EMA/207916/2022

European Medicines Agency decision

P/0164/2022

of 13 May 2022

on the acceptance of a modification of an agreed paediatric investigation plan for bis-choline tetrathiomolybdate (EMEA-002232-PIP02-19-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 Union procedures for the authorisation and supervision of medicinal products for human use and establishing a European Medicines Agency[2],

Having regard to the European Medicines Agency's decision P/0234/2020 issued on 19 June 2020 and the decision P/0261/2021 issued on 7 July 2021,

Having regard to the application submitted by Alexion Europe S.A.S. on 15 December 2021 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 25 March 2022, 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, as amended.
[2] OJ L 136, 30.4.2004, p. 1, as amended.

Has adopted this decision:

Article 1

Changes to the agreed paediatric investigation plan for bis-choline tetrathiomolybdate, coated tablet, age-appropriate oral solid dosage form, 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 Alexion Europe S.A.S., 103-105 rue Anatole France, 92300 - Levallois-Perret, France.

EMA/PDCO/7778/2022
Amsterdam, 25 March 2022

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

EMEA-002232-PIP02-19-M02

Scope of the application

Active substance(s):

Bis-choline tetrathiomolybdate

Condition(s):

Treatment of Wilson disease

Pharmaceutical form(s):

Coated tablet

Age-appropriate oral solid dosage form

Route(s) of administration:

Oral use

Name/corporate name of the PIP applicant:

Alexion Europe S.A.S.

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Alexion Europe S.A.S. submitted to the European Medicines Agency on 15 December 2021 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/0234/2020 issued on 19 June 2020 and the decision P/0261/2021 issued on 7 July 2021.

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

The procedure started on 31 January 2022.

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 Paediatric Committee member of Norway 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 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 Wilson disease

The waiver applies to:

- the paediatric population from birth to less than 3 years of age;
- coated tablet, age-appropriate oral solid dosage form, oral 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 Wilson disease

2.1.1. Indication(s) targeted by the PIP

Treatment of children and adolescents from 3 years to less than 18 years of age with Wilson disease

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

From 3 years to less than 18 years of age

2.1.3. Pharmaceutical form(s)

Coated tablet

Age-appropriate oral solid dosage form

2.1.4. Measures

Area	**Description**
Quality-related studies	**Study 1** Development of an age-appropriate oral solid dosage form.
Non-clinical studies	Not applicable.
Clinical studies	**Study 2 (WTX101-301)** Randomised, rater-blinded efficacy and safety study of bis-choline tetrathiomolybdate (ALXN1840) compared with standard of care in adolescents 12 years to less than 18 years of age (and adults) with Wilson disease.

	Study 3 (ALXN1840-WD-302) Open-label, randomised, controlled, efficacy and safety study of bis-choline tetrathiomolybdate (ALXN1840) compared with standard of care in children from 3 years to less than 18 years of age with Wilson disease.
Extrapolation, modelling and simulation studies	**Study 4 (ALXN1840-WD-303)** Extrapolation study to evaluate the efficacy of bis-choline tetrathiomolybdate (ALXN1840) in paediatric patients from 3 to less than 18 years of age with Wilson disease.
Other studies	Not applicable.
Other measures	Not applicable.

3. Follow-up, completion and deferral of PIP

Concerns on potential long term safety/efficacy issues in relation to paediatric use:	No
Date of completion of the paediatric investigation plan:	By November 2024
Deferral for one or more measures contained in the paediatric investigation plan:	Yes