EMA/494326/2023

European Medicines Agency decision

P/0484/2023

of 30 November 2023

on the acceptance of a modification of an agreed paediatric investigation plan for human fibrinogen concentrate (BT524) (EMEA-001931-PIP01-16-M03) 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/0106/2017 issued on 11 April 2017, the decision P/0285/2018 issued on 12 September 2018 and the decision P/0232/2020 issued on 19 June 2020,

Having regard to the application submitted by Biotest AG on 29 June 2023 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 13 October 2023, 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 human fibrinogen concentrate (BT524), powder for solution for injection or infusion, intravenous 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 Biotest AG, 5 Landsteinerstrasse, 63303 – Dreieich, Germany.

EMA/PDCO/321311/2023 Corr[1]
Amsterdam, 13 October 2023

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

EMEA-001931-PIP01-16-M03

Scope of the application

Active substance(s):

Human fibrinogen concentrate (BT524)

Invented name and authorisation status:

See Annex II

Condition(s):

Treatment of congenital fibrinogen deficiency

Pharmaceutical form(s):

Powder for solution for injection or infusion

Route(s) of administration:

Intravenous use

Name/corporate name of the PIP applicant:

Biotest AG

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Biotest AG submitted to the European Medicines Agency on 29 June 2023 an application for modification of the agreed paediatric investigation plan as set out in the European Medicines Agency's decision P/0106/2017 issued on 11 April 2017, the decision P/0285/2018 issued on 12 September 2018 and the decision P/0232/2020 issued on 19 June 2020.

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

The procedure started on 14 August 2023.

[1] 26 October 2023

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 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 congenital fibrinogen deficiency

2.1.1. Indication(s) targeted by the PIP

Treatment of congenital fibrinogen deficiency

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 injection or infusion

2.1.4. Measures

Area	Description
Quality-related studies	Not applicable
Non-clinical studies	Not applicable
Clinical studies	**Study 1** Open-label, non-controlled PK/PD and safety and efficacy study: Part I: investigation of 14 day single-dose pharmacokinetics and pharmacodynamics of human fibrinogen concentrate in patients from birth to less than 18 years of age (and adults) with congenital fibrinogen deficiency (afibrinogenemia or severe hypofibrinogenemia). Part II: investigation of hemostatic efficacy and safety of single and/or repetitive intravenous infusions of human fibrinogen concentrate for on-demand prophylaxis and/or on-demand treatment of bleeding events in patients from birth to less than 18 years of age (and adults) with congenital fibrinogen deficiency (afibrinogenemia or severe hypofibrinogenemia) (Study 984).
Extrapolation, modelling and	**Study 2** Semi-mechanistic population PK/PD model (Fibrinogen Antigen/Fibrinogen Activity) with subsequent link to the surrogate of efficacy (maximum clot

simulation studies	firmness) to support the dosing strategy and activity of human fibrinogen concentrate in patients from birth to less than 18 years of age with congenital fibrinogen deficiency (afibrinogenemia or severe hypofibrinogenemia). **Study 3** Extrapolation study to support the understanding of dosing and efficacy of human fibrinogen concentrate in patients from birth to less than 18 years of age with congenital fibrinogen deficiency (afibrinogenemia or severe hypofibrinogenemia) using data obtained in part I and II of Study 984 (PIP study 1).
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:	Yes
Date of completion of the paediatric investigation plan:	By September 2020
Deferral for one or more measures contained in the paediatric investigation plan:	No

Annex II

Information about the authorised medicinal product

Information provided by the applicant:

The product is not authorised anywhere in the European Community.