




> Return address PO Box 16114, 2500 BC The Hague, the Netherlands

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|--|---|
| <b>MANUFACTURERS AUTHORISATION OF PROXY LABORATORIES B.V., LEIDEN, THE NETHERLANDS, LASTLY CHANGED ON 27 AUGUST 2010</b>   |   |
| <b>1. AUTHORISATION NUMBER</b>   |   |
| 4577 F   |   |
| <b>2. NAME AND SITE OF AUTHORISATION HOLDER</b>  |   |
| Proxy Laboratories B.V., in Leiden   |   |
| <b>3. SITE(S)</b>  |   |
| LEIDEN   | Archimedesweg 25, 2333 CM Leiden                    |
| <b>4. ADDRESS OF AUTHORISATION HOLDER</b>  |   |
| Archimedesweg 25, 2333 CM in Leiden, the Netherlands   |   |
| <b>5. SCOPE OF AUTHORISATION</b>   |   |
| Manufacturing of medicinal products  |   |
| Importation of medicinal products  |   |
| Manufacturing of investigational medicinal products  |   |
| Importation of investigational medicinal products  |   |
| <b>6. LEGAL BASIS OF AUTHORISATION</b>   |   |
| Article 18, subsection 1, of the Dutch Medicines Act [ <i>Geneesmiddelenwet</i> ]  |   |
| <b>7. ANNEXES TO THE AUTHORISATION</b>   |   |
| Annex 1, part 1:   | Manufacturing of medicinal products                 |
| Annex 1, part 2:   | Importation of medicinal products                   |
| Annex 2, part 1:   | Manufacturing of investigational medicinal products |
| Annex 2, part 2:   | Importation of investigational medicinal products   |
| Annex 3:   | QP's  |
| <b>8. SIGNATURE</b>  |   |
| The Minister for Health, Welfare and Sport,<br>On his behalf<br>Unit Farmatec-BMC<br>was signed<br><br>Ms A.J. Hennis<br>Acting Unit Head |   |





## SCOPE OF AUTHORISATION

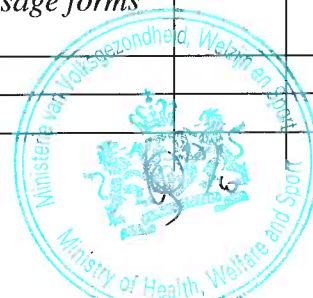
Address of the site: Archimedesweg 25 at LEIDEN

|   |   |
|---|---|
| X | <b>HUMAN MEDICINAL PRODUCTS</b>           |
| X | <b>INVESTIGATIONAL MEDICINAL PRODUCTS</b> |

|                              |  |
|------------------------------|--|
| <b>AUTHORISED OPERATIONS</b> |  |
| X                            | <b>MANUFACTURING OPERATIONS (ACCORDING TO PART 1)</b>          |
| X                            | <b>IMPORTATION OF MEDICINAL PRODUCTS (ACCORDING TO PART 2)</b> |

|   |  |
|---|--|
| <b>PART 1 – MANUFACTURING OPERATIONS</b>  |  |
| <p>- authorised manufacturing operations include total and partial manufacturing (including various processes of dividing up, packaging and presentation), batch release and certification, importation, storage and distribution of specified dosage forms unless informed to the contrary;</p> <p>- quality control testing and/or release and batch certification activities without manufacturing operations should be specified under the relevant items;</p> <p>- if the company is engaged in the manufacture of products with special requirements, e.g. radiopharmaceuticals or products containing penicillin, sulphonamides, cytotoxics, cephalosporins, substances with hormonal activity or other potentially hazardous active ingredients, this should be stated under the relevant product type and dosage form.</p> |  |

