applications are jointly held with our collaboration partners. One of our main collaboration partners is Biolex with whom we share ownership of certain patents.

OCTOPLUS is a registered trademark in the European Union. OCTODEX is a registered trademark in the Benelux countries. POLYACTIVE is a registered trademark in the United States, the European Union, Australia, Japan and Norway.

### Intellectual Property of our PolyActive Drug Delivery Technology

Our PolyActive drug delivery technology platform is covered by 11 patent families. These patents were filed between 1996 and 2006 and it is therefore expected that these patents will expire between 2016 and 2026.

Our patent protection in this area focuses on basic physical and pharmaceutical features, and in particular on the uses of the copolymers as drug carriers. Among the patents which play a key role in the PolyActive portfolio are the granted European patents EP 830 859 and EP 1 247 522, and the corresponding granted US patent US 5,980,948, which are expected to expire in 2016/2017. We own certain further patents and patent applications that relate to specific manufacturing methods, medical devices and components as well as other extensions of the PolyActive technology.

As a result of the product rights acquisition agreement between OctoPlus and Biolex, OctoPlus transferred to or co-owns with Biolex 3 patent families. Biolex has the exclusive rights in the Biolex field from the granted patents, while OctoPlus retains exclusively all other rights subject to certain non-compete provisions. These patents include, EP 830 859, EP 1 247 522, US 5 980 948, EP 1 481 017, EP 1 843 749, US 7 736 674, US 7 867 518, EP 1 306 127, EP 1 587 830, US 7 468 151 and US 7 807 617. The Biolex field concerns the controlled release formulations of interferon alpha by using our PolyActive technology.

#### Intellectual Property of Locteron and OP-145 CSOM

In 2006, we filed an international patent application WO 2006/085747 related to the formulation of Locteron which is based on a combination of interferon (including interferon alpha) with our PolyActive delivery technology. Any European, US and other national patents that may issue based on this international patent application may cover controlled release interferon compositions, and may potentially cover both Locteron and comparable products involving the delivery of interferon by microparticles. In the event that these patents are granted, they could provide patent protection for Locteron (and the production thereof) until at least 2026. In 2009, EP 1 843 749 has been granted in all the member states of the European Union for a method for the preparation of microparticles comprising interferon.

We have exclusively licensed WO 2004/067563 from LUMC and have jointly with the LUMC filed international patent application WO 2006/011792 covering OP-145 CSOM and its therapeutic applications in several important jurisdictions. If these applications lead to granted patents, they could provide exclusive protection until at least 2025. Thus far, patens derived from WO 2004/067563 have been granted in Europe, United States of America, Australia, China, Hong Kong, India, Indonesia, Japan, Philippines, South Africa, Singapore, South Korea, New Zealand and the Russian Federation. A patent derived from WO 2006/011792 has been granted in Singapore. In many other jurisdictions examination is in progress.

In spite of our efforts, it is possible that our patent applications will not be granted, or that the scope of protection allowed under any issued patent may not provide adequate protection for our technologies and products. In addition, it is possible that any of our patents may be challenged and invalidated by third parties. Competitors may find means to circumvent patents held by us or our licensors. We may not be able to enforce any of our patents against infringers in any country or region important to our business.

Even though we believe that we are free to commercialize our technologies and products with respect to their major envisioned uses, there is a risk that we may inadvertently infringe prior or future patents owned by others. We may need to acquire licenses for patents held by third parties to re-establish or maintain our freedom to operate, possibly on unfavorable terms.

#### Competition

We believe that we have established a strong competitive position in the market for contract formulation and manufacturing of clinical-grade products, particularly in the area of pharmaceutical formulations for therapeutic proteins. We were among the first independent companies to be active in this specific segment in Europe, allowing us to build significant in-house expertise in formulation and manufacturing processes. We believe that our capabilities in the contract formulation and manufacturing business are demonstrated by our high rates of repeat business.

We face competition from companies that are active in providing formulation and clinical trial manufacturing services. A number of large contract manufacturing organizations offer development services and manufacturing of clinical trial material as part of an extensive service package, such as Patheon and Recipharm. In addition, we face significant competition from smaller, more specialized firms such as Aptuit and Evotec. These firms typically focus on a specific part of the formulation market, such as biopharmaceuticals, low soluble compounds, oral drug delivery or pulmonary drug delivery. Such firms may offer formulation development services in combination with proprietary drug delivery technologies or bulk manufacturing capacity.

The ability to compete successfully will depend largely on the joint ability of us to:

- provide ongoing superior expertise on the field of formulation development, analytical method development, process development and cGMP manufacturing;
- successfully collaborate with pharmaceutical and biotechnology companies in the discovery, development and commercialisation of new products;
- obtain intellectual property protection for such products and technologies; and
- attract and retain qualified personnel.

There are many other companies developing controlled release drug delivery systems, some of which may compete with us in respect of our products in the future. However, to our knowledge, there has not been another controlled release technology to have reached the clinical stage as we have with the application of our PolyActive technology in Locteron.

### Locteron competition

Biolex and we believe that in the near to medium term interferon alpha will remain a principal element in the treatment of HCV. Biolex and we anticipate that Locteron, if approved, would compete with two approved pegylated interferon alpha products for the treatment of chronic HCV infections. These two products are Pegasys (marketed by Roche) and PEG-Intron (marketed by Schering-Plough). In addition, Locteron would face competition from conventional, unmodified forms of interferon, including Roferon-A (marketed by Roche), Intron A (marketed by Schering-Plough), and Infergen (marketed by Valeant Pharmaceuticals International and Astellas Pharma Inc.). In addition, Biolex and we are aware of other long-acting interferon alpha treatments currently in active clinical development, including product candidates from Flamel Technologies SA and Intarcia Therapeutics, Inc. In hepatitis C therapy a lot of development is ongoing with regard to new antiviral compounds, and we see Locteron serving as a core component of new combination therapies as the treatment of hepatitis C evolves to triple- and quad-drug regimens.

### **OP-145 CSOM competition**

We anticipate that OP-145 CSOM, if approved, would compete with other products that are currently marketed for the treatment of chronic middle ear infection. The current standard of care for the treatment of chronic middle ear infection involves the use of antibiotics, which are delivered either systemically or as eardrops to destroy bacteria and corticosteroids which reduce inflammation of the mucus of the middle ear. For patients who become refractory to this therapy, a surgical intervention may be necessary in order to remove resistant infectious material and inflamed mucus. Antibiotics prescribed for chronic middle ear infection include Cipro HC Otic, a combination of ciprofloxacin hydrochloride and hydrocortisone otic suspension that is marketed by Alcon, Inc., and Floxin Otic (ofloxacin otic), which is marketed by Daiichi Sankyo, Inc.

### Facilities

Our registered main place of business is located at Zernikedreef 12, 2333 CL Leiden, the Netherlands. These approximately 5,250 m<sup>2</sup> facilities comprise laboratories, a cGMP manufacturing facility and offices. We entered into a long-term lease for these premises with a remaining period of 10 - 14 years.

### **Employees**

We believe that our success will depend on our ability to attract, retain and motivate key employees. At the end of 2008, 2009 and 2010, we had 144, 132 and 95 employees (excluding temporary personnel), respectively. As of 31 December 2010, we had a total of 95 employees (representing 88 full-time employees), all of which were located in the Netherlands. As of 31 October 2011, headcount increased to a total of 112 employees (106 full-time employees; excluding temporary personnel).

We believe that relations with our employees are good. None of our employees is represented under a collective bargaining agreement. As required by Dutch law, we have established a Works Council. We believe that relations with our Works Council are good (see "Management and Employees – Works Council").

# **Management and Employees**

### General

Set out below is a summary of relevant information concerning our Executive Board, Supervisory Board, Scientific Advisory Board and other employees. In addition, we set out a brief summary of certain significant provisions of Dutch corporate law and our Articles of Association in respect of our Executive Board and Supervisory Board. See "Description of Share Capital and Corporate Governance".

### **Management Structure**

We have a two-tier board structure, consisting of an Executive Board (*Directie*) and a Supervisory Board (*Raad van Commissarissen*).

### **Executive Board**

### Powers, Composition and Function

The Executive Board is responsible for the day-to-day management of our operations under the supervision of the Supervisory Board. The Executive Board is required to keep the Supervisory Board informed, consult with the Supervisory Board on important matters and submit certain important decisions to the Supervisory Board for its approval, as more fully described below.

The Executive Board may perform all acts necessary or useful for achieving our corporate purpose, save for those acts that are prohibited by law or by our Articles of Association. The Executive Board as a whole is authorized to represent us, as are any two members of the Executive Board acting jointly.

Our Articles of Association provide that the number of members of the Executive Board will be determined by the Supervisory Board, and that the Executive Board will consist of at least one member. In the event that the Executive Board comprises two or more members, the Supervisory Board may attribute specific titles to individual members of the Executive Board, such as "Chief Executive Officer", "Chief Financial Officer" and "Chief Operating Officer".

Members of the Executive Board are appointed by the General Meeting of Shareholders following a proposal by the Supervisory Board. In view of the Dutch Corporate Governance Code, our Articles of Association provide (i) that new members of the Executive Board are appointed for a maximum term of four years, unless provided otherwise in the resolution to appoint such member, and (ii) that a retiring member of the Executive Board can be reappointed immediately for a term of not more than four years at a time.

The General Meeting of Shareholders may suspend or dismiss Executive Board members at any time. The Supervisory Board may also suspend Executive Board members at any time.

Under our Articles of Association, the following decisions of the Executive Board must be approved by the Supervisory Board:

- the acquiring, alienating, encumbering, leasing, letting and in any other way obtaining and giving the use or benefit of registered property;
- entering into agreements, whereby bank credit is granted to us;
- lending and borrowing money, with the exception of acquiring money under a credit already granted to us by a bank;

- entering into agreements by which we bind ourselves as guarantor or as severally liable co-debtor, or otherwise bind ourselves as security for a debt of a third party;
- adoption of our annual budget;
- our operational and financial objectives;
- the parameters to be applied in relation to our strategy, for example in respect of our financial ratios;
- strategic issues and alliances;
- the sale or disposition of all, or an essential part of our assets;
- the issuance and acquisition of shares and debentures chargeable against us or chargeable against a limited partnership, or a general partnership of which we are a fully liable partner;
- petition for quotation, or withdrawal of quotation from a price list of any stock exchange of any listed securities;
- entering into or terminating long-term co-operation by us with another legal entity, company, or with a limited partnership or general partnership of which we are the fully liable partner, if such cooperation or termination of co-operation is of major significance to us;
- participating by us in the capital of another company;
- investments requiring an amount equal to at least one fourth of our issued capital plus reserves, according to our balance sheet and explanatory notes;
- a proposal to amend the Articles of Association;
- a proposal to dissolve us, a proposal for a legal merger or a legal split-up, within the meaning of Title 7, Book 2 of the Dutch Civil Code, a proposal to decrease our issued capital, and a proposal to amend our Articles of Association;
- filing a petition for bankruptcy or for suspension of payments;
- the termination of the employment of a considerable number of our or a dependent company's employees simultaneously or within a short period of time;
- a significant change in the employment conditions of a substantial number of our employees and a termination of the employment of a considerable number of our employees simultaneously or within a short period of time;
- appointing staff members as officer with the general or limited power to represent us and determining their authority and title;
- making settlements;
- being a party to legal proceedings, including conducting arbitration proceedings, with the exception of taking legal measures that cannot be delayed;
- the entering into and changing of employment agreements, whereby a remuneration is granted which exceeds the annual maximum amount determined by the Supervisory Board and notified to the Executive Board in writing;

- the entering into and termination of employment agreements with (proposed) members of our management;
- establishing pension plans and granting pension rights in excess of those arising from existing arrangements;
- adoption of employee stock-option plans;
- a proposal to decrease our issued capital; and
- a proposal for a legal merger or a legal split-up, within the meaning of Title 7, Book 2 of the Dutch Civil Code.

The Supervisory Board may determine that a resolution as referred to above shall not require its approval if the amount involved does not exceed a value fixed by the Supervisory Board and notified to the Executive Board in writing. The Supervisory Board shall be entitled to require further resolutions of the Executive Board in addition to those listed above to be subject to their approval. Such further resolutions shall be clearly specified and notified to the Executive Board in writing. The authority of the Executive Board or its members to represent the Company.

Furthermore, the Executive Board shall at least once a year inform the Supervisory Board in writing of the key elements of our strategic policy, our general and financial risks and our management and control system.

