ASSURING THE SAFETY, QUALITY AND EFFICACY OF VETERINARY MEDICINES

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(Reference Member State)

DECENTRALISED PROCEDURE

PUBLICLY AVAILABLE ASSESSMENT REPORT FOR A VETERINARY MEDICINAL PRODUCT

Alfamed Fipronil 2.5 mg/ml, solution for cats and dogs (UK)
Fipralone 2.5 mg/ml cutaneous spray, solution for cats and dogs (IT) (NL)
Fipromedic 2.5 mg/ml cutaneous spray, solution for cats and dogs (FR)

PuAR correct as of 14/09/2018 when RMS was transferred to FR.
Please contact the RMS for future updates.

MODULE 1

PRODUCT SUMMARY

EU Procedure number	UK/V/0305/001/DC
Name, strength and pharmaceutical form	Alfamed Fipronil 2.5 mg/ml, solution for cats and dogs
Applicant	Francodex Sante Animale S.A.S.
Active substance(s)	Fipronil
ATC Vetcode	QP53AX15
Target species	Dogs Cats
Indication for use	Treatment of flea infestation (*Ctenocephalides* spp.) in dogs and cats. Treatment of tick infestation (*Ixodes ricinus, Rhipicephalus sanguineus*) in dogs and cats. Treatment of biting lice infestations in dogs (*Trichodectes canis*) and cats (*Felicola subrostratus*). The product can be used as part of a treatment strategy for the control of Flea Allergy Dermatitis (FAD).

MODULE 2

The Summary of Product Characteristics (SPC) for this product is available on the Heads of Medicines Agencies (veterinary) (HMA(v)) website (www.hma.eu).

MODULE 3

PUBLIC ASSESSMENT REPORT

Legal basis of original application	Application in accordance with Article 13.3 of Directive 2001/82/EC as amended by Directive 2004/28/EC
Date of completion of the original decentralised procedure	17 April 2009
Concerned Member States for original procedure	France Italy Netherlands

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 slight reactions observed are indicated in the SPC.
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 2.5 mg/ml fipronil as active substance and copovidone, isopropyl alcohol and water purified as excipients.

The container/closure system is a high density polyethylene white opaque bottle containing 100 ml of the product hermetically closed with a mechanical pump spray delivering 0.5 ml per spray (plunger in low density polyethylene) or high density polyethylene white opaque bottle containing 250 ml of the product hermetically closed with a mechanical trigger pump delivering 1.5 ml per spray (plunger in low density polyethylene) or high density polyethylene white opaque bottle containing 500 ml of the product hermetically closed with a mechanical trigger pump delivering 3 ml per spray (plunger in low density polyethylene).

The particulars of the containers and controls performed are provided and conform to the regulation.

The choice of the formulation is justified.

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.

C. Control of Starting Materials

The active substance is fipronil which is almost unabsorbed through the skin and the formulation is designed to deposit the active substance easily onto the animal.

There are three excipients used in the formulation and each has been used previously in veterinary medicines. Copovidone is employed in the formulation as a film-forming agent, with isopropyl alcohol and purified water being employed as solvents.

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.

All the excipients used in the final product have monographs in the Ph. Eur. and each comply with the requirements of the current edition of the Ph. Eur.

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 sites have been provided demonstrating compliance with the specification.

G. Stability

Stability data on the active substance have been provided in accordance with applicable European guidelines, demonstrating the stability of the active substance when stored under the approved conditions.

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.

H. Genetically Modified Organisms

Not applicable

J. Other information

Shelf life
Shelf-life of the veterinary medicinal product as packaged for sale: 3 years.
Shelf-life after first opening the immediate packaging: 1 year.

Special precautions for storage

Highly flammable.
Do not store above 25°C.
Protect from direct sunlight.

III. SAFETY AND RESIDUES ASSESSMENT (PHARMACO-TOXICOLOGICAL)

III.A Safety Testing

Pharmacological Studies
Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not required.

Toxicological Studies

Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not required.

Other Studies

Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not required.

User Safety

The applicant has not submitted a user risk assessment. The Expert has made reference to the user risk assessment for the reference product and has claimed that because the proposed formulation is “clinically equivalent” to the reference product, the user risk is the same and has proposed the same user warnings.

Ecotoxicity

The applicant provided a first phase environmental risk assessment in compliance with the relevant guidelines. The assessment ended at Phase I based on use in companion animals only. Warnings and precautions as listed on the product literature are adequate to ensure safety to the environment when the product is used as directed.

IV CLINICAL ASSESSMENT (EFFICACY)

Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not submitted.

IV.A Pre-Clinical Studies

Pharmacology

Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not provided.

Tolerance in the Target Species of Animals

Since the application is made in accordance with Article 13(3) of Directive 2004/28/EC, on the basis of essential similarity, data on this section of the dossier were not provided.

IV.B Clinical Studies

The applicant has provided two dose confirmation studies. The aim of one study was to determine and compare the efficacy of the Fiproline cutaneous spray solution with the reference product (Frontline spray) against fleas *(Ctenocephalides felis)* on dogs. The study was conducted on dogs. In the study it was demonstrated that the residual efficacies for the Fiproline cutaneous spray solution were not dissimilar to that of the reference product.

Another study was conducted to determine and compare the efficacy of the Fiproline cutaneous spray solution with Frontline spray against a French strain of the tick *Rhipicephalus sanguineus* on dogs. The study was conducted on dogs. The study concluded that 48 hours after application the therapeutic efficacy (based on geometric means) of both the Fiproline cutaneous spray solution and the reference product was more than 90%, when the two products were applied

at a dosage of 3-6 ml/kg bodyweight to dogs with an *R.sanguineus* (French strain) tick infestation.

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 benefit risk profile for the target species is favourable and the quality and safety of the product for humans and the environment is acceptable.

MODULE 4

POST-AUTHORISATION ASSESSMENTS

The SPC and package leaflet may be updated to include new information on the quality, safety and efficacy of the veterinary medicinal product. The current SPC is available on the Heads of Medicines Agencies (veterinary) (HMA(v)) website (www.hma.eu).

This section contains information on significant changes which have been made after the original procedure which are important for the quality, safety or efficacy of the product.

•	19 January 2015	Addition of an active substance manufacturer. Changes to the specification limits.
•	14 March 2014	Change in dimension of immediate packaging.
•	04 December 2013	Renewal procedure.
•	27 September 2012	Minor changes to the purification process of the active substance. Deletion of a non-specific specification parameter in the manufacture of the active substance. Increase in batch size range of active substance.
•	01 August 2012	Change of legal entity to POM-V.
•	11 August 2011	Grouped variation to increase the batch size of the finished product.
•	11 August 2011	Grouped variation to change the shelf life of the veterinary medicinal product as packaged for sale from 2 years to 3 years.