|     |   | Yes | No |
|-----|---|-----|----|
| 1.1 | <b>STERILE PRODUCTS</b>   |     | X  |
|     | <i>1.1.1 Aseptically prepared (list of dosage forms)</i>  |     | X  |
|     | 1.1.1.1 Large volume liquids  |     | X  |
|     | 1.1.1.2 Lyophilisates   |     | X  |
|     | 1.1.1.3 Semi-solids   |     | X  |
|     | 1.1.1.4 Small volume liquids  |     | X  |
|     | 1.1.1.5 Solids and implants   |     | X  |
|     | 1.1.1.6 Other aseptically prepared products   |     | X  |
|     | <i>1.1.2 Terminally sterilised (list of dosage forms)</i>   |     | X  |
|     | 1.1.2.1 Large volume liquids  |     | X  |
|     | 1.1.2.2 Semi-solids   |     | X  |
|     | 1.1.2.3 Small volume liquids  |     | X  |
|     | 1.1.2.4 Solids and implants   |     | X  |
|     | 1.1.2.5 Other terminally sterilised products, namely:   |     | X  |
|     | <i>1.1.3 Batch certification only (in case you would have dosage forms in this product group contract manufactured)</i> |     | X  |
| 1.2 | <b>NON-STERILE PRODUCTS</b>   |     | X  |
|     | <i>1.2.1 Non-sterile products (list of dosage forms)</i>  |     | X  |
|     | 1.2.1.1 Capsules, hard shell  |     | X  |







|     |   |   |   |
|-----|---|---|---|
|     | <i>1.4.1 Manufacture of:</i>  |   | X |
|     | 1.4.1.1 Herbal medicinal products   |   | X |
|     | 1.4.1.2 Homeopathic medicinal products                                      |   | X |
|     | 1.4.1.3 Biological active substances  |   | X |
|     | 1.4.1.4 Other   |   | X |
|     | <i>1.4.2 Sterilisation of active substances/excipients/finished product</i> |   | X |
|     | 1.4.2.1 Filtration  |   | X |
|     | 1.4.2.2 Dry heat  |   | X |
|     | 1.4.2.3 Moist heat  |   | X |
|     | 1.4.2.4 Chemical  |   | X |
|     | 1.4.2.5 Gamma irradiation   |   | X |
|     | 1.4.2.6 Electron beam   |   | X |
|     | <i>1.4.3 Other</i>  |   | X |
| 1.5 | <b>PACKAGING ONLY</b>   |   | X |
|     | <i>1.5.1 Primary packing</i>  |   | X |
|     | 1.5.1.1 Capsules, hard shell  |   | X |
|     | 1.5.1.2 Capsules, soft shell  |   | X |
|     | 1.5.1.3 Chewing gums  |   | X |
|     | 1.5.1.4 Impregnated matrices  |   | X |
|     | 1.5.1.5 Liquids for external use  |   | X |
|     | 1.5.1.6 Liquids for internal use  |   | X |
|     | 1.5.1.7 Medicinal gases   |   | X |
|     | 1.5.1.8 Other solid dosage forms  |   | X |
|     | 1.5.1.9 Pressurised preparations  |   | X |
|     | 1.5.1.10 Radionuclide generators  |   | X |
|     | 1.5.1.11 Semi-solids  |   | X |
|     | 1.5.1.12 Suppositories  |   | X |
|     | 1.5.1.13 Tablets  |   | X |
|     | 1.5.1.14 Transdermal patches  |   | X |
|     | 1.5.1.15 Intraruminal devices   |   | X |
|     | 1.5.1.16 Veterinary premixes  |   | X |
|     | 1.5.1.17 Other non-sterile medicinal products, namely:                      |   | X |
|     | <i>1.5.2 Secondary packing</i>  |   | X |
| 1.6 | <b>QUALITY CONTROL TESTING</b>  | X |   |
|     | <i>1.6.1 Microbiological: sterility</i>                                     | X |   |
|     | <i>1.6.2 Microbiological: non-sterility</i>                                 | X |   |
|     | <i>1.6.3 Chemical/Physical</i>  | X |   |





|                  |   |  |
|------------------|---|--|
| 1.6.4 Biological | X |  |
|------------------|---|--|