### Members of the Executive Board

At our Annual General Meeting of Shareholders of 20 May 2011, Mr. Jan Egberts was appointed Chief Executive Officer.

The Executive Board is composed of the following members:

Name	Age	Position	Member Since	Term expires	Term
Jan Egberts	53	CEO	20 May 2011	20 May 2015	Four years
Susan Swarte	43	CFO	12 May 2010	12 May 2014	Four years
Gerben Moolhuizen	45	CBO	6 November 2008	12 May 2014	Four years

The business address of the members of the Executive Board is Zernikedreef 12, 2333 CL Leiden, the Netherlands.

### Jan Egberts – Chief Executive Officer

Mr. Egberts, M.D., has over 20 years of experience in the pharmaceutical sector. He graduated from Erasmus University Medical School in the Netherlands and he pursued the clinical part of his training at Harvard Medical School. He obtained his MBA from Stanford and started his business career in clinical research at Organon Teknika in Belgium. He worked for four years as a strategic consultant to life sciences companies at McKinsey & Company, and from 1994 onwards held business development and general management positions of increasing responsibility in the USA at Merck, Johnson & Johnson and Mölnycke Health Care. He became CEO of Novadel Pharmaceuticals Inc. in 2005. Subsequently, Mr. Egberts served as Senior Advisor in healthcare investments to private equity firm 3i Group and also gained experience in non-executive positions. Mr. Egberts joined OctoPlus on 1 January 2011 and was appointed Chief Executive Officer at our Annual General Meeting of Shareholders of 20 May 2011.

### Susan Swarte - Chief Financial Officer

Mrs. Swarte obtained a Master's Degree in Business Economics from Erasmus University in Rotterdam and she is a registered controller (RC), which qualification she obtained from VU University Amsterdam. She has over 16 years of experience in financial and strategic management. She worked at Unilever and Numico (now Danone), two large, international and publicly listed companies, where she was responsible for financial, logistic and reporting aspects. At Numico she professionalized the finance function in China, was Senior Business Controller at the Division Baby Food and most recently she was responsible for all financial aspects of the export business of Danone Baby Nutrition as Finance Director, Global Export. Mrs. Swarte joined OctoPlus on 1 August 2009 and was appointed to the Executive Board in May 2010.

#### Gerben Moolhuizen – Chief Business Officer

Mr. Moolhuizen received a Master's Degree in Medical Biology from Utrecht University in 1991, studied at Tohoku University, Sendai, Japan and received an MBA from the Erasmus University of Rotterdam School of Management. He then joined Pharming Group N.V. where he held positions in Business Development, eventually becoming Director, Business Development. In 1997, Mr. Moolhuizen joined ASD B.V. (currently Primagen B.V.) as Vice-President, Business Development in 1999. He joined us in 2001, as senior manager, Business Development. He became Chief Business Officer in January 2006 and joined the Executive Board in November 2008.

#### **Supervisory Board**

#### Powers, Composition and Function

The Supervisory Board is responsible for supervising the conduct of and providing advice to the Executive Board and supervising our business generally. In performing its duties, the Supervisory Board is required to act in the interests of our business as a whole. The members of the Supervisory Board are not, however, authorized to represent us in dealings with third parties.

Our Articles of Association provide that the number of Supervisory Board members will be determined by the Supervisory Board.

Our Articles of Association provide that the General Meeting of Shareholders appoints the members of the Supervisory Board following a proposal by the Supervisory Board, provided that as long as Signet Healthcare Partners, in its capacity as general partner and manager of Life Sciences Opportunities Fund L.P. and Life Sciences Opportunities Fund (Institutional) L.P., holds at least 10% of our total issued ordinary share capital, one member of the Supervisory Board shall be appointed from a nomination, drawn up by Signet Healthcare Partners. A nomination drawn up by Signet Healthcare Partners containing the names of at least two persons shall be binding, provided that the General Meeting of Shareholders may deprive such nomination of its binding character by a resolution adopted by a majority of not less than two thirds of the votes cast, representing more than half of the total issued share capital.

The current members of the Supervisory Board have been appointed for the term set out in the table below. In view of the Dutch Corporate Governance Code, the Articles of Association will provide that any newly appointed member of our Supervisory Board will serve for a maximum of four years, unless provided otherwise in the resolution to appoint the Supervisory Board member in question, and may only be reappointed twice. The General Meeting of Shareholders appoints a chairperson and the Supervisory Board appoints a deputy chairperson from amongst its members.

Under our Articles of Association, the General Meeting of Shareholders may suspend or dismiss Supervisory Board members at any time, provided that, as long as Signet Healthcare Partners holds at least 10% of our total issued ordinary share capital, any resolution to suspend or dismiss a member of the Supervisory Board, who is appointed from a nomination drawn up by Signet Healthcare Partners, other than on the proposal of Signet Healthcare Partners, may only be adopted with a majority of not less than two thirds of the votes cast, representing more than half of total issued share capital. The Articles of Association provide that the Supervisory Board members shall retire periodically in accordance with a rotation plan to be drawn up by the Supervisory Board.

Under the Articles of Association, the Supervisory Board can only adopt resolutions by an absolute majority of the total number of votes to be cast in a meeting where the majority of the Supervisory Board members then in office are present or represented. Each member of the Supervisory Board shall be entitled to one vote.

### Members of the Supervisory Board

The Supervisory Board is composed of the following members:

Name	Age	Position	Member Since	End of term
Hans Stellingsma	55	Chairperson	1 April 2001	2014
René Kuijten <sup>1</sup>	47	Member	19 January 2005	2012
Frans Eelkman Rooda	59	Member	6 November 2008	2012
James Gale <sup>2</sup>	61	Member	23 April 2009	2013
Nancy de Ruiter	38	Member	12 May 2010	2014

1 Mr. Kuijten is General Partner of Life Sciences Partners, one of our Major Shareholders, and therefore not independent within the meaning of the Dutch Corporate Governance Code.

2 Mr. Gale is General Partner of Signet Healthcare Partners, being the general partner and manager of Life Sciences Opportunities Fund L.P. and Life Sciences Opportunities Fund (Institutional) L.P., together one of our Major Shareholders, and therefore not independent within the meaning of the Dutch Corporate Governance Code.

The business address of all members of our Supervisory Board is Zernikedreef 12, 2333 CL Leiden, the Netherlands.

### Hans Stellingsma – Chairperson

Mr. Stellingsma obtained a Master's Degree from Utrecht University and a postgraduate degree (M.Phil.) from Glasgow University. He attended the Advanced Management Program at Harvard Business School. He has held numerous senior positions at a range of businesses in the Netherlands. In particular, he has served as a Managing Director at Microsoft's Dutch subsidiary and as Senior Vice President at KPN N.V. between 1993 and 1996. From 1996 to 1998, he was a member of the managing board of Origin N.V. and then he became the Chief Executive Officer of Content N.V. He was also the Managing Partner at Arthur D. Little in the Netherlands and a Senior Partner with Monitor, both global strategy consulting firms (1999 – 2004). Currently, Mr. Stellingsma is self-employed and serves on the supervisory boards of MTel B.V., De Sleutels van Zijl en Vliet and Twinning Holding B.V.

### René Kuijten – Member

Mr. Kuijten obtained his Medical Doctor Degree from Utrecht University, having obtained additional training at Harvard Medical School and the Mayo Clinics. He completed his Ph.D. at the University of Pennsylvania, where he published, among others, in the New England Journal of Medicine and Cancer Research. He received research awards from the World Health Organization and the International Union Against Cancer, and was honored with the Talma Eijkman Prize and the U-Gene Research Award for his scientific endeavors. He received an MBA from INSEAD in Fontainebleau, France. From 1992 to 2000, Mr. Kuijten was a Senior Consultant at McKinsey & Company, where he was co-leader of the European Pharmaceuticals and Healthcare Practice. He joined Life Sciences Partners (LSP) in 2001 as a General Partner. On behalf of LSP, he serves or has served on the supervisory boards or as a non-executive director of KuDOS Ltd., DNage B.V., Kreatech Holding B.V., Hybrigenics S.A., BMEYE, Nexstim and Syntaxin. Mr. Kuijten is currently board member of the NVP (Nederlandse Vereniging van Participatiemaatschappijen), member of the Dutch Life Sciences and Health Steering Committee (Regiegroep LSH), member of the NGI Valorisation Advisory Board, and chairman of the Stichting Steun Emma Kinderziekenhuis.

### Frans Eelkman Rooda –Member

Mr. Eelkman Rooda obtained a Master's Degree in Econometrics from Erasmus University Rotterdam and a Master's Degree in Business Administration from Dartmouth College. In 1977, he started his career with Esso Nederland B.V. and was subsequently employed by Esso Europe Inc. After that, he held positions of increasing responsibility at McKinsey and Company from 1982 to 1987 and 1989 to 1997. From 1987 to 1989, he worked as Department Manager Securities and Syndicates for the Algemene Bank Nederland (ABN) N.V. From 1997 to 2008, he was Chief Financial Officer of OPG Groep N.V. From 2008 until June 2011, Mr. Eelkman Rooda was Chief Financial Officer of Wessanen N.V. He is also a member of the supervisory board of De Lage Landen International B.V.

### James Gale –Member

Mr. Gale obtained a Master's Degree in Business Administration from Chicago Booth School of Business. In 1977 he started his career with E.F. Hutton & Co and Home Insurance Co. Prior to cofounding Signet Healthcare Partners in 1998, he was head of principal investment activities and head of investment banking for Gruntal & Co. Mr. Gale is a board member of AlpexPharma SA (chairman), Cedarburg Pharmaceuticals Inc., Paladin Laboratories Inc., IGI Labs, Inc., Pfenex, Inc. and SpePharm Holding B.V. During the past five years, Mr. Gale held board positions at Avantium Holding B.V., Cydex Pharmaceuticals Inc. (chairman), Relm Wireless Corp., Indevus Pharmaceuticals, Inc., Molecular Medicine Biosciences and Abrika Pharmaceuticals, Inc. Mr. Gale resigned from the board of Indevus Pharmaceuticals, Inc. early this year upon the completion of the sale to Endo Pharmaceuticals Inc.

#### Nancy de Ruiter – Member

Mrs. de Ruiter has a scientific background and holds a Ph.D. in Medical Biology from the Universities of Utrecht and San Diego. She has held several Supervisory Board positions, including at Kreatech and De Ruiter Seeds.

#### **Supervisory Board Committees**

Our Supervisory Board has appointed from among its members an Audit Committee and a Remuneration and Nominating Committee. These committees prepare the decision making of the Supervisory Board.

#### Audit Committee

Our Audit Committee assists the Supervisory Board in monitoring our systems of internal controls, the integrity of our financial reporting process and the contents of our financial statements and reports. The Audit Committee also assists the Supervisory Board in assessing and mitigating our business and financial risks.

The Audit Committee shall meet at least twice a year and shall also meet each time we propose to issue a press release containing our financial figures.

The Audit Committee consists of Mr. Eelkman Rooda (chairperson), Mr. Kuijten and Mr. Stellingsma.

### **Remuneration and Nominating Committee**

Our Remuneration and Nominating Committee advises the Supervisory Board on the remuneration of the members of the Executive Board and monitors our remuneration policy, which covers bonus plans for the members of the Executive Board. The Remuneration and Nominating Committee further advises on the selection criteria and appointment procedures for members of the Executive Board and members of the Supervisory Board, and the proposals for appointments and reappointments. It also assesses the functioning of individual members of the Supervisory Board and the Executive Board.

The Remuneration and Nominating Committee consists of Mrs. de Ruiter (chairperson), Mr. Stellingsma and Mr. Kuijten.

### **Remuneration Policy**

According to our Articles of Association, the General Meeting of Shareholders adopts the remuneration policy in respect of the remuneration of the Executive Board. The Supervisory Board establishes the remuneration of the individual members of the Executive Board, taking into account the policy adopted by the General Meeting of Shareholders, provided that arrangements in the form of Shares or rights to subscribe for Shares are subject to the approval of the General Meeting of Shareholders. Such a proposal must include the number of Shares or rights to subscribe for Shares that may be granted to the members of the Executive Board and which criteria apply to a grant or modification. The remuneration of the members of the Supervisory Board consists of a fixed cash remuneration, which is determined by the General Meeting of Shareholders.

The objective of our remuneration policy is to ensure a high direct involvement and to encourage high performance by our personnel (including members of the Executive Board). The remuneration system is based on achieving performance criteria that are determined on a yearly basis.

### **Executive Board**

The total remuneration we paid to or for the benefit of (former) members of our Executive Board in 2010 amounted to approximately € 1,018,000. The following table denotes the breakdown in remuneration of (former) members of the Executive Board in 2010.

