EMA/371377/2020

European Medicines Agency decision

P/0279/2020

of 24 July 2020

on the acceptance of a modification of an agreed paediatric investigation plan for imipenem (monohydrate) / cilastatin (sodium) / relebactam (MK-7655A) (Recarbrio), (EMEA-001809-PIP01-15-M02) in accordance with Regulation (EC) No 1901/2006 of the European Parliament and of the Council

Disclaimer

This decision does not constitute entitlement to the rewards and incentives referred to in Title V of Regulation (EC) No 1901/2006.

Only the English text is authentic.

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The European Medicines Agency,

Having regard to the Treaty on the Functioning of the European Union,

Having regard to Regulation (EC) No 1901/2006 of the European Parliament and of the Council of 12 December 2006 on medicinal products for paediatric use and amending Regulation (EEC) No. 1768/92, Directive 2001/20/EC, Directive 2001/83/EC and Regulation (EC) No 726/2004[1],

Having regard to Regulation (EC) No 726/2004 of the European Parliament and of the Council of 31 March 2004 laying down Community procedures for the authorisation and supervision of medicinal products for human and veterinary use and establishing a European Medicines Agency[2],

Having regard to the European Medicines Agency's decision P/0163/2016 issued on 15 June 2016 and the decision P/0173/2019 issued on 15 May 2019,

Having regard to the application submitted by Merck Sharp & Dohme (Europe), Inc. on 24 March 2020 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan with a deferral,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 26 June 2020, in accordance with Article 22 of Regulation (EC) No 1901/2006,

Having regard to Article 25 of Regulation (EC) No 1901/2006,

Whereas:

(1) The Paediatric Committee of the European Medicines Agency has given an opinion on the acceptance of changes to the agreed paediatric investigation plan.

(2) It is therefore appropriate to adopt a decision on the acceptance of changes to the agreed paediatric investigation plan.

[1] OJ L 378, 27.12.2006, p.1.
[2] OJ L 136, 30.4.2004, p. 1.

Has adopted this decision:

Article 1

Changes to the agreed paediatric investigation plan for imipenem (monohydrate) / cilastatin (sodium) / relebactam (MK-7655A) (Recarbrio), powder for solution for infusion, parenteral use, are hereby accepted in the scope set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices.

Article 2

This decision is addressed to Merck Sharp & Dohme (Europe), Inc., Clos du Lynx, 5, 1200 - Brussels, Belgium.

EMA/PDCO/201936/2020
Amsterdam, 26 June 2020

Opinion of the Paediatric Committee on the acceptance of a modification of an agreed Paediatric Investigation Plan

EMEA-001809-PIP01-15-M02

Scope of the application

Active substance(s):

Imipenem (monohydrate) / cilastatin (sodium) / relebactam (MK-7655A)

Invented name:

Recarbrio

Condition(s):

Treatment of infections caused by gram-negative organisms

Authorised indication(s):

See Annex II

Pharmaceutical form(s):

Powder for solution for infusion

Route(s) of administration:

Parenteral use

Name/corporate name of the PIP applicant:

Merck Sharp & Dohme (Europe), Inc.

Information about the authorised medicinal product:

See Annex II

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Merck Sharp & Dohme (Europe), Inc. submitted to the European Medicines Agency on 24 March 2020 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/0163/2016 issued on 15 June 2016 and the decision P/0173/2019 issued on 15 May 2019.

The application for modification proposed changes to the agreed paediatric investigation plan.

The procedure started on 30 April 2020.

Scope of the modification

Some measures and timelines of the Paediatric Investigation Plan have been modified.

Opinion

1. The Paediatric Committee, having assessed the application in accordance with Article 22 of Regulation (EC) No 1901/2006 as amended, recommends as set out in the appended summary report:

   - to agree to changes to the paediatric investigation plan in the scope set out in the Annex I of this opinion.

   The Norwegian Paediatric Committee member agrees with the above-mentioned recommendation of the Paediatric Committee.

2. The measures and timelines of the paediatric investigation plan are set out in the Annex I.

This opinion is forwarded to the applicant and the Executive Director of the European Medicines Agency, together with its annexes and appendix.

Annex I

The subset(s) of the paediatric population and condition(s) covered by the waiver and the measures and timelines of the agreed paediatric investigation plan (PIP)

1. Waiver

Not applicable.

2. Paediatric investigation plan

2.1. Condition:

Treatment of infections caused by gram-negative organisms

2.1.1. Indication(s) targeted by the PIP

Treatment of hospital-acquired bacterial pneumonia (HABP) and ventilator-associated bacterial pneumonia (VABP)

Treatment of bacterial infections caused by aerobic gram-negative organisms in patients with limited treatment options

2.1.2. Subset(s) of the paediatric population concerned by the paediatric development

From birth to less than 18 years of age

2.1.3. Pharmaceutical form(s)

Powder for solution for infusion

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	0	Not applicable.
Non-clinical studies	2	**Study 1** Dose range-finding juvenile toxicity study **Study 2** Definitive juvenile toxicity study

Clinical studies	2	**Study 3 (Ped001)** Open-label study to evaluate the pharmacokinetics, safety and tolerability of a single dose of imipenem (monohydrate) / cilastatin (sodium) / relebactam (MK-7655A) and to identify the appropriate dose of MK-7655A in children from birth to less than 18 years of age with gram-negative bacterial infections **Study 4 (Ped002)** Open-label, randomised, active-controlled trial to evaluate safety, tolerability and efficacy of MK-7655A in children from birth to less than 18 years of age with gram-negative bacterial infections ***Study 5:*** *deleted during procedure EMEA-001809-PIP01-15-M01*
Extrapolation, modelling and simulation studies	2	**Study 6 (Modelling & Simulation Study)** Modelling and simulation study to select doses in children from birth to less than 18 years of age with gram-negative bacterial infections **Study 7 (Extrapolation Study)** Extrapolation study to evaluate the use of MK-7655A in the proposed paediatric indications in children from birth to less than 18 years of age with serious gram-negative bacterial infections
Other studies	0	Not applicable.
Other measures	0	Not applicable.

3. Follow-up, completion and deferral of PIP

Concerns on potential long term safety/efficacy issues in relation to paediatric use:	Yes
Date of completion of the paediatric investigation plan:	By June 2023
Deferral for one or more measures contained in the paediatric investigation plan:	Yes

Annex II

Information about the authorised medicinal product

Condition(s) and authorised indication(s):

1. Treatment of infections caused by gram-negative organisms

Authorised indication(s):

- Recarbrio is indicated for the treatment of infections due to aerobic gram-negative organisms in adults with limited treatment options.

Authorised pharmaceutical form(s):

Powder for solution for infusion

Authorised route(s) of administration:

Intravenous use