**Any restrictions or clarifying remarks related to the scope of these manufacturing operations**





|   |  |
|---|--|
| <b>PART 2 – IMPORTATION OF MEDICINAL PRODUCTS</b>   |  |
| - authorised importation activities without manufacturing activity;                                   |  |
| - authorised importation activities include storage and distribution unless informed to the contrary. |  |

|     |  | Yes | No |
|-----|--|-----|----|
| 2.1 | <b>QUALITY CONTROL TESTING OF IMPORTED MEDICINAL PRODUCTS</b>  | X   |    |
|     | <i>2.1.1 Microbiological: sterility</i>                        | X   |    |
|     | <i>2.1.2 Microbiological: non-sterility</i>                    | X   |    |
|     | <i>2.1.3 Chemical/Physical</i>                                 | X   |    |
|     | <i>2.1.4 Biological</i>  | X   |    |
| 2.2 | <b>BATCH CERTIFICATION OF IMPORTED MEDICINAL PRODUCTS</b>      |     | X  |
|     | <i>2.2.1 Sterile products</i>                                  |     | X  |
|     | 2.2.1.1 Aseptically prepared                                   |     | X  |
|     | 2.2.1.2 Terminally sterilised                                  |     | X  |
|     | <i>2.2.2 Non-sterile products</i>                              |     | X  |
|     | <i>2.2.3 Biological medicinal products</i>                     |     | X  |
|     | 2.2.3.1 Blood products   |     | X  |
|     | 2.2.3.2 Immunological products                                 |     | X  |
|     | 2.2.3.3 Cell therapy products                                  |     | X  |
|     | 2.2.3.4 Gene therapy products                                  |     | X  |
|     | 2.2.3.5 Biotechnology products                                 |     | X  |
|     | 2.2.3.6 Human or animal extracted products                     |     | X  |
|     | 2.2.3.7 Other biological medicinal products                    |     | X  |
|     | <i>2.2.4 Other importation activities (any other relevant)</i> |     | X  |
|     | 2.2.4.1 Radiopharmaceuticals                                   |     | X  |
|     | 2.2.4.2 Medicinal gases  |     | X  |
|     | 2.2.4.3 Herbal products  |     | X  |
|     | 2.2.4.4 Homeopathic products                                   |     | X  |
|     | 2.2.4.5 Biological active starting materials                   |     | X  |
|     | 2.2.4.6 Other  |     | X  |

**Any restrictions or clarifying remarks related to the scope of these manufacturing operations**





**PART 1 – MANUFACTURING OPERATIONS RELATING TO INVESTIGATIONAL MEDICINAL PRODUCTS**

- authorised manufacturing operations include total and partial manufacturing (including various processes of dividing up, packaging and presentation), batch release and certification, importation, storage and distribution of specified dosage forms unless informed to the contrary;
- quality control testing and/or release and batch certification activities without manufacturing operations should be specified under the relevant items;
- if the company is engaged in the manufacture of products with special requirements, e.g. radiopharmaceuticals or products containing penicillin, sulphonamides, cytotoxics, cephalosporins, substances with hormonal activity or other potentially hazardous active ingredients, this should be stated under the relevant product type and dosage form.

|     |   | Yes | No |
|-----|---|-----|----|
| 1.1 | <b>STERILE INVESTIGATIONAL MEDICINAL PRODUCTS</b>   | X   |    |
|     | <i>1.1.1 Aseptically prepared (list of dosage forms)</i>  | X   |    |
|     | 1.1.1.1 Large volume liquids  |     | X  |
|     | 1.1.1.2 Lyophilisates   |     | X  |
|     | 1.1.1.3 Semi-solids   |     | X  |
|     | 1.1.1.4 Small volume liquids  | X   |    |
|     | 1.1.1.5 Solids and implants   |     | X  |
|     | 1.1.1.6 Other aseptically prepared products   |     | X  |
|     | <i>1.1.2 Terminally sterilised (list of dosage forms)</i>   |     | X  |
|     | 1.1.2.1 Large volume liquids  |     | X  |
|     | 1.1.2.2 Semi-solids   |     | X  |
|     | 1.1.2.3 Small volume liquids  |     | X  |
|     | 1.1.2.4 Solids and implants   |     | X  |
|     | 1.1.2.5 Other terminally sterilised prepared products   |     | X  |
|     | <i>1.1.3 Batch certification only (in case you would have dosage forms in this product group contract manufactured)</i> |     | X  |
| 1.2 | <b>NON-STERILE INVESTIGATIONAL MEDICINAL PRODUCTS</b>   | X   |    |
|     | <i>1.2.1 Non-sterile products (list of dosage forms)</i>  | X   |    |
|     | 1.2.1.1 Capsules, hard shell  | X   |    |
|     | 1.2.1.2 Capsules, soft shell  |     | X  |
|     | 1.2.1.3 Chewing gums  |     | X  |
|     | 1.2.1.4 Impregnated matrices  |     | X  |
|     | 1.2.1.5 Liquids for external use  |     | X  |
|     | 1.2.1.6 Liquids for internal use  |     | X  |
|     | 1.2.1.7 Medicinal gases   |     | X  |
|     | 1.2.1.8 Other solid dosage forms  |     | X  |
|     | 1.2.1.9 Pressurised preparations  |     | X  |





