EMA/192367/2021

European Medicines Agency decision

P/0167/2021

of 14 April 2021

on the acceptance of a modification of an agreed paediatric investigation plan for upadacitinib (Rinvoq), (EMEA-001741-PIP01-14-M04) 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/0288/2015 issued on 27 November 2015, the decision P/0363/2017 issued on 1 December 2017 and the decision P/0322/2019 issued on 11 September 2019 and the decision P/0347/2020 issued on 9 September,

Having regard to the application submitted by AbbVie Ltd on 30 November 2020 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan with a deferral and a waiver,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 26 February 2021, 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 and to the deferral.

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

[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 upadacitinib (Rinvoq), age-appropriate oral solid dosage form, age-appropriate oral liquid dosage form, prolonged-release tablet, oral use, including changes to the deferral, 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 AbbVie Ltd, AbbVie House, Vanwall Road, SL6 4UB – Maidenhead, United Kingdom.

EMA/PDCO/658961/2020
Amsterdam, 26 February 2021

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

EMEA-001741-PIP01-14-M04

Scope of the application

Active substance(s):

Upadacitinib

Invented name:

Rinvoq

Condition(s):

Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, psoriatic arthritis, spondyloarthritis and juvenile idiopathic arthritis)

Authorised indication(s):

See Annex II

Pharmaceutical form(s):

Age-appropriate oral solid dosage form

Age-appropriate oral liquid dosage form

Prolonged-release tablet

Route(s) of administration:

Oral use

Name/corporate name of the PIP applicant:

AbbVie Ltd

Information about the authorised medicinal product:

See Annex II

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, AbbVie Ltd submitted to the European Medicines Agency on 30 November 2020 an application for modification of the agreed paediatric investigation plan with a deferral and a waiver as set out in the European Medicines Agency's decision P/0288/2015 issued on 27 November 2015, the decision P/0363/2017 issued on 1 December 2017 and the decision P/0322/2019 issued on 11 September 2019 and the decision P/0347/2020 issued on 9 September 2020.

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

The procedure started on 4 January 2021.

Scope of the modification

Some 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 and to the deferral 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 and the subset(s) of the paediatric population and condition(s) covered by the waiver 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

1.1. Condition

Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, psoriatic arthritis, spondyloarthritis and juvenile idiopathic arthritis)

The waiver applies to:

- the paediatric population from birth to less than 1 year of age;
- age-appropriate oral solid dosage form, age-appropriate oral liquid dosage form, prolonged-release tablet, oral use;
- on the grounds that the disease or condition for which the specific medicinal product is intended does not occur in the specified paediatric subset(s).

2. Paediatric investigation plan

2.1. Condition:

Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, psoriatic arthritis, spondyloarthritis and juvenile idiopathic arthritis)

2.1.1. Indication(s) targeted by the PIP

Treatment of juvenile idiopathic arthritis

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

From 1 to less than 18 years of age

2.1.3. Pharmaceutical form(s)

Age-appropriate oral solid dosage form

Age-appropriate oral liquid dosage form

Prolonged-release tablet

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	1	**Study 1** Development of age-appropriate oral solid dosage form (dispersible tablet or multi-particulate granules) or age-appropriate oral liquid dosage form

Non-clinical studies	2	**Study 2** Dose range-finding juvenile toxicity study **Study 3** Definitive juvenile toxicity study to evaluate toxicity and impact of upadacitinib on neonatal/juvenile development
Clinical studies	3	**Study 4** Open-label, multiple dose study to evaluate the pharmacokinetics, safety, and tolerability and to confirm the dosing regimen of upadacitinib in children with active polyarticular course JIA **Study 5** Randomised, placebo-controlled, double-blind withdrawal study to evaluate the safety and efficacy of upadacitinib in children with active polyarticular course JIA **Study 6** Randomised, placebo-controlled, double-blind withdrawal study to evaluate the safety and efficacy of multiple doses of upadacitinib in children with active systemic JIA
Extrapolation, modelling and simulation studies	1	**Study 7** Population pharmacokinetic two compartment model that characterizes the pharmacokinetic parameters, the inter- and intra-subject variability, and relationship between pharmacokinetic parameters and the relevant covariates
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 August 2028
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 Chronic Idiopathic Arthritis (including rheumatoid arthritis, psoriatic arthritis, spondyloarthritis and juvenile idiopathic arthritis)

Authorised indication(s):

Rinvoq is indicated for the treatment of moderate to severe active rheumatoid arthritis in adults

Authorised pharmaceutical form(s):

Prolonged-release tablet

Authorised route(s) of administration:

Oral use