Name	Base Salary	Bonus	Pension Contributions	Other Payments <sup>2</sup>	Total Remuneration
Jan Egberts <sup>1</sup>				€ 52,000	€ 52,00
Susan Swarte	€ 155,000	€ 28,000	€9,	€ 34,000	€ 226,00
Gerben Moolhuizen	€ 175,000	€ 28,000	€ 12,0	€ 44,000	€ 259,00
Simon Sturge	€ 375,000	€ 46,000³		€ 60,000	€ 481,00
				·	
Total	€ 705,00(	€ 102,000	€ 21,0	€ 190,000	€ 1,018,000

1 Mr. Egberts joined OctoPlus per 1 January 2011 but spent already significant time with OctoPlus during the transition period in November and December 2010 for which he received a consultancy fee of € 42,000 included under 'Other'.

2 Included under 'Other Payments' are option costs for Mr. Egberts (€ 10,000), Mrs. Swarte (€ 30,000) and Mr. Moolhuizen (€ 36,000).

3 Mr. Sturge stepped down as OctoPlus CEO per 31 December 2010. The € 46,000 bonus accrual is the maximum exposure based on 8% achievement of targets. This is subject to further discussion with Mr. Sturge.

#### Share ownership

The number of Shares currently owned by members of our Executive Board are as follows:

	Number of Shares Owned
Jan Egberts	427,119
Gerben Moolhuizen	22,500
Total	449,619

The numbers of options currently owned by members of our Executive Board are described below.

### Supervisory Board

The remuneration of the members of the Supervisory Board is determined by the General Meeting of Shareholders. As of 1 January 2008, the chairperson of the Supervisory Board receives an annual remuneration of  $\in$  31,250 and the other members of the Supervisory Board receive an annual

remuneration of  $\in$  25,000 each. In addition,  $\in$  5,000 remuneration is received by a member of the Supervisory Board for each committee participated in.

Mrs. de Ruiter owns 813,560 Shares in the Company. None of the other members of the Supervisory Board owns Shares and none of the members of the Supervisory Board own options to acquire Shares.

### **Other Information**

Except as indicated below, none of the members of the Executive Board and Supervisory Board is, or has been, subject to (i) any convictions in relation to fraudulent offences in the last five years, (ii) any bankruptcies, receiverships or liquidations of any entities in which such members held any office, directorships or senior management positions in the last five years, or (iii) any official public incrimination and/or sanctions of such person by statutory or regulatory authorities (including designated professional bodies), or disqualification by a court from acting as a member of the administrative, management or supervisory bodies of an issuer or from acting in the management or conduct of the affairs of any issuer for at least the previous five years.

Mr. Stellingsma served as Chief Operating Officer of Bikkair B.V. until February 2009. Bikkair B.V. was declared bankrupt on 4 February 2009.

#### Administrative, Management and Supervisory Bodies Conflicts of Interest

We are not aware of any potential conflict of interest between the private interests or other duties of the members of our Executive Board or Supervisory Board and their duties and responsibilities to us.

No family ties exist among the members of our Executive Board and Supervisory Board.

### **Option Plans**

#### 2003 Option Plan

In 2003, we adopted our share option plan as one of the elements of our remuneration policy for senior personnel (see "Management and Employees – Remuneration Policy"). In 2004 and 2006, we granted options pursuant to a standard share option agreement with an exercise price between  $\in$  3.43 and  $\in$  4.34 (2004) and between  $\in$  2.70 and  $\in$  4.55 (2006), subject to customary adjustment provisions and an exercise period of the options between 37 months and 60 months following the date of grant. In addition, the exercise prices of the options would be adjusted if Shares were to be issued below the applicable exercise price of the options, subject to certain conditions.

For all options granted in 2004 and 2006, any unexercised options held by an employee shall lapse in case of termination of the employment. The Supervisory Board and Executive Board have the authority to decide, in favor of the respective employee, to deviate from these terms and may decide that the exercise period for some of the options held by former employees shall be extended.

#### 2006 Option Plan

With a view to the listing of our Shares in October 2006, we adopted a new share option plan which entered into force following our listing. The share option agreements entered into prior to that date remain in place.

Under the 2006 Option Plan, the Supervisory Board has a discretionary power to grant options to our employees. The criteria for the granting of options, as well as the exercise price, will be determined by the Supervisory Board. The exercise period of the options shall be 60 months following the date of grant. The granting of options to the Executive Board can be made subject to the approval of the General Meeting of Shareholders.

All unexercised options held by an employee will lapse if we terminate that person's employment or that person resigns, except in cases of death, permanent disability or retirement. In the event that the option holder ceases to be an employee for any reason other than death, permanent disability or retirement, we may force that employee to disgorge a specified portion of any profits realized from the exercise of those options. The Executive Board has the authority to decide, in favor of the respective employee, to deviate from the above described terms.

We will not grant options to our employees and members of the Executive Board, which if exercised, would represent more than 7.5% of our issued share capital, unless the General Meeting of Shareholders approves otherwise.

In 2006, we granted options pursuant to the 2006 Option plan with an exercise price between  $\in$  2.70 and  $\in$  4.55.

#### 2010 Option Plan

On 12 May 2010, a revised option plan was approved by the General Meeting of Shareholders and the Supervisory Board. The revised plan is applicable for all options granted from 1 January 2010 onwards. Under the plan, each unconditional option has a vesting period of three years during which no options can be exercised followed by an exercise period of two years. Options rights automatically forfeit when an employee leaves the Company. Repayment of part of the award is no longer required. The exercise price of all granted options is equal to or higher than the market price of the shares on the date of the (conditional) grant.

In December 2010, Mr. Egberts received options as part of the 2010 Option plan. Mr. Egberts' option contract has an anti-dilution clause. The contract stipulates that for each follow-on financing round after the December 2010 financing round he will receive additional options with an exercise price of € 1.27 so that the potential pro rata participation of Mr. Egberts remains unchanged. As a result, Mr. Egberts received 184,888 unconditional options and 65,255 conditional options after the Private Placement of 31 October 2011. The options were received under identical conditions as the options initially granted in 2010. The vesting period for the unconditional options granted has been reduced to two years so that the end of the vesting period of the unconditional anti-dilution options is aligned with the end of the vesting period of the unconditional options in December 2010.

In 2010, we granted options pursuant to the 2010 Option plan with an exercise price between  $\in$  1.27 and  $\in$  1.41. In 2011, we granted options pursuant to the 2010 Option plan to Mr. Egberts, as referred to above, with an exercise price of  $\in$  1.27.

### **Options Granted to the Executive Board and Other Parties**

The options indicated in the table below were granted to members of the Executive Board and other employees, former employees and consultants for the periods indicated. Conditional and unconditional options are presented in separate tables.

Name	Currently Out- standing Options	Options Granted in 2004	Options Granted in 2006	Options Granted in 2010	Options Granted in 2011	End of Exercise Period of 2004 Options	End of Exercise Period of 2006 Options	End of Exercise Period of 2010 Options	End of Exercise Period of 2011 Options	Average Exercise Price of All Outstand ing Options
Jan Egberts <sup>1</sup>	1,034,888	-	-	850,000	184,888	n.a.	n.a.	2015	n.a.	1.27
Susan Swarte	178,133	-		167,000	11,133	n.a.	n.a.	2015	2016	1.41
Gerben Moolhuizen	178,133	-		167,000	11,133	n.a.	n.a.	2015	2016	1.41
Other employees	266,150	-		266,150	-	n.a.	n.a.	2014	n.a.	1.41
Former employees <sup>2</sup>	150,370	73,500	76,870	-	-	2014	2014	n.a.	n.a.	3.05
Total	1,807,674	73,500	76,870	1,450,150	207,154					€ 1.47

1 Mr. Egberts' option contract has an anti-dilution clause. It states that for each follow-on financing round after the December 2010 financing round he will receive additional options so that the potential pro rata participation of Mr. Egberts remains unchanged. As a result, Mr. Egberts received 184,888 unconditional options after the Private Placement of 31 October 2011.

2 Former employees are options held by Mr. Holthuis, our co-founder who no longer works for OctoPlus.

Name	Currently Out- standing Conditional Options	Conditional Options Granted in 2010	Conditional Options Granted In 2011	End of Exercise Period of 2011 Conditional	End of Exercise Period of 2012 Conditional	End of Exercise Period of 2013 Conditional	Average Exercise Price of All Outstanding
	-			Options	Options	Options	Options
Jan Egberts <sup>1</sup>	365,255	300,000	65,255	2017	2018	2019	1.27
Susan Swarte	155,867	155,867	-	2017	2018	2019	1.41
Gerben Moolhuizen	155,867	155,867	-	2017	2018	2019	1.41
Total	676,989	611,734	65,255				€ 1.33

1 Mr. Egberts' option contract has an anti-dilution clause. It states that for each follow-on financing round after the December 2010 financing round he will receive additional options so that the potential pro rata participation of Mr. Egberts remains unchanged. As a result, Mr. Egberts received 65,255 conditional options after the Private Placement of October 2011.

### **Employment Agreements**

We have employment agreements with each of the members of the Executive Board. All employment agreements have an indefinite term and can be terminated, subject to a notice period of two months for the employee and four months for us.

The employment agreements with each of the members of the Executive Board provide for severance payments in the event of termination. If OctoPlus terminates the employment agreement for either of them, we are obliged to pay a severance amount equal to 1.5 times the monthly salary per year of service, up to a maximum of a full year salary. Furthermore, if the employment agreement is terminated within six months after a change of control, they are entitled to a severance amount of 1.5 times the

monthly salary per year of service inclusive of the average bonus payment received over the three years preceding the change of control, up to a maximum of a full year's salary including the average bonus payment received.

Members of the Supervisory Board are not eligible to severance payments in case of departure.

### **Directors Indemnification and Insurance**

In order to attract and retain qualified and talented persons to serve as members of the Executive Board or the Supervisory Board, we provide such persons with protection through a directors' and officers' insurance policy.

Furthermore, we provide indemnification for members of our Executive Board and Supervisory Board against (i) substantiated costs made within the bounds of reasonableness with respect to conducting a defense (including lawyers fees), at law and otherwise, against third party claims for reimbursement of damages, or payment of fines, (judicially imposed) penalty payments and the like; and (ii) financial consequences of court rulings and resolutions of governmental authorities and amounts due relating to settlements that actually and in reasonableness have been paid by such member to third parties, due to an act or failing to act in the performance of his duties as member of the Executive Board or Supervisory Board or any other function he performs at our request, save where such act or the failing to act could be characterized as seriously culpable, or to the extent the loss of capital is covered by an insurance.

### **Pension Plan and Other Benefits**

We provide our employees with a collective pension plan. From 1 February 2006, we have had a defined contribution plan, which replaced our defined benefit plan, which was closed on 31 January 2005.

Mr. Moolhuizen is the only Executive Board member participating in OctoPlus' pension scheme. Mr. Egberts does not get any compensation for pension. Mrs. Swarte receives the equivalent of the employer's contribution to OctoPlus' pension scheme in cash. All but one other employees participate in OctoPlus' pension scheme.

We provide our employees with collectively negotiated health and retirement benefits in line with market practices in the Netherlands.

### **Works Council**

As required by Dutch law, we have established a Works Council. Works councils in the Netherlands have the authority to advise on certain company decisions proposed by the general meeting of shareholders or the management board, including but not limited to a change of control. Employers are also required to submit certain statutory defined matters that are viewed as 'social policy' (affecting employment terms and conditions) to the works council for prior approval.

# **Major Shareholders**

The following table presents information about the ownership of our Shares as of the date of this Prospectus for each existing shareholder we know to beneficially own 5% or more of our Shares. This information is sourced from the public online register of the AFM website.

Shareholder	%	Date of notification
LSP III Management B.V. <sup>1</sup>	14.9%	21 December 2010
Signet Healthcare Partners (USA) <sup>2</sup>	14.4%	23 December 2010
Glaxo Smithkline plc (USA)	9,7%	31 October 2011
J.J.M. Holthuis (the Netherlands) <sup>3</sup>	9.5%	31 December 2009
Innoven Partenaires S.A. (France) <sup>4</sup>	9.3%	25 February 2009
A Strating (the Netherlands)	5.7%	21 January 2011

1. Life Sciences Partners has invested in OctoPlus through its group company LSP III Management B.V.

2. Signet Healthcare Partners acts as general partner and manager of Life Sciences Opportunities Fund II, L.P. and Life Sciences Opportunities Fund (Institutional) II, L.P.

