

EMA/541006/2018

European Medicines Agency decision

P/0321/2018

of 12 September 2018

on the agreement of a paediatric investigation plan and on the granting of a deferral and on the granting of a waiver for inclisiran sodium (EMEA-002214-PIP01-17) 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 application submitted by The Medicines Company UK Ltd. on 6 July 2017 under Article 16(1) of Regulation (EC) No 1901/2006 also requesting a deferral under Article 20 of said Regulation and a waiver under Article 13 of said Regulation,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 27 July 2018, in accordance with Article 17 of Regulation (EC) No 1901/2006 and Article 21 of said Regulation and Article 13 of said Regulation,

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 agreement of a paediatric investigation plan and on the granting of a deferral and on the granting of a waiver.
- (2) It is therefore appropriate to adopt a decision agreeing a paediatric investigation plan.
- (3) It is therefore appropriate to adopt a decision granting a deferral.
- (4) It is therefore appropriate to adopt a decision granting a waiver.

¹ OJ L 378, 27.12.2006, p.1.

² OJ L 136, 30.4.2004, p. 1.

Has adopted this decision:

Article 1

A paediatric investigation plan for inclisiran sodium, solution for injection, subcutaneous use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby agreed.

Article 2

A deferral for inclisiran sodium, solution for injection, subcutaneous use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby granted.

Article 3

A waiver for inclisiran sodium, solution for injection, subcutaneous use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby granted.

Article 4

This decision is addressed to The Medicines Company UK Ltd., 115L Milton Park, OX14 4SA - Abingdon, United Kingdom.

EMA/367714/2018
London, 27 July 2018

Opinion of the Paediatric Committee on the agreement of a Paediatric Investigation Plan and a deferral and a waiver

EMA-002214-PIP01-17

Scope of the application

Active substance(s):

Inclisiran sodium

Condition(s):

Treatment of elevated cholesterol

Pharmaceutical form(s):

Solution for injection

Route(s) of administration:

Subcutaneous use

Name / corporate name of the PIP applicant:

The Medicines Company UK Ltd.

Basis for opinion

Pursuant to Article 16(1) of Regulation (EC) No 1901/2006 as amended, The Medicines Company UK Ltd. submitted for agreement to the European Medicines Agency on 6 July 2017 an application for a paediatric investigation plan for the above mentioned medicinal product and a deferral under Article 20 of said Regulation and a waiver under Article 13 of said Regulation.

The procedure started on 15 August 2017.

Supplementary information was provided by the applicant on 7 May 2018. The applicant proposed modifications to the paediatric investigation plan.

A meeting with the Paediatric Committee took place on 25 July 2018

Opinion

1. The Paediatric Committee, having assessed the proposed paediatric investigation plan in accordance with Article 17 of Regulation (EC) No 1901/2006 as amended, recommends as set out in the appended summary report :

- to agree the paediatric investigation plan in accordance with Article 17(1) of said Regulation;
- to grant a deferral in accordance with Article 21 of said Regulation;
- to grant a waiver for one or more subsets of the paediatric population in accordance with Article 13 of said Regulation and concluded in accordance with Article 11(1)(c) of said Regulation, on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments for paediatric patients.

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

2. The measures and timelines of the agreed 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 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

1.1. Condition:

Treatment of elevated cholesterol

The waiver applies to:

- the paediatric population below 6 years of age;
- solution for injection, subcutaneous use;
- on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments.

2. Paediatric investigation plan

2.1. Condition:

Treatment of elevated cholesterol.

2.1.1. Indication(s) targeted by the PIP

Treatment of heterozygous familial hypercholesterolemia (HeFH).

Treatment of homozygous familial hypercholesterolemia (HoFH).

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

From 6 to less than 18 years of age.

2.1.3. Pharmaceutical form(s)

Solution for injection.

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	0	Not applicable.
Non-clinical studies	0	Not applicable.
Clinical studies	4	Study 1 Two-stage, randomized study to evaluate the safety, tolerability and efficacy of inclisiran versus placebo, used in combination with other lipid lowering therapy in paediatric subjects with homozygous familial hypercholesterolaemia (HoFH) from 12 to less than 18 years of age [ORION-13 (MDCO-PCS-17-13)].

Area	Number of measures	Description
		Study 2 Two-stage, randomized, study to evaluate the safety, tolerability and efficacy of inclisiran versus placebo, used in combination with other lipid lowering therapy in paediatric patients with heterozygous familial hypercholesterolaemia (HeFH) from 12 to less than 18 years of age [ORION-16 (MDCO-PCS-17-11)]. Study 3 Two-stage, randomised, controlled versus placebo study to evaluate the safety, tolerability and efficacy of inclisiran, used in combination with other lipid lowering therapy in children with homozygous familial hypercholesterolaemia (HoFH) from 6 to less than 12 years of age [ORION-19 (MDCO-PCS-17-10)]. Study 4 Two-stage, randomized, study to evaluate the safety, tolerability and efficacy of inclisiran versus placebo, used in combination with other lipid lowering therapy in paediatric patients with heterozygous familial hypercholesterolaemia (HeFH) from 6 to less than 12 years of age [ORION-20 (MDCO-PCS-17-12)].
Extrapolation, modelling and simulation studies	0	Not applicable.
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 2025
Deferral for one or more measures contained in the paediatric investigation plan:	Yes