FRENCH AGENCY FOR VETERINARY MEDICINAL PRODUCTS
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DECENTRALISED PROCEDURE

PUBLICLY AVAILABLE ASSESSMENT REPORT FOR A VETERINARY MEDICINAL PRODUCT

Alfadexx 2 mg/mL solution for injection for horses, cattle, goats, pigs, dogs and cats

DATE: 07/07/2021

MODULE 1

PRODUCT SUMMARY

EU Procedure number	FR/V/0430/001/DC
Name, strength and pharmaceutical form	Alfadexx 2 mg/ml solution for injection for horses, cattle, goats, pigs, dogs and cats
Applicant	Alfasan Nederland B.V. Kuipersweg 9 3449 JA Woerden The Netherlands
Active substance(s)	Dexamethasone (as dexamethasone sodium phosphate)
ATC Vetcode	QH02AB02
Target species	Horses, cattle, goats, pigs, dogs and cats
Indication for use	<u>Horses, cattle, goats, pigs, dogs and cats:</u> Treatment of inflammation and allergic reactions. <u>Horses:</u> Treatment of arthritis, bursitis or tenosynovitis. <u>Cattle:</u> Treatment of primary ketosis (Acetonemia). Induction of parturition. <u>Goats:</u> Treatment of primary ketosis (Acetonemia).

MODULE 2

The Summary of Product Characteristics (SPC) for this product is available on the website http://www.anmv.anses.fr/

MODULE 3

PUBLIC ASSESSMENT REPORT

Legal basis of original application	Article 13(1) generic application of Directive 2001/82/EC as amended
Date of completion of the original decentralised procedure	30/06/2021
Date product first authorised in the Reference Member State (MRP only)	Not applicable
Concerned Member States for original procedure	AT, BE, BG, CY, CZ, DE, DK, EE, EL, ES, FI, HR, HU, IE, IS, IT, LT, LU, LV, NL, NO, PL, PT, RO, SE, SK

I. SCIENTIFIC OVERVIEW

The product is produced and controlled using validated methods and tests, which ensure the consistency of the product released on the market.
It has been shown that the product can be safely used in the target species.
The product is safe for the user, the consumer of foodstuffs from treated animals and for the environment, when used as recommended. Suitable warnings and precautions are indicated in the SPC.
The efficacy of the product was demonstrated according to the claims made in the SPC. The overall risk/benefit analysis is in favour of granting a marketing authorisation.

II. QUALITY ASPECTS

A. Composition

The product contains the active substance Dexamethasone (as dexamethasone sodium phosphate) at a concentration of 2 mg/ml and the following excipients :
Benzyl alcohol, Sodium chloride, Sodium citrate dehydrate, Citric acid, Sodium hydroxide and water for injections.
The packaging of the finished product is as described on the SPC. The particulars of the containers and controls performed are provided and conform to the regulation.

The product is an established pharmaceutical form and its development is adequately described in accordance with the relevant European guidelines.

B. Method of Preparation of the Product

The product is manufactured fully in accordance with the principles of good manufacturing practice from a licensed manufacturing site.
Process validation data on the product have been presented in accordance with the relevant European guidelines.

The product is manufactured in accordance with the European Pharmacopoeia and relevant European guidelines.

C. Control of Starting Materials

The active substance is Dexamethasone (as dexamethasone sodium phosphate) an established active substance described in the European Pharmacopoeia. The active substance is manufactured in accordance with the principles of good manufacturing practice.
The active substance specification is considered adequate to control the quality of the material. Batch analytical data demonstrating compliance with this specification have been provided.

D. Specific Measures concerning the Prevention of the Transmission of Animal Spongiform Encephalopathies

There are no substances within the scope of the TSE Guideline present or used in the manufacture of this product.

E. Control on intermediate products

Not applicable.

F. Control Tests on the Finished Product

The finished product specification controls the relevant parameters for the pharmaceutical form. The tests in the specification, and their limits, have been justified and are considered appropriate to adequately control the quality of the product.
Satisfactory validation data for the analytical methods have been provided.
Batch analytical data from the proposed production site have been provided demonstrating compliance with the specification.