3. The shares owned by Mr. Holthuis, our co-founder, are held via Sodoro B.V.

4. Innoven Partenaires S.A. acts as management company of various FCPI Funds (*Fonds Commun de Placement dans l'Innovation*).

Certain members of our Executive Board and one member of our Supervisory Board hold Shares as described in "Management and Employees – Share ownership" and "Management and Employees – Supervisory Board". Our Executive Board members hold options to purchase Shares. We describe these options in more detail in "Management and Employees – Option Plans".

Except as disclosed above, we are not aware of any person who, as of the date of this Prospectus, directly or indirectly, has a beneficial interest in 5% or more of our Shares. Our Major Shareholders have the same voting rights as other holders of the Shares.

# **Related Party Transactions**

Except as disclosed in "Management and Employees" and except as disclosed below, the members of the Executive Board, the members of the Supervisory Board, and the Major Shareholders have had no interest in any transactions to which we were a party since 1 January 2006, or in any transactions which we entered into prior thereto and under which we or the other parties still have ongoing obligations.

In, April 2008, we entered into a convertible loan agreement with Life Sciences Partners and S.R. One, two of our Major Shareholders, as a result of which Life Sciences Partners and S.R. One provided a convertible loan of € 2.0 million each. The convertible loan carried 15% interest. This convertible loan agreement including the interest thereon was redeemed through the issuance of 5,996,249 New Shares to Life Sciences Partners and S.R. One on 25 February 2009 at € 0.75 per share.

On 25 February 2009, Mr. Sturge, CEO of OctoPlus from 15 September 2008 until 31 December 2010, acquired 133,333 ordinary shares as part of a private placement at € 0.75 per share.

In December 2009, we raised € 4.0 million in gross proceeds in a private placement. Signet Healthcare Partners participated in the private placement. Mr. James Gale, a member of our Supervisory Board, is managing partner at Signet Healthcare Partners. As a result of his involvement in the financing, Mr. Gale did not have voting rights during the Supervisory Board meeting in which the private placement was authorized. Mrs. Nancy de Ruiter participated in the private placement by purchasing 500,000 shares at identical conditions as the other participants. Mrs. De Ruiter was appointed as a new member of the Supervisory Board on 12 May 2010.

In December 2010, we raised € 3.9 million in gross proceeds in a private placement. Mr. Jan Egberts, our Chief Executive Officer, participated in the private placement and acquired 127,119 shares at identical conditions as the other participants. Mrs. Nancy de Ruiter, a member of our Supervisory Board, participated in the private placement and acquired 63,560 shares at identical conditions as the other participants. As a result of her involvement in the financing, Mrs. de Ruiter did not take part in any discussion or decision-making during the Supervisory Board meeting in which the private placement was authorized. Life Sciences Partners participated in the private placement. Mr. René Kuijten, a member of our Supervisory Board, is general partner at Life Sciences Partners. As a result of his involvement in the financing, Mr. Kuijten did not take part in any discussion or decision-making during the supervisory Board meeting in which the private placement was authorized.

In October 2011, we raised  $\in$  4.0 million in gross proceeds in the Private Placement. Mr. Jan Egberts, our Chief Executive Officer, participated in the Private Placement and acquired 300.000 New Shares at identical conditions as the other participants. As a result of his involvement in the financing, Mr. Egberts did not take part in any discussion or decision-making during the Executive Board meeting in which it was resolved to pursue the Private Placement, Mrs. Nancy de Ruiter, a member of our Supervisory Board. participated in the Private Placement and acquired 250,000 New Shares at identical conditions as the other participants. As a result of her involvement in the financing, Mrs. De Ruiter did not take part in any discussion or decision-making during the Supervisory Board meeting in which the Private Placement was authorized. Life Sciences Partners participated in the Private Placement. Mr. René Kuijten, a member of our Supervisory Board, is general partner at Life Sciences Partners. As a result of his involvement in the financing, Mr. Kuijten did not take part in any discussion or decision-making during the Supervisory Board meeting in which the Private Placement was authorized. Signet Healthcare Partners participated in the Private Placement. Mr. James Gale, a member of our Supervisory Board, is managing partner at Signet Healthcare Partners. As a result of his involvement in the Private Placement, Mr. Gale did not take part in any discussion or decision-making during the Supervisory Board meeting in which the Private Placement was authorized.

# **Description of Share Capital and Corporate Governance**

## General

Our business was commenced by a company incorporated under Dutch law as a private company with limited liability (*besloten vennootschap met beperkte aansprakelijkheid*), by deed executed on 20 July 1995 under the name OctoPlus B.V., which is currently named OctoPlus Development B.V. and is one of our 100% subsidiaries. We were incorporated on 29 May 1997 under Dutch law, under the name OctoPlus B.V. on 4 October 2006, we converted OctoPlus B.V. into OctoPlus N.V. a limited liability company under Dutch law. We trade under the name OctoPlus. We are registered with the Trade Register of the Chamber of Commerce for The Hague, the Netherlands under number 28075073. Our corporate seat is in Leiden, the Netherlands and our office address is Zernikedreef 12, 2333 CL Leiden, the Netherlands. We can be contacted by telephone on + 31 (0)71 5244044, by fax on +31 (0)71 5244043, by email at octoplus@octoplus.nl, or through our website which is <u>www.octoplus.nl</u>. The contents of our website are expressly not incorporated by reference into this Prospectus.

Our articles of association were last amended by deed of amendment, executed on 30 May 2011, before Mr. D.F.M.M. Zaman, civil law notary in Rotterdam, the Netherlands. The certificate of no objection of the Ministry of Justice for that amendment was granted on 26 May 2011, under number N.V. 597.465.

Set out below is a summary of certain relevant information concerning our share capital, certain significant provisions of Dutch corporate law and a brief summary of certain provisions of our articles of association (the "Articles of Association").

This summary does not purport to give a complete overview and should be read in conjunction with the Articles of Association, together with relevant provisions of Dutch law, and does not constitute legal advice regarding these matters and should not be considered as such.

### **Corporate Objects**

Pursuant to Article 3 of our Articles of Association, our corporate objects are:

- to incorporate, to participate in any way whatsoever in, to manage and to supervise businesses and companies, in particular, but not limited to those involved in the pharmaceutical and (bio)medical industries, and, more in particular, those involved in the development and exploitation of pharmaceutical and biomedical processes and products and the rendering of research and development services;
- to develop and trade in patents, trade marks, licenses, know-how and other intellectual property rights;
- to render advice and services to businesses and companies with which we form a group and to third parties;
- to borrow, to lend and to raise funds, including the issue of bonds, promissory notes or other securities or evidence of indebtedness, as well as to enter into agreements in connection with the aforementioned activities;
- to grant guarantees, to bind ourselves and to pledge our assets for obligations of businesses and companies with which we form a group and on behalf of third parties;
- to acquire, dispose of, manage and exploit registered property and items of property in general; and

• to do all that is connected therewith or may be conducive thereto, all to be interpreted in the broadest sense.

## **Share Capital**

### Authorized and Issued Share Capital

At the date of this Prospectus, our authorized share capital amounts to  $\in$  9,600,000 and is divided in 80,000,000 ordinary shares, with a nominal value of  $\in$  0.12 each. As a result of the Private Placement there are 44,778,974 ordinary shares issued and outstanding.

The following table sets forth information about our issued share capital as of the date of this Prospectus.

	As of the date of this Prospectus
Ordinary shares	44,778,974
Unconditional options <sup>1</sup>	2,007,674
Total	46,786,648

<sup>1</sup> 1,807,674 unconditional options issued under our share option plans and 200,000 unconditional options issued under our agreement with Theratechnologies (see "Description of Share Capital and Corporate Governance – Share Capital – Outstanding Options" below). In addition, 676,989 conditional options were granted to members of our Executive Board.

Currently, neither we nor any of our subsidiaries hold any of our shares. All shares that are outstanding as of the date of this Prospectus are fully paid up.

## Form and Trading of Shares

Our ordinary shares are in registered form (*aandelen op naam*) and are traded through the book-entry facilities of Euroclear Netherlands. No share certificates will be issued. We are responsible for keeping a shareholders' register.

## **Outstanding Options**

On various occasions in 2004, 2006 and 2010, we entered into share option agreements with employees. In 2011, we granted both conditional and unconditional options pursuant to the anti-dilution clause in the option contract of Mr. Egberts. The option contract with Mr. Egberts stipulates that for each follow-on financing round after the December 2010 financing round he will receive additional options so that the potential pro rata participation of Mr. Egberts remains unchanged. Pursuant to all option agreements, a total of 1,807,674 unconditional options and 676,989 conditional options are currently outstanding. Each unconditional option entitles its holder to acquire an ordinary share for an exercise price, depending on the calendar year in which the unconditional option was granted, between € 3.43 and € 4.34 (2004; only options with an exercise price of € 3.43 currently outstanding), between € 2.70 and € 4.55 (2006), between € 1.27 and € 1.41 (2010) and € 1.27 (2011). For all unconditional option agreements entered into in 2004 and 2006, the exercise period starts at the date of grant and in the event that the option holder ceases to be an employee for any reason other than death, permanent disability or retirement, we may force that employee to disgorge a specified portion of any profits realized from the exercise of those options. For all unconditional option agreements entered into in 2010, each unconditional option has a vesting period of three years during which no options can be exercised followed by an exercise period of two years. The yesting period for the unconditional anti-dilution options granted to Mr. Egberts in 2011 is two years so that the end of the vesting period of these anti-dilution options is aligned with the end of the vesting period for the unconditional options initially granted to Mr. Egberts in December 2010. Options rights automatically forfeit when an employee leaves the Company. Repayment of part of the award is no longer required. Except for these differences, all unconditional option agreements have identical terms (see "Management and Employees - Option Plans").

In March 2010, the Supervisory Board granted 167,000 conditional options to acquire shares at an exercise price of € 1.41 per share to both Mrs. Swarte and Mr. Moolhuizen. The number of unconditional options each person will receive depends on certain pre-defined performance criteria for each person in the years 2010, 2011 and 2012, with 1/3 of the conditional options related to each of the three years. Any conditional options not granted based upon the 2010, 211 and 2012 performance may be granted in 2013 when certain pre-defined performance criteria related to the 2013 performance of each of the Executive Board members are met. Mrs. Swarte and Mr. Moolhuizen both achieved 20% of their 2010 performance criteria and as a consequence each person received 11,133 unconditional options from the 55,667 conditional options not received can still be granted to both Mrs. Swarte and Mr. Moolhuizen in 2014 in case certain pre-defined performance criteria are met by Mrs. Swarte and Mr. Moolhuizen in the year 2013.

In December 2010, the Company granted 300,000 conditional options at an exercise price of € 1.27 per share to Mr. Egberts, who formally started working for OctoPlus on 1 January 2011. The number of unconditional options Mr. Egberts will receive from the conditional options granted depends on certain pre-defined performance criteria in the years 2011, 2012 and 2013, with 1/3 of the conditional options related to each of the three years. The pre-defined performance criteria might include market conditions.

In December 2011, the Company granted 65,255 conditional options at an exercise price of € 1.27 per share to Mr. Egberts related to the anti-dilution clause in Mr. Egberts' option contract. The number of unconditional options Mr. Egberts will receive from the conditional options granted depends on certain pre-defined performance criteria in the years 2011, 2012 and 2013, with 1/3 of the conditional options related to each of the three years. The pre-defined performance criteria might include market conditions.

On 26 September 2007, we granted options to Theratechnologies to acquire 200,000 Shares at an exercise price of € 3.95 per Share, equal to the average closing price of the Shares for the ten trading days immediately prior to the date of our license agreement with Theratechnologies. These options are exercisable at any time until the earlier of ten years following date of execution or the fifth anniversary of the date of termination of the agreement with Theratechnologies.

#### Issue of Shares and Pre-emptive Rights

In general, each holder of our ordinary shares shall have a pre-emptive right to subscribe for newly issued ordinary shares, pro rata to the aggregate amount of that holder's ordinary shares. Such preemptive rights do not apply, however, in respect of (i) ordinary shares issued for a non-cash contribution, and (ii) ordinary shares issued to our employees.

On 20 May 2011, the General Meeting of Shareholders has designated the Executive Board, for a period that ends on 20 November 2012, as the corporate body authorized to issue ordinary shares, subject to the limitation that the authority is limited to 30% of the entire issued share capital as it stands per 20 May 2011, and/or to limit or exclude pre-emptive rights in relation to an issuance of shares with the prior approval of our Supervisory Board. This designation may be extended, either by an amendment to the Articles of Association, or by a resolution of the General Meeting of Shareholders, for a period not exceeding five years in each case. A designation pursuant to a resolution of the General Meeting of Shareholders shall require the proposal of the Executive Board, which is subject to the prior approval of the Supervisory Board.

