EMA/245271/2021

# European Medicines Agency decision

P/0190/2021

of 10 May 2021

on the agreement of a paediatric investigation plan and on the granting of a deferral for deferoxamine (mesylate) / histidine / tryptophan / aspartic acid / n-acetyl-histidine (monohydrate) / glycine / alpha-ketoglutaric acid / arginine / potassium chloride / magnesium chloride (hexahydrate) / calcium chloride (dihydrate) / sodium chloride / alanine / 3,4-dimethoxy-N-methylbenzohydroxamic acid (Custodiol-N) (EMEA-002735-PIP03-20) 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 application submitted by Dr. Franz Köhler Chemie GmbH on 21 December 2020 under Article 16(1) of Regulation (EC) No 1901/2006 also requesting a deferral under Article 20 of said Regulation,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 26 March 2021, in accordance with Article 17 of Regulation (EC) No 1901/2006 and Article 21 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.

(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.

[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 deferoxamine (mesylate) / histidine / tryptophan / aspartic acid / n-acetyl-histidine (monohydrate) / glycine / alpha-ketoglutaric acid / arginine / potassium chloride / magnesium chloride (hexahydrate) / calcium chloride (dihydrate) / sodium chloride / alanine / 3,4-dimethoxy-N-methylbenzohydroxamic acid (Custodiol-N), solution for organ preservation, 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 deferoxamine (mesylate) / histidine / tryptophan / aspartic acid / n-acetyl-histidine (monohydrate) / glycine / alpha-ketoglutaric acid / arginine / potassium chloride / magnesium chloride (hexahydrate) / calcium chloride (dihydrate) / sodium chloride / alanine / 3,4-dimethoxy-N-methylbenzohydroxamic acid (Custodiol-N), solution for organ preservation, 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***

This decision is addressed to Dr. Franz Köhler Chemie GmbH, Werner-von-Siemens-Str. 14-28, 64625 – Bensheim, Germany.

EMA/PDCO/103789/2021
Amsterdam, 26 March 2021

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

EMEA-002735-PIP03-20

## Scope of the application

**Active substance(s):**

Deferoxamine (mesylate) / histidine / tryptophan / aspartic acid / n-acetyl-histidine (monohydrate) / glycine / alpha-ketoglutaric acid / arginine / potassium chloride / magnesium chloride (hexahydrate) / calcium chloride (dihydrate) / sodium chloride / alanine / 3,4-dimethoxy-N-methylbenzohydroxamic acid (Custodiol-N)

**Condition(s):**

Heart transplantation

**Pharmaceutical form(s):**

Solution for organ preservation

**Route(s) of administration:**

Not applicable

**Name/corporate name of the PIP applicant:**

Dr. Franz Köhler Chemie GmbH

## Basis for opinion

Pursuant to Article 16(1) of Regulation (EC) No 1901/2006 as amended, Dr. Franz Köhler Chemie GmbH submitted for agreement to the European Medicines Agency on 21 December 2020 an application for a paediatric investigation plan for the above mentioned medicinal product and a deferral under Article 20 of said Regulation.

The procedure started on 26 January 2021.

## 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.

   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 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:*

Heart transplantation

### 2.1.1. Indication(s) targeted by the PIP

Preservation of hearts prior to heart transplantation

### 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)

Solution for organ preservation

### 2.1.4. Measures

| Area | Number of measures | Description |
|---|---|---|
| Quality-related studies | 0 | Not applicable. |
| Non-clinical studies | 2 | **Study 1 (FKC-001_02_rat_tox_iv)**<br>Single dose intravenous toxicity study of 3,4-dimethoxy-N-methylbenzohydroxamic acid in rats.<br>**Study 2 (FKC-003_01_dev_reprotox Rat)**<br>Reproductive toxicity study of 3,4-dimethoxy-N-methylbenzohydroxamic acid in rats. |
| Clinical studies | 1 | **Study 3 (CL-N-HTX-Paed-II/10/20)**<br>Randomised single-blind study to compare the safety of the investigational preservation solution to that of its precursor product in children from birth to less than 18 years of age undergoing heart transplantation. |
| 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: | No |
| Date of completion of the paediatric investigation plan: | By October 2023 |
| Deferral for one or more measures contained in the paediatric investigation plan: | Yes |