

Public Assessment Report

Scientific discussion

Rasagiline ratio (rasagiline mesilate)

SE/H/1529/01/DC

This module reflects the scientific discussion for the approval of Rasagiline ratio. The procedure was finalised on 2015-11-18. For information on changes after this date please refer to the module 'Update'.

I. INTRODUCTION

The application for Rasagiline ratio, 1 mg, tablet is a generic application made according to Article 10(1) of Directive 2001/83/EC. The applicant, Teva Sweden AB applies through the Decentralised Procedure with Sweden acting as reference member state (RMS) and ES as concerned member state (CMS).

The reference medicinal product chosen for the purposes of establishing the expiry of the data protection period is Azilect, 1mg, tablet authorised in Union since 2005, with Teva Pharma GmbH as marketing authorisation holder.

No bioequivalence study was performed since the applied product is identical with the reference product Azilect (which has the same 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 was submitted with this application since the applied product is identical with the reference product Azilect. Thus, no bioequivalence study was performed which is found acceptable.

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 Rasagiline ratio.

Safety specification

Summary table of safety concerns as approved in RMP

IMPORTANT IDENTIFIED RISKS	• Orthostatic hypotension • Serotonin syndrome • Impulse control disorders • Concomitant use with antidepressants (SSRI, SnRI, tricyclic and tetracyclic antidepressants), CYP1A2 inhibitors or MAO inhibitors
IMPORTANT POTENTIAL RISKS	• Hypertension • Malignant Melanoma • Concomitant use with pethidine or sympathomimetics
MISSING INFORMATION	• Pregnant and lactating women

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 MAH has satisfactorily responded to the questions.

Issue is resolved

The RMP is approved.

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 Azilect EMEA/H/C/000574. The bridging report submitted by the applicant has been found acceptable.

VI. OVERALL CONCLUSION, BENEFIT/RISK ASSESSMENT AND RECOMMENDATION

The benefit/risk ratio is considered positive and Rasagiline ratio, 1 mg, tablet is 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

The Decentralised procedure for Rasagiline ratio, 1 mg, tablet was positively finalised on 2015-11-18.

Public Assessment Report – Update

Scope	Procedure number	Product Information affected	Date of start of the procedure	Date of end of procedure	Approval/ non approval	Assessment report attached
						Y/N (version)