Designation of the Executive Board as the corporate body with these authorities by the Articles of Association may be revoked by an amendment of the Articles of Association. Designation of the Executive Board as the corporate body with these authorities by the General Meeting of Shareholders cannot be revoked, unless determined otherwise at the time of designation.

Following termination of the Executive Board's authority to issue ordinary shares and/or to limit or exclude pre-emptive rights in relation to an issue of shares, the General Meeting of Shareholders shall be authorized to do so, unless it has delegated these authorities to another corporate body.

No resolution of the General Meeting of Shareholders or the Supervisory Board is required for an issue of shares pursuant to the exercise of a previously granted right to subscribe for shares.

#### Granting of Rights to Subscribe for Shares

On 20 May 2011 the General Meeting of Shareholders has designated the Executive Board, for a period that ends on 20 November 2012, as the corporate body authorized to grant rights to subscribe for shares, with the prior approval of our Supervisory Board. This designation may be extended, either by an amendment to the Articles of Association, or by a resolution of the General Meeting of Shareholders, for a period not exceeding five years in each case. A designation pursuant to a resolution of the General Meeting of Shareholders shall require a proposal by the Executive Board, which is subject to prior approval of the Supervisory Board.

Designation of the Executive Board as the corporate body with the authority to grant rights to subscribe for shares by the Articles of Association may be revoked by an amendment of the Articles of Association. Designation of the Executive Board as the corporate body with the authority to grant rights to subscribe for shares by the General Meeting of Shareholders cannot be revoked, unless determined otherwise at the time of designation.

#### Acquisition of Shares in Our Capital

We may acquire our own fully paid shares at any time for no consideration (*om niet*). Furthermore, subject to certain provisions of Dutch law and our Articles of Association, we may acquire fully paid shares in our own capital if (i) our shareholders' equity less the payment required to make the acquisition, does not fall below the sum of the paid-in and called-up share capital plus the reserves as required to be maintained by Dutch law or by our Articles of Association (such excess, the "Distributable Equity") and (ii) we and our subsidiaries would thereafter not hold shares or hold a pledge over our shares with an aggregate nominal value exceeding 50% of our issued share capital.

Other than those shares acquired for no consideration, shares may only be acquired subject to a resolution of the Executive Board, which is approved by the Supervisory Board, and authorized by the General Meeting of Shareholders. Such authorization from the General Meeting of Shareholders for the acquisition of our shares shall specify the number and class of these shares that may be acquired, the manner in which these shares may be acquired and the price range within which shares may be acquired. Such authorization may be valid for no more than 18 months.

The General Meeting of Shareholders has authorized the Executive Board to acquire a maximum of 10% of our issued ordinary shares for a period of 18 months from the meeting of shareholders which was held on 20 May 2011, at a purchase price between the nominal value of the shares and 110% of the average price of our ordinary shares during five trading days before the repurchase.

No authorization from the General Meeting of Shareholders is required for the acquisition of fully paid shares for the purpose of transferring these shares to employees pursuant to our share option plan. Any shares we hold in our own capital may not be voted or counted for voting quorum purposes.

### **Reduction of Share Capital**

The General Meeting of Shareholders may, at the proposal of the Executive Board, which proposal is subject to the approval of the Supervisory Board, resolve to reduce our issued and outstanding share capital by cancelling our shares, or by amending our Articles of Association to reduce the nominal value of our shares.

## **Dividends and Other Distributions**

We may only make distributions to our shareholders in so far as our shareholders' equity exceeds the Distributable Equity.

Under our Articles of Association, we may only make a distribution of dividends to our shareholders if our statutory annual accounts demonstrate that such distribution is legally permitted. The Executive Board may, however, subject to the prior approval of the Supervisory Board, declare an interim dividend. The General Meeting of Shareholders may, furthermore, upon the proposal by the Executive Board, with the prior approval of the Supervisory Board, resolve that a distribution is made from the Distributable Equity and that a distribution of dividends on the ordinary shares shall not be paid in whole or in part in cash, but in shares.

Furthermore, the corporate body authorized to issue ordinary shares, and/or to limit or exclude preemptive rights in relation to an issuance of shares, and/or to grant rights to subscribe for shares, shall be authorized to resolve that in respect of any issuance of shares and/or granting of rights for shares, the nominal value of these shares shall be paid up on account of our equity, to the extent it exceeds the aggregate of the paid in and called up part of our share capital and the reserves which must be maintained pursuant to the law and to resolve that pre-emptive rights in respect of such issuance or granting are restricted or limited.

Claims to dividends and other distributions not made within five years from the date that such dividends or distributions became payable, will lapse and any such amounts will be considered to have been forfeited to us (*verjaring*).

#### Stichting Continuïteit OctoPlus and Preference Shares

In 2007, OctoPlus incorporated Stichting Continuïteit OctoPlus (the 'Foundation') to safeguard OctoPlus' interests and those of OctoPlus' enterprise and to protect, insofar as possible, the Company's continuity, the Company's independence and the Company's corporate identity. In 2010, the Supervisory Board decided that the costs associated with the Foundation were deemed to be too high in relation to the limited risks against which the Foundation would protect the Company. As a result, the Board of the Foundation decided to liquidate the Foundation in March 2010. The foundation was liquidated on 5 May 2010.

#### **General Meetings of Shareholders and Voting Rights**

The annual General Meeting of Shareholders shall be held within six months after the end of each financial year. Our financial year is equal to a calendar year.

An Extraordinary General Meeting of Shareholders may be convened, whenever our interests so require, by the Executive Board or the Supervisory Board. Shareholders representing alone or in aggregate at least one-tenth of our issued and outstanding share capital may, pursuant to the Dutch Civil Code and our Articles of Association, request the district court to authorize such shareholder to convene a General Meeting of Shareholders.

The notice convening any General Meeting of Shareholders shall be sent no later than the 42nd day prior to the meeting and shall include an agenda stating the items to be dealt with. Holders of shares (including holders of the rights conferred by law upon holders of depositary receipts issued with a company's cooperation for shares in its capital) who, alone or in the aggregate, own shares representing at least 1% of our issued and outstanding capital have the right to request the Supervisory Board or the Executive Board to place items on the agenda of the General Meeting of Shareholders. If such proposals are submitted to the Executive Board or the Supervisory Board in time for the Executive Board to put these proposals on the agenda for the next meeting, or announce them prior to the meeting by means of a supplementary notice with due observance of the aforementioned notice period, the Executive Board or the Supervisory Board shall be obliged to do so, provided that no important interest (*zwaarwichtig belang*) we have dictates otherwise.

All notices of General Meetings of Shareholders, all announcements concerning dividend and other distributions, and all other announcements to holders of shares (including holders of rights conferred by law upon holders of depositary receipts issued with a company's cooperation for shares in its capital), shall be effected by means of a press release and on our website.

Shareholders may only attend the General Meeting of Shareholders and participate in the voting in respect of shares which are registered in their name on the record date as specified in the notice of the meeting. The record date will be on the 28th day prior to the date of the meeting.

Each of our ordinary shares is entitled to one vote. Shareholders may vote by proxy. The voting rights attached to any of our shares held by us are suspended as long as they are held in treasury.

Decisions of the General Meeting of Shareholders are taken by an absolute majority of votes cast, except where Dutch law provides for a qualified majority.

#### Amendment of Our Articles of Association and Change of Our Corporate Form

The General Meeting of Shareholders may resolve to amend our Articles of Association, subject to a proposal by the Executive Board, which requires the approval of the Supervisory Board.

The General Meeting of Shareholders may furthermore resolve to change our corporate form. A change of our corporate form shall require a resolution to amend our Articles of Association, subject to a proposal by the Executive Board, which requires the approval of the Supervisory Board.

#### **Statutory Merger and Statutory Demerger**

The General Meeting of Shareholders may resolve that we enter into a statutory merger or demerger (which term includes both a split-up and a spin-off), subject to a proposal by the Executive Board, which requires the approval of the Supervisory Board. In the event we are the acquiring company, the Executive Board may resolve to enter into a statutory merger or demerger, unless one or more shareholders representing at least 5% of our issued share capital request the Executive Board within one month of the announcement of the merger or demerger, to convene a General Meeting of Shareholders.

#### **Dissolution and Liquidation**

We may only be dissolved by a resolution of the General Meeting of Shareholders subject to a proposal by the Executive Board, which requires the approval of the Supervisory Board.

In the event of a dissolution, our business will be liquidated in accordance with Dutch law and our Articles of Association, and the members of the Executive Board will (unless otherwise determined by the General Meeting of Shareholders) become liquidators, acting under supervision of the Supervisory Board. During liquidation, the provisions of our Articles of Association will remain in force to the extent possible.

The balance remaining after settlement of debts shall be transferred to the holders of ordinary shares, in proportion to the aggregate nominal amount of their ordinary shares.

#### Dutch Corporate Governance Code

On 9 December 2003, the Dutch Corporate Governance Committee, also known as the Tabaksblat Committee, released the Dutch Corporate Governance Code. With effect from 1 January 2009, the Dutch Corporate Governance Code has been amended by the Frijns Committee (Code Frijns). The Dutch Corporate Governance Code contains 22 principles and 129 best practice provisions for executive boards, supervisory boards, shareholders and general meetings of shareholders, financial reporting, auditors, disclosure, compliance and enforcement standards.

Dutch companies listed on a government-recognized stock exchange, whether in the Netherlands or elsewhere, are required under Dutch law to disclose in their annual reports whether or not they apply the provisions of the Dutch Corporate Governance Code that relate to the executive board or supervisory board and, if they do not apply, to explain the reasons why. The Dutch Corporate Governance Code provides that if a company's general meeting of shareholders explicitly approves the corporate governance structure and policy and endorses the explanation for any deviation from the best practice provisions, such company will be deemed to have applied the Dutch Corporate Governance Code.

We acknowledge the importance of good corporate governance. The Executive Board and Supervisory Board have reviewed the Dutch Corporate Governance Code, generally agree with its basic provisions, and have taken and will take any further steps they consider appropriate to implement the Dutch Corporate Governance Code.

We fully support the principles and best practice provisions of the Dutch Corporate Governance Code and will apply with the relevant best practice provisions of the Dutch Corporate Governance Code, subject to the exceptions set out below.

#### Exceptions in the application of the Dutch Corporate Governance Code

II.2.4 If options are granted, they shall, in any event, not be exercised in the first three years after the date of granting. The number of options to be granted shall be dependent on the achievement of challenging targets specified beforehand.

As detailed in the section 'Management and Employees – Options Granted to the Executive Board and Other Parties", the Supervisory Board granted Mr. Egberts, Mrs. Swarte and Mr. Moolhuizen a number of conditional and unconditional options in 2010 and granted Mr. Egberts a number of conditional and unconditional options in 2011. The unconditional options granted to Mr. Egberts in 2011 can be exercised after two years. None of the other options can be exercised within three years after unconditional grant. The unconditional options were granted by the Supervisory Board based upon the individual's past performance or as a sign-on bonus. This is a deviation from the Dutch Corporate Governance Code, where unconditional options will only be granted when certain pre-defined, challenging, individual targets have been met. The Supervisory Board perceived that granting of these unconditional options to the Executive Board members was in the best interest of the Company and its stakeholders.

III.2.1 The supervisory board members, with the exception of not more than one person, shall be independent within the meaning of best practice provision III.2.2.

Our Supervisory Board consists of five members. Two members are not independent as they are employed by Signet Healthcare Partners and Life Sciences Partners, both of which have more than 10% Shares in OctoPlus. Signet Healthcare Partners and Life Sciences Partners both have a long-term interest in the Company and were willing to back this up by making senior partners with relevant knowledge and experience available to the Company. The Supervisory Board considers that Mssrs. Gale and Kuijten fit the profile of the Supervisory Board and their contributions outweigh any perceived disadvantage of non-independence.

IV.3.1 Meetings with analysts, presentations to analysts, presentations to investors and institutional investors and press conferences shall be announced in advance on the website and by means of press releases. Provision shall be made for all shareholders to follow these meetings and presentations in real time, for example by means of web casting or telephone lines. After the meetings, the presentations shall be posted on the company's website.

Considering our size, it would create an excessive burden to provide facilities that enable shareholders to follow in real time the meetings and presentations, referred to in the best practice provision. We will, however, ensure that presentations are posted on our website immediately after the meetings in question.

