EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

18 August 2011
EMA/533485/2011
Committee for Medicinal Products for Human Use (CHMP)

Assessment report

Entacapone Orion

entacapone

Procedure No. **EMEA/H/C/002440**

Assessment report as adopted by the CHMP with all information of a commercially confidential nature deleted.

7 Westferry Circus ● Canary Wharf ● London E14 4HB ● United Kingdom
Telephone +44 (0)20 7418 8400 **Facsimile** +44 (0)20 7418 8613
E-mail info@ema.europa.eu **Website** www.ema.europa.eu
An agency of the European Union

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List of abbreviations

CHMP	Committee for Medicinal Products for Human Use
EC	European Commission
EMA	European Medicines Agency
EPAR	European Public Assessment Report
ERA	Environmental Risk Assessment
HDPE	High Density Poly Ethylene
MAH	Marketing Authorisation Holder
PSUR	Periodic Safety Update Report

1. Background information on the procedure

1.1. Submission of the dossier

The applicant Orion Corporation submitted on 7 April 2011 an application for Marketing Authorisation to the European Medicines Agency (EMA) for Entacapone Orion, through the centralised procedure under Article 3 (2)(a) of Regulation (EC) No 726/2004. The eligibility to the centralised procedure was agreed upon by the EMA/CHMP on 26 October 2010.

The applicant applied for the following indication:

"Entacapone is indicated as an adjunct to standard preparations of levodopa/benserazide or levodopa/carbidopa for use in adult patients with Parkinson's disease and end-of-dose motor fluctuations, who cannot be stabilised on those combinations."

The legal basis for this application refers to:

Article 10(c) of Directive 2001/83/EC – informed consent application.

The application submitted is composed of administrative information, quality, non-clinical and clinical data with a letter from Orion Corporation allowing use to be made of relevant quality, non-clinical and/or clinical data.

This application is submitted in accordance with Article 82.1 of Regulation (EC) No 726/2004 as a multiple of Comtess authorised on 16 September 1998.

Information on Paediatric requirements

Not applicable.

Information relating to orphan market exclusivity

Similarity

Not applicable.

Market Exclusivity

Not applicable.

Scientific Advice

The applicant did not seek scientific advice at the CHMP.

Licensing status

The initial product Comtess has been given a Community Marketing Authorisation on 16 September 1998.

1.2. Steps taken for the assessment of the product

The Rapporteur and Co-Rapporteur appointed by the CHMP were:

Rapporteur: **Jaana Kallio**

Co-Rapporteur: **Jacqueline Genoux-Hames**

- The application was received by the EMA on 7 April 2011.
- The procedure started on 24 April 2011.
- The Rapporteur's first Assessment Report was circulated to all CHMP members on 30 May 2011. The Co-Rapporteur's first Assessment Report was circulated to all CHMP members on 27 May 2011.
- The Rapporteurs circulated the Joint Assessment Report to all CHMP members on 15 June 2011.
- During the meeting on 20-23 June 2011, the CHMP, in the light of the overall data submitted and the scientific discussion within the Committee, issued a positive opinion for granting a Marketing Authorisation to Entacapone Orion on 23 June 2011.

2. Scientific discussion

2.1. Introduction

This marketing authorisation application for Entacapone Orion (entacapone) has been submitted by Orion Corporation as an informed consent application in accordance with Article 10c of Directive 2001/83/EC, as amended.

The MAH (Orion Corporation) for Comtess, which was authorised on 16 September 1998, provided consent to make use of the pharmaceutical, non-clinical and clinical documentation of the initial dossier of this authorised product and any subsequent post-marketing procedures submitted, assessed and approved.

As a consequence, quality, safety and efficacy of the Entacapone Orion medicinal product are identical to the up-to-date quality, non-clinical and clinical profile of Comtess. The application for Entacapone Orion concerns the identical strengths and pack sizes to those approved for Comtess and consists of only Module 1. Information on the scientific discussion can be found in the Comtess CHMP assessment reports and in the European Public Assessment Report (EPAR) published on the EMA website.

