REPUBLIC OF ESTONIA
AGENCY OF MEDICINES

Ravimiamet
Estonian State Agency of Medicines

CERTIFICATE NUMBER: IN-2-14/19/5 (API)

CERTIFICATE OF GMP COMPLIANCE OF A MANUFACTURER [2]

Part 1

Issued following an inspection in accordance with :
Art. 111(5) of Directive 2001/83/EC as amended
Art. 80(5) of Directive 2001/82/EC as amended

The competent authority of Estonia confirms the following:

The manufacturer ***TBD-Biodiscovery (osaühing TBD-Biodiscovery)***

Site address ***Tiigi 61b, Tartu, 50410, Estonia***

Is an active substance manufacturer that has been inspected in accordance with Art. 111(1) of Directive 2001/83/EC and Art. 80(1) of Directive 2001/82/EC, transposed in the following national legislation: National legislation – Medicinal Products Act requires ManA for manufacture of API in Estonia. § 16 subsection (3)

From the knowledge gained during inspection of this manufacturer, the latest of which was conducted on ***2016-05-23*** *(inspection end date),* it is considered that it complies with:

- The principles of GMP for active substances *(3)* referred to in Article 47 of Directive 2001/83/EC and Article 51 of Directive 2001/82/EC

This certificate reflects the status of the manufacturing site at the time of the inspection noted above and should not be relied upon to reflect the compliance status if more than three years have elapsed since the date of that inspection.
This certificate is valid only when presented with all pages and both Parts 1 and 2.

The authenticity of this certificate may be verified with the issuing authority or in EudraGMDP http://eudragmdp.ema.europa.eu. If it does not appear, please contact the issuing authority.

(2) Guidance on the interpretation of this template can be found in the Help menu of EudraGMDP database.
(3) These requirements fulfil the GMP recommendations of WHO.

Part 2

Manufacture of active substance. Names of substances subject to inspection :

RIMANTADINE HYDROCHLORIDE (en)
ANTAZOLINE PHOSPHATE (en)
MAROPITANT (en)
VARENICLINE CITRATE (en)
POMALIDOMIDE (en)
NON-STERILE ACTIVE SUBSTANCES BY CHEMICAL SYNTHESIS (en)

3. MANUFACTURING OPERATIONS - ACTIVE SUBSTANCES

Active Substance : ***RIMANTADINE HYDROCHLORIDE***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Crystallisation
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Drying 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Active Substance : ***ANTAZOLINE PHOSPHATE***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Crystallisation
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Drying 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Active Substance : ***MAROPITANT***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Crystallisation
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Drying 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Active Substance : ***VARENICLINE CITRATE***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Salt formation. Crystallisation
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Drying. De-agglomeration 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Active Substance : ***POMALIDOMIDE***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Crystallisation
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Washing. Drying. 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Active Substance : ***NON-STERILE ACTIVE SUBSTANCES BY CHEMICAL SYNTHESIS***	
3.1	**Manufacture of Active Substance by Chemical Synthesis**
	3.1.1 Manufacture of active substance intermediates 3.1.2 Manufacture of crude active substance 3.1.3 Salt formation / Purification steps : Salt formation. Crystallisation. Chromatographic purification
3.5	**General Finishing Steps**
	3.5.1 Physical processing steps : Drying. De-agglomeration. Milling. Sieving 3.5.2 Primary Packaging (enclosing / sealing the active substance within a packaging material which is in direct contact with the substance) 3.5.3 Secondary Packaging (placing the sealed primary package within an outer packaging material or container. This also includes any labelling of the material which could be used for identification or traceability (lot numbering) of the active substance)
3.6	**Quality Control Testing**
	3.6.1 Physical / Chemical testing

Any restrictions related to the scope of this certificate:
Clarifying remarks:

NON-STERILE ACTIVE SUBSTANCES BY CHEMICAL SYNTHESIS - Small-scale production of non-sterile active substances by chemical synthesis (e.g. for investigational medicinal products)

2019-08-15

Name and signature of the authorised pers
of the competent authority of Estonia

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