IV.3.13 The company shall formulate an outline policy on bilateral contacts with the shareholders and publish this policy on its website.

We will formulate an outline policy on bilateral contacts with the shareholders and publish this policy on our website in the near future.

V.3.1 The external auditor and the audit committee shall be involved in drawing up the work schedule of the internal auditor. They shall also take cognizance of the findings of the internal auditor.

We feel that our financial reporting will be sufficiently monitored by our audit committee and will, in view of our size, initially not appoint an internal auditor. The Audit Committee will evaluate the need for an internal auditor every year. Based on this evaluation will the Supervisory Board make a recommendation to the Executive Board whether or not to appoint an internal auditor.

#### **Disclosure of Information**

As a Dutch company listed on Euronext Amsterdam, we will be required to publish our annual accounts within four months after the end of each financial year and our half-yearly figures within two months after the end of the first six months of each financial year. In addition, we will also be obliged to publish interim management statements (*inter alia* containing an overview of important transactions and their financial consequences) in the period starting ten weeks after and six weeks before the first and second half of each financial year, or, alternatively, we are obliged to publish quarterly financial statements.

We must also make public certain inside information by means of a press release. Pursuant to the Financial Supervision Act, inside information is knowledge of concrete information directly or indirectly relating to the issuer or the trade in its securities which has not been made public and publication of which could significantly affect the trading price of the securities. The Financial Supervision Act contains specific rules intended to prevent insider trading.

#### **Obligations of Shareholders to Make a Public Offer**

The European Directive on Takeover Bids (2004/25/EC) has been implemented in Dutch legislation in the Financial Supervision Act. Pursuant to the Financial Supervision Act, a shareholder who has acquired 30% of our voting rights has the obligation to launch a public offer for all shares and depositary receipts issued for shares. The legislation also applies to persons acting in concert who jointly acquire substantial control.

#### **Squeeze Out Procedures**

Pursuant to section 2:92a of the Dutch Civil Code, a shareholder who for his own account contributes at least 95% of our issued capital may institute proceedings against our other shareholders jointly for the transfer of their shares to the claimant. The proceedings are held before the Enterprise Chamber and can be instituted by means of a writ of summons served upon each of the minority shareholders in accordance with the provisions of the Dutch Code of Civil Procedure (*Wetboek van Burgerlijke Rechtsvordering*). The Enterprise Chamber may grant the claim for squeeze out in relation to all minority shareholders and will determine the price to be paid for the shares, if necessary upon advice of one or three experts. Once the order to transfer becomes final before the Enterprise Chamber, the person acquiring the shares shall give written notice of the date and place of payment and the price to the holders of the shares to be acquired whose addresses are known to him. Unless the addresses of all of them are known to him, he shall also publish the same in a newspaper with a national circulation.

The offeror under a public offer is also entitled to start a squeeze out procedure, within three months after the public offer, if following the public offer he contributes at least 95% of the class of shares and represents at least 95% of the total voting rights attached to these shares. In the event of a mandatory offer, the mandatory offer price is in principal deemed to be a reasonable price, which has to be accepted by minority shareholders. In the event of a voluntary public offer, the offered price is considered reasonable as long as 90% of the shares have been acquired.

The same legislation also entitles each remaining minority shareholder to demand a squeeze out if the offeror has acquired at least 95% of the class of shares held by him, representing at least 95% of the total voting rights in that class. This procedure must be initiated with the Enterprise Chamber within three months after the end of the period for tendering shares in the public offer. With regard to price, the same procedure as for squeeze out proceedings initiated by the offeror applies.

#### Notification of Holdings of Voting Rights and Capital Interest

Pursuant to the Financial Supervision Act, certain notification requirements apply to us as well as to holders of our shares due to the fact that we are a listed company. The notification requirements are summarized below.

Pursuant to the Financial Supervision Act, each person whose holding of voting rights and/or capital interest, directly or indirectly, amounts to 5% or more must notify the AFM without delay by means of a standard form or through the automated notification system of the AFM. Any person who, directly or indirectly, acquires or disposes of an interest in our share capital or voting rights must without delay give written notice to the AFM, if, as a result of such acquisition or disposal, the percentage of capital interest or voting rights held by such person, directly or indirectly, reaches, exceeds or falls below the following thresholds: 5%, 10%, 15%, 20%, 25%, 30%, 40%, 50%, 60%, 75% and 95%.

We are required to notify the AFM of any changes in our share capital and voting rights. More specifically, we are required to notify the AFM without delay of any changes in our share capital if our share capital has changed by 1% or more compared to the previous disclosure in respect of our share capital. We are also required to notify the AFM without delay of any changes in the voting rights, insofar as it has not already been notified at the same time as a related change in our share capital. Changes in our share capital and voting rights of less than 1% must also be notified; these changes can be notified at any time but at the latest within eight days after the end of each calendar quarter. The AFM will publish such notifications in a public register. If, as a result of such change, a person's direct or indirect interest in our share capital or voting rights passively reaches, exceeds or falls below the abovementioned thresholds, the person in question must give notice to the AFM no later than the fourth trading day after the AFM has published the change in our share capital and/or voting rights in the public register.

In addition, annually within four weeks after the end of the calendar year, every holder of 5% or more of our shares or voting rights whose interest has changed in the period after his most recent notification to the AFM, which change relates to the composition of the notification as a result of certain acts (e.g., the exchange of shares (an actual interest) for depositary receipts for shares (which is a potential interest) or the exercise of a right to acquire shares (pursuant to which the potential interest becomes an actual interest)) must notify the AFM of such changes.

A person is deemed to hold the interest in our share capital or voting rights that is held by its controlled undertakings as defined in the Financial Supervision Act. The controlled undertaking does not have a duty to notify the AFM because the interest is attributed to the undertaking in control, which as a result has to notify the interest as an indirect interest. Any person, including an individual, may qualify as an undertaking in control for the purposes of the Financial Supervision Act. A person who has a 5% or larger interest in our share capital or voting rights and who ceases to be a controlled undertaking for purposes of the Financial Supervision Act must without delay notify the AFM. As of that moment, all notification obligations under the Financial Supervision Act will become applicable to the former controlled undertaking.

For the purpose of calculating the percentage of capital interest or voting rights, amongst others, the following interests must be taken into account: (i) shares or depositary receipts for shares or voting rights directly held (or acquired or disposed of) by any person, (ii) shares or depositary receipts for shares or voting rights held (or acquired or disposed of) by such person's controlled undertakings or by a third party for such person's account or by a third party with whom such person has concluded an oral or written voting agreement (including a discretionary power of attorney), and (iii) shares or depositary receipts for shares for shares or voting rights which such person, or any controlled undertaking or third party referred to above, may acquire pursuant to any option or other right held by such person (including, but not limited to, on the basis of convertible bonds). As a consequence, the notification should indicate whether the interest is held directly or indirectly, and whether the interest is an actual or a potential interest.

In addition, a notification obligation for certain cash settled instruments has been introduced. As a result hereof the notification obligation is extended to holders of financial instruments where the increase in

value of the instruments is dependent on the increase in value of the (underlying) shares or related dividends.

A holder of a pledge or right of usufruct in respect of shares or depositary receipts for shares can also be subject to the reporting obligations of the Financial Supervision Act, if such person has, or can acquire, the right to vote on the shares or, in the case of depositary receipts for shares, the underlying shares. If a pledgee or usufructuary acquires the voting rights on the shares or depositary receipts for shares, this may trigger a corresponding reporting obligation for the holder of the shares or depositary receipts for shares or shares. Special rules apply with respect to the attribution of shares or depositary receipts for shares or voting rights which are part of the property of a partnership or other community of property.

The Financial Supervision Act contains detailed rules that set out how its requirements apply to certain categories of holders, including but not limited to (managers of) investment funds, investment managers, custodians, market makers, clearing and settlement institutions, brokers and credit institutions.

Pursuant to the Financial Supervision Act, members of our Executive Board and Supervisory Board must notify the AFM of their interest in our share capital and voting rights within two weeks of their appointment as a member of our Executive Board or our Supervisory Board. Any subsequent change of their interest in our share capital and voting rights must be notified to the AFM without delay.

The notifications referred to in this paragraph should be made in writing by means of a standard form or electronically through the notification system of the AFM.

It is envisaged that the threshold for shareholders of 3% will be added, pursuant to a draft bill on the amendment of the Financial Supervision Act. In addition, on the basis of this draft bill, the Company would be obliged to publish its strategy on its website. In connection therewith, shareholders with a capital interest of 3% or more will have to disclose whether they have any objections against the published strategy.

Another draft bill, also amending the Financial Supervision Act, includes an extension of the notification obligations in respect of substantial holdings on the basis of economic long positions. Pursuant to the proposal the notification obligations would be extended to voting rights and capital holdings in financial instruments of which the value depends on the increase in value of the shares or dividend rights and which will be settled other than in those shares. On the basis of this proposal, (legal) persons which/who hold certain financial instruments such as contracts for differences and total return equity swaps should notify their interest as of 3%.

It is unclear if and when the above described proposed legislation will become effective. Once effective, the definitive amendment of the Financial Supervision Act may deviate from the above described proposals, due to amendments during the legislative process.

#### **Market Abuse Regime**

The rules on preventing market abuse set out in the Financial Supervision Act are applicable to us, the members of our Executive Board and Supervisory Board, other insiders and persons performing or conducting transactions in our securities. Certain important market abuse rules set out in the Financial Supervision Act that are relevant for investors are described hereunder.

We are required to make inside information public. Pursuant to the Financial Supervision Act, inside information is knowledge of concrete information directly or indirectly relating to the issuer or the trade in its securities which has not yet been made public and publication of which could significantly affect the trading price of the securities. We must also provide the AFM with this inside information at the time of publication. Furthermore, we must without delay publish the inside information on our website and keep it available on our website for at least one year.

It is prohibited for any person to make use of inside information within or from the Netherlands or a non-European Union member state by conducting or effecting a transaction in our shares. In addition, it is prohibited for any person to pass on inside information to a third party or to recommend or induce, on the basis of inside information, any person to conduct a transaction. Furthermore, it is prohibited for any person to manipulate the market, for instance by conducting transactions which could lead to an incorrect or misleading signal of the supply of, the demand for or the price of the securities.

Our insiders within the meaning of the Financial Supervision Act are obliged to notify the AFM when they carry out or cause to be carried out, for their own account, a transaction in our shares or in securities the value of which is at least in part determined by the value of our shares. Insiders within the meaning of the Financial Supervision Act in this respect are: (i) members of our Executive Board and our Supervisory Board, (ii) other persons who have a managerial position and in that capacity are authorized to make decisions which have consequences for our future development and business prospects and who, on a regular basis, can have access to inside information relating, directly or indirectly, to us, and (iii) certain persons closely associated with the persons mentioned under (i) and (ii) designated by the Dutch Market Abuse Decree (*Besluit marktmisbruik Wft*).

This notification must be made no later than the fifth business day after the transaction date on a standard form drawn up by the AFM. This notification obligation does not apply to transactions based on a discretionary management agreement as described in section 8 of the Dutch Market Abuse Decree. Under certain circumstances, the notification may be delayed until the date on which the value of the transactions amounts to  $\in$  5,000 or more in the calendar year in question.

If a member of our Executive Board or Supervisory Board has notified a transaction to the AFM under the Financial Supervision Act as described above under "Notification of Holdings of Voting Rights and Capital Interest", such notification is sufficient for purposes of the Financial Supervision Act as described in this paragraph.

We have adopted an internal code on inside information in respect of the holding of and carrying out of transactions in our shares by the members of our Executive Board and Supervisory Board and our employees. Further, we have drawn up a list of those persons working for the Company who could have access to inside information on a regular or incidental basis and we have informed the persons concerned of the rules on insider trading and market manipulation including the sanctions which can be imposed in the event of a violation of those rules.

# **Market Information**

### **Euronext Amsterdam**

Our Shares are listed and traded on Euronext Amsterdam. We are subject to Dutch securities regulations and supervision by the relevant Netherlands authorities.

### **Market Regulation**

The AFM is the market regulator in the Netherlands and supervises market conduct of the parties active on the securities markets. The AFM has supervisory powers with respect to the application of takeover regulations and compliance with financial reporting requirements. It also supervises financial intermediaries and investment advisers. Since the implementation of the Prospectus Directive on 1 July 2005, the AFM is furthermore the competent authority for approving all prospectuses published for admission of securities to trading on Euronext Amsterdam, except for prospectuses approved in other European Economic Area states that are used in the Netherlands in accordance with applicable passporting rules. Due to the implementation of the Market Abuse Directive and related Commission Directives on 1 October 2005, the AFM has taken over from Euronext Amsterdam its supervisory powers with respect to publication of inside information by listed companies. The surveillance unit of Euronext Amsterdam continues to monitor and supervise all trading operations.