G. Stability

A re-test period for the active substance is set in the certificate of suitability issued by EDQM.

Stability data on the finished product have been provided in accordance with applicable European guidelines, demonstrating the stability of the product throughout its shelf life when stored under the approved conditions.

An in-use shelf-life as detailed on the SPC has been supported by appropriate data.

H. Genetically Modified Organisms

Not applicable.

J. Other Information

Not applicable.

III. SAFETY AND RESIDUES ASSESSMENT (PHARMACO-TOXICOLOGICAL)

III.A Safety Testing

Pharmacological Studies

As this is a generic application according to Article 13 and bioequivalence with the reference product has been demonstrated, results of pharmacological tests are not required.

The pharmacological aspects of this product are identical to the reference product.

Toxicological Studies

As this is a generic application according to Article 13, and bioequivalence with a reference product has been demonstrated, results of toxicological tests are not required.

The toxicological aspects of this product are identical to the reference product.

User Safety

Warnings and precautions as listed on the product literature are adequate to ensure safety to users of the product.

Environmental Risk Assessment

The applicant provided a first phase environmental risk assessment in compliance with the relevant guideline, which showed that no further assessment is required.
If used as recommended, the product will have a negligible impact on the environment.

III.B Residues documentation

Residue Studies

No depletion data was provided.

MRLs

The active substance, dexamethasone, is included in table 1 of the annex of the Commission regulation (EU) No. 37/2010, as follows:

Marker residue	Animal Species	MRL	Target Tissues	Other Provisions	Therapeutic Classification	Regulation
Dexamethasone	Bovine, caprine, porcine, Equidae	0.75 µg/kg 2.00 µg/kg 0.75 µg/kg	Muscle Liver Kidney	No entry	Cortocoïdes/ Glucocorticoïdes	37/2010 of 22.12.2009
	Bovine, caprine	0.30 µg/kg	Milk			

The MRL status of excipients of the product is indicated in the following table.

Excipient	MRL status
Benzyl alcohol	Table 1, all food species, no MRL required
Sodium chloride	Table 1, all food species, no MRL required
Sodium citrate dihydrate	*
Citric acid	*
Sodium hydroxyde	*
Water for injection	Out of scope list

* Covered with food additives (substance with a valid E number approved as additives in foodstuffs for human consumption)

The composition of the product is acceptable according to the European Regulation (EC) 470/2009.

Withdrawal Periods

It is a generic application submitted according to Article 13(1) of Directive 2001/82/EC as amended. The withdrawal periods are the same as those for the reference product.

Cattle and goats:
Meat and offal: 8 days
Milk: 72 hours

Pigs:
Meat and offal: 2 days following intramuscular administration
Meat and offal: 6 days following intravenous administration

Horses:
Meat and offal: 8 days
Not authorized for use in horses producing milk for human consumption.

IV. CLINICAL ASSESSMENT (EFFICACY)

IV.A Pre-Clinical Studies

Pharmacology

It is a generic application submitted according to Article 13(1) of Directive 2001/82/EC as amended by 2004/28/EC. The cited reference product is DEXADRESON (Intervet).

Pharmaceutical form
The test and the reference products have the same pharmaceutical form: solution for injection.

Active substance qualitative and quantitative composition
The test and reference products have the same qualitative and quantitative composition in active substance: 2.0 mg of dexamethasone per mL.

Bioequivalence studies
No bioequivalence study was performed.
In line with the current bioequivalence guideline (EMA/CVMP/016/00 –Rev.2), an exemption from bioequivalence study is claimed based on the similar formulations of the two products.

IV.B Clinical Studies

As this is a generic application according to Article 13(1) of Directive 2001/82/EC, as amended by Directive 2004/28/EC, and bioequivalence with a

reference product has been demonstrated, efficacy studies are not required. The efficacy claims for this product are equivalent to those of the reference product.

V. OVERALL CONCLUSION AND BENEFIT– RISK ASSESSMENT

The data submitted in the dossier demonstrate that when the product is used in accordance with the Summary of Product Characteristics, the risk benefit profile for the target species is favourable and the quality and safety of the product for humans and the environment is acceptable.