The approved indication is:

“Entacapone is indicated as an adjunct to standard preparations of levodopa/benserazide or levodopa/carbidopa for use in adult patients with Parkinson’s disease and end-of-dose motor fluctuations, who cannot be stabilised on those combinations.”

2.2. Quality aspects

Since this application is an informed consent of the Comtess application, the quality data in support of the Entacapone Orion application are identical to the up-to-date quality data of the Comtess dossier, which have been assessed and approved (including all post-marketing procedures).

2.3. Non-clinical aspects

Since this application is an informed consent of the Comtess application, the non-clinical data in support of the Entacapone Orion application are identical to the up-to-date non-clinical data of the Comtess dossier, which have been assessed and approved (including all post-marketing procedures).

The ERA provided for this application consists of an adequate justification for the absence of specific study data. The medicinal product subject to this application is intended to be administered at comparable dose levels and for indications that are already approved in the European Community for Comtess. Based on the assumption that the product is intended to substitute for identical products on

the market, the approval of the referred product should not result in an increase of the total quantity of the active ingredients released in to the environment. Therefore, it should not result in an increase of risk to the environment during storage, distribution, use and disposal.

2.4. Clinical aspects

Since this application is an informed consent of the Comtess application, the clinical data in support of the Entacapone Orion application are identical to the up-to-date clinical data of the Comtess dossier, which have been assessed and approved (including all post-marketing procedures).

2.5. Pharmacovigilance

Detailed description of the pharmacovigilance system

The CHMP considered that the Pharmacovigilance system as described by the applicant fulfils the legislative requirements.

Risk Management Plan

The CHMP did not require the applicant to submit a risk management plan because the safety profile of the medicinal product subject to this application is well established, and no safety concerns requiring risk minimisation activities have been identified for Comtess.

The CHMP, having considered the data submitted, was of the opinion that routine pharmacovigilance was adequate to monitor the safety of the product.

2.6. User consultation

A justification for not performing a full user consultation with target patient groups on the package leaflet has been submitted by the applicant and has been found acceptable for the following reasons: The Package Leaflet of Comtess has been successfully user tested in the framework of variation EMEA/H/C/0170/II/0043/G for which the CHMP opinion was adopted on 22 April 2010. Since the proposed Package Leaflet for the current application is identical to the Package Leaflet for Comtess except for the product-specific information, no further testing is warranted.

3. Benefit–Risk Balance

Since this application has been submitted by Orion Corporation as an informed consent application to Comtess in accordance with Article 10c of Directive 2001/83/EC, as amended, the CHMP considered that the benefit-risk balance of Entacapone Orion 200 mg film-coated tablets was favourable and therefore recommended the granting of the marketing authorisation for the following indication:

“Entacapone is indicated as an adjunct to standard preparations of levodopa/benserazide or levodopa/carbidopa for use in adult patients with Parkinson’s disease and end-of-dose motor fluctuations, who cannot be stabilised on those combinations.”

4. Recommendations

Outcome

Based on the CHMP review of data on quality, safety and efficacy, the CHMP considered by consensus that the risk-benefit balance of Entacapone Orion in the treatment of Parkinson’s diseases is favourable

and therefore recommends the granting of the marketing authorisation subject to the following conditions:

Conditions or restrictions regarding supply and use

Medicinal product subject to medical prescription.

Conditions and requirements of the Marketing Authorisation

Risk Management System and PSUR cycle

The MAH must ensure that the system of pharmacovigilance, presented in Module 1.8.1 of the marketing authorisation, is in place and functioning before and whilst the medicinal product is on the market.

The PSUR submission schedule should follow the PSUR submission schedule for Comtess.

Conditions or restrictions with regard to the safe and effective use of the medicinal product

Not applicable.

Conditions or restrictions with regard to the safe and effective use of the medicinal product to be implemented by the Member States.

Not applicable.