# Taxation

The following is a general summary and the tax consequences as described here may not apply to a holder of Shares. Any potential investor should consult his tax adviser for more information about the tax consequences of acquiring, owning and disposing of Shares in his particular circumstances.

## **Dutch Taxation**

This taxation summary solely addresses the principal Dutch tax consequences of the acquisition, ownership and disposal of Shares. It does not consider every aspect of taxation that may be relevant to a particular holder of Shares under special circumstances or who is subject to special treatment under applicable law. Where in this summary English terms and expressions are used to refer to Dutch concepts, the meaning to be attributed to such terms and expressions shall be the meaning to be attributed to the equivalent Dutch concepts under Dutch tax law. Where in this summary the terms "the Netherlands" and "Dutch" are used, these refer solely to the European part of the Netherlands. This summary also assumes that we are organized, and that our business will be conducted, in the manner outlined in this Prospectus. A change to such organizational structure or to the manner in which we conduct our business may invalidate the contents of this summary, which will not be updated to reflect any such change.

This summary is based on the tax law of the Netherlands (unpublished case law not included) as it stands at the date of this Prospectus. The law upon which this summary is based is subject to change, perhaps with retroactive effect. Any such change may invalidate the contents of this summary, which will not be updated to reflect such change.

Where in this section "Dutch Taxation" reference is made to "your Shares", that concept includes, without limitation, that:

- 1. you own one or more Shares and in addition to the title to such Shares, you have an economic interest in such Shares;
- 2. you hold the entire economic interest in one or more Shares;
- 3. you hold an interest in an entity, such as a partnership or a mutual fund, that is transparent for Dutch tax purposes, the assets of which comprise one or more Shares, within the meaning of 1. or 2. above; or
- 4. you are deemed to hold an interest in Shares, as referred to under 1. to 3., pursuant to the attribution rules of article 2.14a, of the Dutch Income Tax Act 2001 (*Wet inkomstenbelasting 2001*), with respect to property that has been segregated, for instance in a trust or a foundation.

#### **Dutch Taxation – Taxes on Income and Capital Gains**

#### **Resident Holders of Shares**

#### General

The summary set out in this section "Dutch Taxation – Taxes on Income and Capital Gains – Resident Holders of Shares" applies only to a holder of Shares who is a "Dutch Individual" or a "Dutch Corporate Entity".

For the purposes of this section you are a "Dutch Individual" if you satisfy the following tests:

- a. you are an individual;
- b. you are resident, or deemed to be resident, in the Netherlands for Dutch income tax purposes, or you have elected to be treated as a resident of the Netherlands for Dutch income tax purposes;
- c. your Shares and any benefits derived or deemed to be derived therefrom have no connection with your past, present or future employment, if any; and
- d. your Shares do not form part of a substantial interest (*aanmerkelijk belang*) or a deemed substantial interest in us within the meaning of Chapter 4 of the Dutch Income Tax Act 2001 (*Wet inkomstenbelasting 2001*).

Generally, if a person holds an interest in us, such interest forms part of a substantial interest, or a deemed substantial interest, in us if any one or more of the following circumstances is present:

- 1. You either alone or, in the case of an individual, together with your partner (*partner*), if any, or pursuant to article 2.14a, of the Dutch Income Tax Act 2001 (*Wet inkomstenbelasting 2001*) own or are deemed to own, directly or indirectly, either a number of Shares in us representing five per cent. or more of our total issued and outstanding capital (or the issued and outstanding capital of any class of our Shares), or rights to acquire, directly or indirectly, Shares, whether or not already issued, representing five per cent. or more of our total issued and outstanding capital (or the issued and outstanding capital (or the issued and outstanding capital (or the issued and outstanding capital of any class of our Shares), or profit participating certificates (*winstbewijzen*) relating to five per cent. or more of our annual profit or to five per cent. or more of our liquidation proceeds.
- 2. Your Shares, profit participating certificates or rights to acquire Shares in us are held by you or deemed to be held by you following the application of a non-recognition provision.
- 3. Your partner or any of your relatives by blood or by marriage in the direct line (including fosterchildren) or of those of your partner has a substantial interest (as described under 1. and 2. above) in us.

If you are entitled to the benefits from Shares or profit participating certificates (for instance if you are a holder of a right of usufruct) you are deemed to be a holder of Shares or profit participating certificates, as the case may be, and your entitlement to benefits is considered a share or profit participating certificate, as the case may be.

If you are an individual and if you satisfy test b., but do not satisfy test c. and/or test d., your Dutch income tax position is not discussed in this Prospectus. If you are an individual who does not satisfy test b., please refer to the section "Dutch Taxation – Taxes on Income and Capital Gains – Non-Resident Holders of Shares". For the purposes of this section you are a "Dutch Corporate Entity" if you satisfy the following tests:

- i. you are a corporate entity (*lichaam*), including an association that is taxable as a corporate entity, that is subject to Dutch corporation tax in respect of benefits derived from its Shares;
- ii. you are resident, or deemed to be resident, in the Netherlands for Dutch corporation tax purposes;
- iii. you are not an entity that, although in principle subject to Dutch corporation tax, is, in whole or in part, specifically exempt from that tax; and
- iv. you are not an investment institution (*beleggingsinstelling*) as defined in article 28 of the Dutch Corporation Tax Act 1969 (*Wet op de vennootschapsbelasting 1969*).

If you are not an individual and if you do not satisfy any one or more of these tests, with the exception of test ii., your Dutch corporation tax position is not discussed in this Prospectus. If you are not an individual that does not satisfy test ii., please refer to the section "Dutch Taxation – Taxes on Income and

# Capital Gains – Non-Resident Holders of Shares". Dutch Individuals Deriving Profits or Deemed To Be Deriving Profits from an Enterprise

If you are a Dutch Individual and if you derive or are deemed to derive any benefits from your Shares, including any capital gain realized on the disposal of such Shares, that are attributable to an enterprise from which you derive profits, whether as an entrepreneur (*ondernemer*) or pursuant to a co-entitlement to the net value of an enterprise, other than as a shareholder, such benefits are generally subject to Dutch income tax at progressive rates.

#### Dutch Individuals Deriving Benefits from Miscellaneous Activities

If you are a Dutch Individual and if you derive or are deemed to derive any benefits from your Shares, including any gain realized on the disposal of such Shares, that constitute benefits from miscellaneous activities (*resultaat uit overige werkzaamheden*), such benefits are generally subject to Dutch income tax at progressive rates.

If you are a Dutch Individual you may, *inter alia*, derive, or be deemed to derive, benefits from Shares that are taxable as benefits from miscellaneous activities in the following circumstances:

- a. if your investment activities go beyond the activities of an active portfolio investor, for instance in the case of use of insider knowledge (*voorkennis*) or comparable forms of special knowledge; or
- b. if any benefits to be derived from your Shares, whether held directly or indirectly, are intended, in whole or in part, as remuneration for activities performed by you or by a person who is a connected person to you as meant by article 3.92b, paragraph 5, of the Dutch Income Tax Act 2001 (Wet inkomstenbelasting 2001).

#### Other Dutch Individuals

If you are a Dutch Individual and your situation has not been discussed before in this section "Dutch Taxation – Taxes on Income and Capital Gains – Resident Holders of Shares", benefits from your Shares are taxed annually as a benefit from savings and investments (*voordeel uit sparen en beleggen*). Such benefit is deemed to be 4 per cent per annum of your "yield basis" (*rendementsgrondslag*), generally to be determined at the beginning of the year, to the extent that such yield basis exceeds the "exempt net asset amount" (*heffingvrij vermogen*) for the relevant year. The benefit is taxed at the rate of 30 per cent. The value of your Shares forms part of your yield basis. Actual benefits derived from your Shares, including any gain realized on the disposal of such Shares, are not as such subject to Dutch income tax.

### Attribution Rule

Benefits derived or deemed to be derived from certain miscellaneous activities by, and yield basis for benefits from savings and investments of, a child or a foster child who is under eighteen years of age are attributed to the parent who exercises, or to the parents who exercise, authority over the child, irrespective of the country of residence of the child.

#### Dutch Corporate Entities

If you are a Dutch Corporate Entity, any benefits derived or deemed to be derived by you from your Shares, including any gain realized on the disposal thereof, are generally subject to Dutch corporation tax, except to the extent that the benefits are exempt under the participation exemption as laid down in the Dutch Corporation Tax Act 1969 (*Wet op de Vennootschapsbelasting 1969*).

## Non-Resident Holders of Shares

The summary set out in this section "Dutch Taxation – Taxes on Income and Capital Gains – Non-Resident Holders of Shares" applies only to a holder of Shares who is a Non-resident holder of Shares.

For the purposes of this section, you are a "Non-resident holder of Shares" if you satisfy the following tests:

- a. you are neither resident, nor deemed to be resident, in the Netherlands for purposes of Dutch income tax or corporation tax, as the case may be, and, if you are an individual, you have not elected to be treated as a resident of the Netherlands for Dutch income tax purposes;
- b. your Shares and any benefits derived or deemed to be derived from such Shares have no connection with your past, present or future employment or membership of a management board (*bestuurder*) or a supervisory board (*commissaris*);
- c. your Shares do not form part of a substantial interest or a deemed substantial interest in us within the meaning of Chapter 4 of the Dutch Income Tax Act 2001 (*Wet inkomstenbelasting 2001*), unless such interest forms part of the assets of an enterprise; and
- d. if you are not an individual, no part of the benefits derived from your Shares is exempt from Dutch corporation tax under the participation exemption as laid down in the Dutch Corporation Tax Act 1969 (*Wet op de Vennootschapsbelasting 1969*).

See the section "Dutch Taxation – Taxes on Income and Capital Gains – Resident Holders of Shares" for a description of the circumstances under which Shares form part of a substantial interest or a deemed substantial interest in us.

If you satisfy test a., but do not satisfy any one or more of tests b., c., and d., your Dutch income tax position or corporation tax position, as the case may be, is not discussed in this Prospectus. If you are a Non-resident holder of Shares you will not be subject to any Dutch taxes on income or capital gains (other than the dividend withholding tax described below) in respect of any benefits derived or deemed to be derived by you from your Shares, including any capital gain realised on the disposal thereof, except if

- you derive profits from an enterprise, directly or pursuant to a co-entitlement to the net value of such enterprise, other than as a shareholder, if you are an individual, or other than as a holder of securities, if you are not an individual such enterprise is either managed in the Netherlands or carried on, in whole or in part, through a permanent establishment or a permanent representative in the Netherlands, and your Shares are attributable to such enterprise; or
- 2. you are an individual and you derive benefits from Shares that are taxable as benefits from miscellaneous activities in the Netherlands.

See the section "Dutch Taxation – Taxes on Income and Capital Gains – Resident Holders of Shares" for a description of the circumstances under which the benefits derived from Shares may be taxable as benefits from miscellaneous activities, on the understanding that such benefits will be taxable in the Netherlands only if such activities are performed or deemed to be performed in the Netherlands.

## Attribution rule

Benefits derived or deemed to be derived from certain miscellaneous activities by a child or a foster child who is under eighteen years of age are attributed to the parent who exercises, or the parents who exercise, authority over the child, irrespective of the country of residence of the child.

### **Dutch Taxation – Dividend Withholding Tax**

#### General

We are generally required to withhold Dutch dividend withholding tax at a rate of 15 per cent. from dividends distributed by us.

The concept "dividends distributed by us' as used in this section "Dutch Taxation – Dividend Withholding Tax" includes, but is not limited to, the following:

- distributions in cash or in kind, deemed and constructive distributions and repayments of capital not recognized as paid-in for Dutch dividend withholding tax purposes;
- liquidation proceeds and proceeds of repurchase or redemption of Shares in excess of the average capital recognized as paid-in for Dutch dividend withholding tax purposes;
- the par value of Shares issued by us to a holder of Shares or an increase of the par value of Shares, as the case may be, to the extent that it does not appear that a contribution, recognized for Dutch dividend withholding tax purposes, has been made or will be made; and
- partial repayment of capital, recognized as paid-in for Dutch dividend withholding tax purposes, if and to the extent that there are net profits (*zuivere winst*), unless (a) the general meeting of our shareholders has resolved in advance to make such repayment and (b) the par value of the Shares concerned has been reduced by an equal amount by way of an amendment to our articles of association.

