

EMA/812166/2018

European Medicines Agency decision P/0360/2018

of 7 December 2018

on the acceptance of a modification of an agreed paediatric investigation plan for risdiplam (RO7034067), (EMA-002070-PIP01-16-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 726/2004¹,

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 Agency²,

Having regard to the European Medicines Agency's decision P/0284/2017 issued on 4 October 2017 and the decision P/0080/2018 issued on 16 March 2018,

Having regard to the application submitted by Roche Registration GmbH on 13 July 2018 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 19 October 2018, 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 risdiplam (RO7034067), powder for oral solution, oral 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 Roche Registration GmbH, Emil-Barell-Strasse 1, 79639 - Grenzach-Wyhlen, Germany.

EMA/PDCO/499176/2018
London, 19 October 2018

Opinion of the Paediatric Committee on the acceptance of a modification of an agreed Paediatric Investigation Plan EMA-002070-PIP01-16-M02

Scope of the application

Active substance(s):

Risdiplam (RO7034067)

Condition(s):

Treatment of spinal muscular atrophy

Pharmaceutical form(s):

Powder for oral solution

Route(s) of administration:

Oral use

Name / corporate name of the PIP applicant:

Roche Registration GmbH

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Roche Registration GmbH submitted to the European Medicines Agency on 13 July 2018 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/0284/2017 issued on 4 October 2017 and the decision P/0080/2018 issued on 16 March 2018.

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

The procedure started on 21 August 2018.

Scope of the modification

Some measures 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 annex 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 spinal muscular atrophy

2.1.1. Indication(s) targeted by the PIP

Treatment of spinal muscular atrophy

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 oral solution

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	1	Study 1: Development of an age-appropriate oral liquid dosage form
Non-clinical studies	1	Study 2: In-vitro study in plasma samples of infants and children to investigate RO7034067 plasma free fraction in the human paediatric population
Clinical studies	4	Study 3: Two-part multi-centre study to investigate the efficacy, safety and tolerability, pharmacokinetics (PK) and pharmacodynamics (PD) of RO7034067 in patients with type 2 and type 3 SMA Study 4: Two-part, multi-centre, single-arm, open-label study to investigate the efficacy, safety and tolerability, pharmacokinetics and pharmacodynamics of RO7034067 in patients with type 1 SMA

		Study 5: Multi-centre, exploratory, non-comparative and open-label study to investigate the safety, tolerability, pharmacokinetics and PK/PD relationship of RO7034067 patients with SMA who have previously participated in the Moonfish (BP29420) study with the splicing modifier RO6885247 or previously been treated with nusinersen (SPINRAZA), AVXS-101 or olesoxime Study 6: Multi-centre, single-arm, open-label study to investigate the efficacy, safety and tolerability, and PK/PD of RO7034067 in infants genetically diagnosed with SMA and pre-symptomatic
Extrapolation, modelling and simulation studies	4	Study 7: Physiologically based pharmacokinetic (PBPK) model of RO7034067 Study 8: Population pharmacokinetic (PopPK) model of RO7034067 Study 9: Extrapolation study to support the use of RO7034067 for the treatment of children with SMA Type 1, 2 and 3 aged between 7 months and 2 years based on extrapolation of in house data from younger infants (Type 1 SMA) and from older children and young adults (Type 2 and 3 SMA) Study 10: Extrapolation study to support the use of RO7034067 for the treatment of ambulant Type 3 SMA patients based on extrapolation of in house data from younger infants (Type 1 SMA) and from older children and young adults (Type 2 and 3 SMA)
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 September 2020
Deferral for one or more measures contained in the paediatric investigation plan:	Yes