|     |  |   |   |
|-----|--|---|---|
|     | 1.2.1.10 Radionuclide generators   |   | X |
|     | 1.2.1.11 Semi-solids   |   | X |
|     | 1.2.1.12 Suppositories   |   | X |
|     | 1.2.1.13 Tablets   |   | X |
|     | 1.2.1.14 Transdermal patches   |   | X |
|     | 1.2.1.15 Other non-sterile medicinal products:   |   | X |
|     | <i>1.2.2 Batch certification only (in case you would have dosage forms in this product group contract manufactured)</i>                        |   | X |
| 1.3 | <b>BIOLOGICAL INVESTIGATIONAL MEDICINAL PRODUCTS</b>   | X |   |
|     | <i>1.3.1 Biological medicinal products</i>   | X |   |
|     | 1.3.1.1 Blood products   |   | X |
|     | 1.3.1.2 Immunological products   |   | X |
|     | 1.3.1.3 Cell therapy products  |   | X |
|     | 1.3.1.4 Gene therapy products  |   | X |
|     | 1.3.1.5 Biotechnology products   | X |   |
|     | 1.3.1.6 Human or animal extracted products   |   | X |
|     | 1.3.1.7 Other biological medicinal products  |   | X |
|     | <i>1.3.2 Batch certification only (in case you would have dosage forms in this product group contract manufactured; list of product types)</i> |   | X |
|     | 1.3.2.1 Blood products   |   | X |
|     | 1.3.2.2 Immunological products   |   | X |
|     | 1.3.2.3 Cell therapy products  |   | X |
|     | 1.3.2.4 Gene therapy products  |   | X |
|     | 1.3.2.5 Biotechnology products   |   | X |
|     | 1.3.2.6 Human or animal extracted products   |   | X |
|     | 1.3.2.7 Other biological medicinal products  |   | X |
| 1.4 | <b>OTHER INVESTIGATIONAL MEDICINAL PRODUCTS OR MANUFACTURING ACTIVITY</b>  | X |   |
|     | <i>1.4.1 Manufacture of:</i>   |   | X |
|     | 1.4.1.1 Herbal products  |   | X |
|     | 1.4.1.2 Homeopathic products   |   | X |
|     | 1.4.1.3 Biological active starting materials   |   | X |
|     | 1.4.1.4 Other  |   | X |
|     | <i>1.4.2 Sterilisation of active substances/excipients/finished product</i>  | X |   |
|     | 1.4.2.1 Filtration   | X |   |
|     | 1.4.2.2 Dry heat   |   | X |
|     | 1.4.2.3 Moist heat   |   | X |
|     | 1.4.2.4 Chemical   |   | X |
|     | 1.4.2.5 Gamma irradiation  |   | X |
|     | 1.4.2.6 Electron beam  |   | X |