## **Dutch Individuals and Dutch Corporate Entities**

If you are a Dutch Individual (other than an individual who has elected to be treated as a resident of the Netherlands for Dutch income tax purposes) or a Dutch Corporate Entity, you can generally credit Dutch dividend withholding tax against your Dutch income tax or your Dutch corporation tax liability, as applicable, and you are generally entitled to a refund in the form of a negative assessment of Dutch income tax or Dutch corporation tax, as applicable, insofar as such dividend withholding tax, together with any other creditable domestic and/or foreign taxes, exceeds your aggregate Dutch income tax or your aggregate Dutch corporation tax liability.

Pursuant to domestic rules to avoid dividend stripping, Dutch dividend withholding tax will only be creditable by or refundable if you are the beneficial owner (*uiteindelijk gerechtigde*) of dividends distributed by us. If you receive proceeds from your Shares, you shall not be recognized as the beneficial owner of such proceeds if, in connection with the receipt of the proceeds, you have given a consideration, in the framework of a composite transaction including, without limitation, the mere acquisition of one or more dividend coupons or the creation of short-term rights of enjoyment of Shares (*kortlopende genotsrechten op aandelen*), whereas it may be presumed that (i) such proceeds in whole or in part, directly or indirectly, inure to a person who would not have been entitled to an exemption from, reduction or refund of, or credit for, dividend withholding tax, or who would have been entitled to a smaller reduction or refund of, or credit for, dividend withholding tax than you, the actual recipient of the proceeds; and (ii) such person acquires or retains, directly or indirectly, an interest in Shares or similar instruments, comparable to its interest in Shares prior to the time the composite transaction was first initiated.

If you are an individual who is not resident or deemed to be resident in the Netherlands, but if you have elected to be treated as a resident of the Netherlands for Dutch income tax purposes, you may be eligible for relief from Dutch dividend withholding tax on the same conditions as an individual who is a Non-resident holder of Shares, as discussed below.

See the section "Dutch Taxation – Dividend Withholding Tax – General" for a description of the concept "dividends distributed by us".

See the section "Dutch Taxation – Taxes on Income and Capital Gains – Resident Holders of Shares" for a description of the terms Dutch Individual and Dutch Corporate Entity.

### Non-Resident Holders of Shares

Relief

If you are a Non-resident holder of Shares and if you are resident in the non-European part of the Kingdom of the Netherlands or in a country that has concluded a double taxation treaty with the Netherlands, you may be eligible for a full or partial relief from the dividend withholding tax, provided such relief is timely and duly claimed. Pursuant to domestic rules to avoid dividend stripping, dividend withholding tax relief will only be available to you if you are the beneficial owner of dividends distributed by us. The Dutch tax authorities have taken the position that this beneficial-ownership test can also be applied to deny relief from dividend withholding tax under double tax treaties and the Tax Arrangement for the Kingdom (*Belastingregeling voor het Koninkrijk*) or the Tax Arrangement for the country of the Netherlands (*Belastingregeling voor het land Nederland*).

In addition, if you are a Non-resident holder of Shares that is not an individual, you are entitled to an exemption from dividend withholding tax, provided that the following tests are satisfied:

- 1. you are, according to the tax law of a Member State of the European Union or a state designated by ministerial decree, that is a party to the Agreement regarding the European Economic Area, resident there and you are not transparent for tax purposes according to the tax law of such state;
- any one or more of the following threshold conditions are satisfied:
   at the time the dividend is distributed by us, you hold Shares representing at least five per set.
  - a. at the time the dividend is distributed by us, you hold Shares representing at least five per cent. of our nominal paid up capital; or
  - b. you have held Shares representing at least five per cent. of our nominal paid up capital for a continuous period of more than one year at any time during the four years preceding the time the dividend is distributed by us; or
  - c. you are connected with us within the meaning of article 10a, paragraph 4, of the Dutch Corporation Tax Act 1969 (Wet op de Vennootschapsbelasting 1969); or
  - d. an entity connected with you within the meaning of article 10a, paragraph 4, of the Dutch Corporation Tax Act 1969 (Wet op de Vennootschapsbelasting 1969) holds at the time the dividend is distributed by us, Shares representing at least five per cent. of our nominal paid up capital;
- 3. you are not considered to be resident outside the Member States of the European Union or the states designated by ministerial decree, that are a party to the Agreement regarding the European Economic Area under the terms of a double taxation treaty concluded with a third State; and
- 4. you do not perform a similar function as an investment institution (*beleggingsinstelling*) as meant by article 6a or article 28 of the Dutch Corporation Tax Act 1969 (*Wet op de Vennootschapsbelasting 1969*).

The exemption from dividend withholding tax is not available to you if you are a Non-resident Holder of Shares and if pursuant to a provision for the prevention of fraud or abuse included in a double taxation treaty between the Netherlands and your country of residence, you would not be entitled to the reduction of tax on dividends provided for by such treaty. Furthermore, the exemption from dividend withholding tax will only be available to you if you are the beneficial owner of dividends distributed by us. If you are a Non-resident holder of Shares and you are resident in a Member State of the European Union with which the Netherlands has concluded a double taxation treaty that provides for a reduction of tax on dividends based on the ownership of the number of voting rights, the test under 2.a. above is also satisfied if you own five per cent. of the voting rights in us.

#### Credit

If your are a Non-resident Holder of Shares and if you are subject to Dutch income tax or Dutch corporation tax in respect of any benefits derived or deemed to be derived from your Shares, including any capital gain realized on the disposal thereof, you can generally credit Dutch dividend withholding tax against your Dutch income tax or your Dutch corporation tax liability, as applicable, and you are generally entitled to a refund pursuant to a negative tax assessment if and to the extent the dividend withholding tax, together with any other creditable domestic and/or foreign taxes, exceeds your aggregate Dutch income tax or your aggregate Dutch corporation tax liability, respectively.

See the section "Dutch Taxation – Dividend Withholding Tax – Dutch Individuals and Dutch Corporate Entities" for a description of the term beneficial owner.

See the section "Dutch Taxation – Dividend Withholding Tax – General" for a description of the concept "dividends distributed by us".

See the section "Dutch Taxation – Taxes on Income and Capital Gains – Non-Resident Holders of Shares" for a description of the term Non-resident holder of Shares.

See the section "Dutch Taxation – Taxes on Income and Capital Gains – Non-resident Holders of Shares" for a description of the circumstances under which a Non-resident Holder of Shares is subject to Dutch income tax or Dutch corporation tax.

## **Dutch Taxation – Gift and Inheritance Taxes**

If you dispose of Shares by way of gift, in form or in substance, or if you die, no Dutch gift tax or Dutch inheritance tax, as applicable, will be due, unless:

- (i) you are, or you were, resident or deemed to be resident in the Netherlands for purposes of Dutch gift tax or Dutch inheritance tax, as applicable; or
- (ii) you made a gift of Shares, then became a resident or deemed resident of the Netherlands, and died as a resident or deemed resident of the Netherlands within 180 days of the date of the gift.

For purposes of the above, a gift of Shares made under a condition precedent (*opschortende voorwaarde*) is deemed to be made at the time the condition precedent is satisfied.

### **Dutch Taxation – Other Taxes and Duties**

No Dutch registration tax, transfer tax, stamp duty or any other similar documentary tax or duty, other than court fees, is payable in the Netherlands by you in respect of or in connection with (i) the subscription, issue, placement, allotment, delivery of Shares, (ii) the delivery and/or enforcement by way of legal proceedings (including the enforcement of any foreign judgment in the courts of the Netherlands) of the documents relating to the issue of Shares or the performance by us of our obligations under such documents, or (iii) the transfer of Shares.

## **General Information**

#### **Available Information**

We will be required to publish our annual accounts, accompanied by an annual report and an auditor's certificate, within four months after the end of each financial year and our half-yearly figures within two months after the end of the first six months of each financial year. In addition, we are also obliged to publish interim management statements.

Our Annual Reports (comprising our annual accounts, an annual report and an auditors' certificate) and our half-yearly figures and interim management statements upon their publication can be inspected by our shareholders without charge at our head office in Leiden, during regular business hours from the day of notice convening the annual General Meeting of Shareholders.

Copies of our Annual Reports for the years ended 31 December 2008, 2009 and 2010, our Condensed Consolidated Interim Financial Statements for the six months ended 30 June 2011 (also including the 30 June 2010 figures), our deed of incorporation and our Articles of Association may be obtained free of charge for the life of this Prospectus by sending a request in writing to us at our business address: Zernikedreef 12, 2333 CL Leiden, the Netherlands. These Annual Reports and Condensed Consolidated Interim Financial Statements are also available on www.octoplus.nl for the life of this Prospectus.

This Prospectus will be available to investors on the website of the AFM at <u>www.afm.nl</u> and through the NYSE Euronext website at <u>www.euronext.com</u>. Alternatively, this Prospectus will be available at no cost upon simple request from the Company, by sending an e-mail to: IR@octoplus.nl

#### **Share Trading Information**

The Shares are traded through the book-entry facilities of Euroclear Netherlands, only. The address of Euroclear Netherlands is: Herengracht 459-469, 1017 BS Amsterdam.

The Shares are traded under the following characteristics:

ISIN Code: NL0000345718 Common Code: 026668441 Euronext Amsterdam Symbol: OCTO

### **The Private Placement**

The Private Placement consisted of a private placement to qualified investors in the Netherlands and certain other jurisdictions of 8,000,000 New Shares at a price of  $\leq 0.50$  each, totalling an amount of  $\leq 4.0$  million. The New Shares were offered and sold outside the United States in reliance on Regulation S. As a result of the issue of the 8,000,000 New Shares pursuant to the Private Placement, the number of outstanding shares increased from 36,778,974 to 44,7778,974. Any shareholder of OctoPlus who has not participated in the Private Placement suffered a dilution of 21.8% as a result of the Private Placement. The rights of holders of the New Shares and our existing Shares rank pari passu with each other.

We have applied for admission of 8,000,000 New Shares issued pursuant to the Private Placement to listing and trading on Euronext Amsterdam and expect that trading in these Shares on Euronext Amsterdam will commence on or about the Listing Date.

#### **Paying Agent**

ABN AMRO Bank N.V. is the Paying Agent with respect to our Shares and will act as the Listing Agent with respect to the New Shares. The address of the Paying and Listing Agent is:

ABN AMRO Bank N.V. Listing and Paying Agency Services HQ7050 Gustav Mahlerlaan 10 P.O. Box 283 (HQ7050) 1000 EA Amsterdam The Netherlands

## **Organizational Structure**

We are a holding company of a number of directly held operating companies. Our subsidiaries and holdings are:

Name	Percentage	Country of Incorporation
OctoPlus Development B.V.	100%	The Netherlands
OctoPlus Technologies B.V.	100%	The Netherlands
Chienna B.V.	100%	The Netherlands
OctoPlus Sciences B.V.	100%	The Netherlands
OctoShare B.V.	100%	The Netherlands
OctoPlus PolyActive Sciences B.V.	100%	The Netherlands

## Advisors

Loyens & Loeff N.V. acted as our Dutch counsel in connection with the Private Placement and this Prospectus.

### **Independent Auditors**

Our audited consolidated financial statements as of and for each of the years in the three-year period ended 31 December 2008, 2009 and 2010, incorporated by reference in this Prospectus, have been audited by an auditor of Deloitte Accountants B.V., independent auditors, as stated in their report thereon appearing elsewhere herein. The auditor is a member of the Royal Netherlands Institute of Chartered Accountants (*Koninklijk Nederlands Instituut voor Registeraccountants*).

## **Legal Proceedings**

There are no governmental, legal or arbitration proceedings, including any such proceedings pending or threatened of which we are aware, during a period covering at least the past 12 months which may have, or have had in the recent past, significant effects on our financial position or profitability.

## **Financial and Trading Position**

Since 30 June 2011, the following significant changes to the financial and trading position of OctoPlus have occurred:

 In October 2011, we completed the Private Placement issuing 8,000,000 New Shares and raising € 4.0 million in gross proceeds and € 3.6 million in net proceeds.

## ISSUER

# OctoPlus N.V.

Business Address Zernikedreef 12 2333 CL Leiden The Netherlands

## LEGAL ADVISORS TO THE ISSUER

# Loyens & Loeff N.V.

Fred. Roeskestraat 100 1076 ED Amsterdam The Netherlands

# **INDEPENDENT AUDITORS**

## Deloitte Accountants B.V. Orlyplein 10 1043 DP Amsterdam The Netherlands