LÄKEMEDELSVERKET
MEDICAL PRODUCTS AGENCY

Public Assessment Report
Scientific discussion

Parapo,
500 mg and 1 g, film-coated tablet
(paracetamol)

Asp no: 2016-1686, 2016-1687

This module reflects the scientific discussion for the approval of Parapo. The procedure was finalised on 2017-12-12. For information on changes after this date please refer to the module 'Update'.

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I. INTRODUCTION

The application for Parapo 500 mg and 1 g, film-coated tablet is a generic application made according to Article 10(1) of Directive 2001/83/EC. The applicant, Evolan Pharma AB applied for marketing authorisations in Sweden through the National Procedure.

The reference medicinal product chosen for the purposes of establishing the expiry of the data protection period is Panodil, 500 mg, film-coated tablet authorised in Sweden since 1958, with Omega Pharma Nordic AB as marketing authorisation holder.

For approved indications, see the Summary of Product Characteristics.

For recommendations to the marketing authorisation not falling under Article 21a/22 of Directive 2001/83 and conditions to the marketing authorisation pursuant to Article 21a or 22 of Directive 2001/83/EC to the marketing authorisation, please see section VI.

II. QUALITY ASPECTS

II.1 Drug Substance

The structure of the drug substance has been adequately proven and its physico-chemical properties are sufficiently described.

The manufacture of the drug substance has been adequately described and satisfactory specifications have been provided for starting materials, reagents and solvents.

The drug substance specification includes relevant tests and the limits for impurities and degradation products have been justified. The analytical methods applied are suitably described and validated.

Stability studies confirm the retest period.

II.2 Medicinal Product

The medicinal product is formulated using excipients listed in section 6.1 in the Summary of Product Characteristics.

The manufacturing process has been sufficiently described and critical steps identified.

The tests and limits in the specification are considered appropriate to control the quality of the finished product in relation to its intended purpose.

Stability studies have been performed and data presented support the shelf life and special precautions for storage claimed in the Summary of Product Characteristics, sections 6.3 and 6.4.

III. NON-CLINICAL ASPECTS

III.1 Discussion on the non-clinical aspects

Since this product has been shown to be essentially similar and refer to a product approved based on a full application with regard to preclinical data, no further such data have been submitted or are considered necessary.

IV. CLINICAL ASPECTS

IV.1 Pharmacokinetics

No bioequivalence study has been submitted since the applicant applies for a BCS-based biowaiver.

From a pharmacokinetic point of view, a BCS class I based biowaiver is acceptable for an immediate release drug product if the drug substance has proven to exhibit complete absorption and the excipients that might affect bioavailability are qualitatively and quantitatively the same. It should also be confirmed that the substance is not a narrow therapeutic index drug.

Paracetamol has a fraction absorbed greater or equal to 85% and is classified as a BCS class I substance. For a BCS class I substance the differences regarding the excipients between the test and reference product are acceptable since it is not likely that these differences would affect the bioavailability. In addition paracetamol is not considered to be a narrow therapeutic index drug. Hence, the justification for a BCS class I based biowaiver can be accepted from a clinical point of view and no bioequivalence study is considered necessary.

IV.2 Discussion on the clinical aspects

Since this product has been shown to be essentially similar and refer to a product approved based on a full application with regard to clinical efficacy/safety data, no further such data have been submitted or are considered necessary.

IV.3 Risk Management Plan

The MAH has submitted a risk management plan, in accordance with the requirements of Directive 2001/83/EC as amended, describing the pharmacovigilance activities and interventions designed to identify, characterise, prevent or minimise risks relating to Parapo.

Safety specification

Summary table of safety concerns as approved in RMP

Important identified risks	Hepatotoxicity in overdose Severe skin reactions
Important potential risks	None
Missing information	None

Pharmacovigilance Plan

Routine pharmacovigilance is suggested and no additional pharmacovigilance activities are proposed by the applicant, which is endorsed.

Risk minimisation measures

Routine risk minimisation is suggested and no additional risk minimisation activities are proposed by the applicant, which is endorsed.

Summary of the RMP

The RMP is approved

The MAH shall perform the required pharmacovigilance activities and interventions detailed in the agreed RMP presented in Module 1.8.2 of the Marketing Authorisation and any agreed subsequent updates of the RMP.

An updated RMP should be submitted:

- At the request of the RMS;
- Whenever the risk management system is modified, especially as the result of new information being received that may lead to a significant change to the benefit/risk profile or as the result of an important (pharmacovigilance or risk minimisation) milestone being reached.

If the dates for submission of a PSUR and the update of a RMP coincide, they can be submitted at the same time, but via different procedures.

V. USER CONSULTATION

A user consultation with target patient groups on the package information leaflet (PIL) has been performed on the basis of a bridging report making reference to Paracetamol Apofri 500 mg film coated tablets (Parent PL) which was assessed and approved 2012-11-08 (Dnr 111:2012/48028). The bridging report submitted by the applicant has been found acceptable.

VI. OVERALL CONCLUSION, BENEFIT/RISK ASSESSMENT AND RECOMMENDATION

The quality of the generic product Parapo, is found adequate. There are no objections to the approval of Parapo, from a non-clinical and clinical point of view. The product information is acceptable.
The application is therefore recommended for approval.

List of recommendations not falling under Article 21a/22 of Directive 2001/83 in case of a positive benefit risk assessment

N/A

List of conditions pursuant to Article 21a or 22 of Directive 2001/83/EC

N/A

VII. APPROVAL

Parapo, 500 mg and 1 g, film-coated tablet was approved in the national procedure on 2017-12-12.

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Public Assessment Report – Update

Procedure number*	Scope	Product Information affected	Date of end of procedure	Approval/ non approval	Summary/ Justification for refuse

*Only procedure qualifier, chronological number and grouping qualifier (when applicable)

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