|     |   |   |   |
|-----|---|---|---|
|     | <i>1.4.3 Other:</i>                           |   | X |
| 1.5 | PACKAGING ONLY                                |   | X |
|     | <i>1.5.1 Primary packing</i>                  |   | X |
|     | 1.5.1.1 Capsules, hard shell                  |   | X |
|     | 1.5.1.2 Capsules, soft shell                  |   | X |
|     | 1.5.1.3 Chewing gums                          |   | X |
|     | 1.5.1.4 Impregnated matrices                  |   | X |
|     | 1.5.1.5 Liquids for external use              |   | X |
|     | 1.5.1.6 Liquids for internal use              |   | X |
|     | 1.5.1.7 Medicinal gases                       |   | X |
|     | 1.5.1.8 Other solid dosage forms              |   | X |
|     | 1.5.1.9 Pressurised preparations              |   | X |
|     | 1.5.1.10 Radionuclide generators              |   | X |
|     | 1.5.1.11 Semi-solids                          |   | X |
|     | 1.5.1.12 Suppositories                        |   | X |
|     | 1.5.1.13 Tablets                              |   | X |
|     | 1.5.1.14 Transdermal patches                  |   | X |
|     | 1.5.1.15 Other non-sterile medicinal products |   | X |
|     | 1.5.1.16 Veterinary premixes                  |   | X |
|     | 1.5.1.17 Other non-sterile medicinal products |   | X |
|     | <i>1.5.2 Secondary packing</i>                |   | X |
| 1.6 | QUALITY CONTROL TESTING                       | X |   |
|     | <i>1.6.1 Microbiological: sterility</i>       | X |   |
|     | <i>1.6.2 Microbiological: non-sterility</i>   | X |   |
|     | <i>1.6.3 Chemical/Physical</i>                | X |   |
|     | <i>1.6.4 Biological</i>                       | X |   |

**Any restrictions or clarifying remarks related to the scope of these manufacturing operations**





**PART 2 – IMPORTATION OF INVESTIGATIONAL MEDICINAL PRODUCTS**

- authorised importation activities without manufacturing activity;  
- authorised importation activities include storage and distribution unless informed to the contrary.

|     |  | Yes | No |
|-----|--|-----|----|
| 2.1 | <b>QUALITY CONTROL TESTING OF IMPORTED MEDICINAL PRODUCTS</b>  | X   |    |
|     | <i>2.1.1 Microbiological: sterility</i>                        | X   |    |
|     | <i>2.1.2 Microbiological: non-sterility</i>                    | X   |    |
|     | <i>2.1.3 Chemical/Physical</i>                                 | X   |    |
|     | <i>2.1.4 Biological</i>  | X   |    |
| 2.2 | <b>BATCH CERTIFICATION OF IMPORTED MEDICINAL PRODUCTS</b>      |     | X  |
|     | <i>2.2.1 Sterile products</i>                                  |     | X  |
|     | 2.2.1.1 Aseptically prepared                                   |     | X  |
|     | 2.2.1.2 Terminally sterilised                                  |     | X  |
|     | <i>2.2.2 Non-sterile products</i>                              |     | X  |
|     | <i>2.2.3 Biological medicinal products</i>                     |     | X  |
|     | 2.2.3.1 Blood products   |     | X  |
|     | 2.2.3.2 Immunological products                                 |     | X  |
|     | 2.2.3.3 Cell therapy products                                  |     | X  |
|     | 2.2.3.4 Gene therapy products                                  |     | X  |
|     | 2.2.3.5 Biotechnology products                                 |     | X  |
|     | 2.2.3.6 Human or animal extracted products                     |     | X  |
|     | 2.2.3.7 Other biological medicinal products                    |     | X  |
|     | <i>2.2.4 Other importation activities (any other relevant)</i> |     | X  |
|     | 2.2.4.1 Radiopharmaceuticals                                   |     | X  |
|     | 2.2.4.2 Medicinal gases  |     | X  |
|     | 2.2.4.3 Herbal products  |     | X  |
|     | 2.2.4.4 Homeopathic products                                   |     | X  |
|     | 2.2.4.5 Biological active starting materials                   |     | X  |
|     | 2.2.4.6 Other  |     | X  |

**Any restrictions or clarifying remarks related to the scope of these manufacturing operations**





|                                       |
|---------------------------------------|
| ANNEX 3: QUALIFIED PERSON(S) (QP('S)) |
| Mr. M.C. Kooij                        |
| Mr. Dr. R.E. Santing                  |

