MINISTERIO
DE SANIDAD, SERVICIOS SOCIALES
E IGUALDAD

agencia española de
medicamentos y
productos sanitarios

DEPARTAMENTO DE
MEDICAMENTOS
VETERINARIOS

Agencia Española de Medicamentos y Productos Sanitarios

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28022 – Madrid
España
(Reference Member State)

DECENTRALISED PROCEDURE

[DRAFT] PUBLICLY AVAILABLE ASSESSMENT REPORT FOR A VETERINARY MEDICINAL PRODUCT

Kelacyl 100 mg/ml, solution for injection for cattle and pigs
Kelbomar 100 mg/ml, solution for injection for cattle and pigs

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MODULE 1

PRODUCT SUMMARY

EU Procedure number	ES/V/0189/001/DC
Name, strength and pharmaceutical form	Kelacyl 100 mg/ml, solution for injection (BG, CY, CZ, DE, EL, FR, HU, IE, IT, LT, PL, PT, RO, SK, UK) Kelbomar 100 mg/ml, solution for injection (BE, NL and LU)
Applicant	KELA N.V. St. Lenaartseweg 48 2320 Hoogstraten BELGIUM
Active substance	Marbofloxacin
ATC Vet code	QJ01MA93
Target species	Cattle and pigs (sows)
Indication for use	In cattle: - treatment of respiratory infections caused by strains of *Histophilus somni, Mannheimia haemolytica*, *Mycoplasma bovis*, *Pasteurella multocida* susceptible to marbofloxacin. - treatment of acute mastitis caused by strains of *Escherichia coli* susceptible to marbofloxacin during the lactation period. In pigs: - treatment of Postpartum Dysgalactia Syndrome – PDS-(Metritis Mastitis Agalactia syndrome), caused by bacterial strains susceptible to marbofloxacin.

MODULE 2

The Summary of Product Characteristics (SPC) for this product is available on the Heads of Medicines Agencies website (http://www.hma.eu).

MODULE 3

PUBLIC ASSESSMENT REPORT

Legal basis of original application	Decentralised application in accordance with Article Article 13 (1) of Directive 2001/82/EEC as amended by Directive 2004/28/EC.
Date of completion of the original decentralised procedure	23/01/2013
Date product first authorised in the Reference Member State (MRP only)	Not applicable
Concerned Member States for original procedure	BE, BG, CY, CZ, DE, EL, FR, HU, IE, IT, LT, LU, NL, PL, PT, RO, SK, UK

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:

Active substance:
Marbofloxacin 100 mg

Excipients:
Gluconolactone
Disodium edetate 0.10 mg
Monothioglycerol 1 mg
Metacresol 2 mg
Water for injection

The container/closure system is composed by an amber glass type II vial closed with bromobutyl rubber stopper and aluminium closure. The particulars of the containers and controls performed are provided and conform to the regulation.

The choice of the presence of preservatives is justified.

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.

C. Control of Starting Materials

The active substance is marbofloxacin, an established 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.

The master file is submitted according to the Active Substance Master File procedure using the CTD format (version 1.1 October 26, 2011)

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

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. The storage condition “Protect from light” is established.

The claim of a 28 days stability after broaching is based on the demonstration of stability for a batch broached and stored 28 days at 25 ± 2 °C / 60 ± 5 % RH.

H. Genetically Modified Organisms

Not applicable.

J. Other Information

Not applicable.

III. SAFETY AND RESIDUES ASSESSMENT (PHARMACOTOXICOLOGICAL)

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of toxicological and pharmacological tests are not required.

The aspects of safety of this product are identical to the reference products.

Warnings and precautions as listed on the product literature are the same as those of the reference products and are adequate to ensure safety of the product to users, the environment and consumers.

III.A Safety Testing

Pharmacological Studies

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of pharmacological tests are not required.

Toxicological Studies

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of toxicological tests are not required.

User Safety

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the applicant has provided a user safety assessment in compliance with the relevant guideline EMEA/CVMP/543/03.

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

Ecotoxicity

The applicant provided a first phase environmental risk assessment in compliance with the relevant guideline which showed that no further assessment is required.

Warnings and precautions as listed on the product literature are adequate to ensure safety to the environment when the product is used as directed.

III.B Residues documentation

Residue Studies

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of residue depletion studies are not required.

MRLs

Marbofloxacin is listed in Commission Regulation (EU) No 37/2010 on pharmacologically active substances and their classification regarding maximum residue limits in foodstuffs of animal origin.

MRLs are listed below:

Pharmacologically active Substance	Marker residue	Animal Species	MRL	Target Tissues	Other Provisions (according to Article 14 (7) of Regulation (EC) No 470/2009)	Therapeutic classification
Marbofloxacin	*Marbofloxacin*	*Bovine, porcine*	*150 µg/kg* *50 µg/kg* *150 µg/kg* *150 µg/kg*	*Muscle* *Fat* *Liver* *Kidney*	*For porcine species the fat MRL, relates to skin and fat in natural proportions*	*Anti-infectious agents/Antibiotics*
		Bovine	*75 µg/kg*	*Milk*		

Withdrawal Periods

Since the bioequivalence with reference products has been demonstrated, the withdrawal period proposed are identical to those authorised for reference products.

Cattle:

<u>8 mg/kg on a single occasion (IM)</u>

Meat and offal: 3 days

Milk: 72 hours

<u>2 mg/kg for 3 to 5 days (IV/SC/IM)</u>

Meat and offal: 6 days

Milk: 36 hours

Pigs (sows):

Meat and offal: 4 days

IV. CLINICAL ASSESSMENT (EFFICACY)

As this is a generic application according to Article 13 (1), 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 products.

IV.A Pre-Clinical Studies

Pharmacology

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of pharmacological tests are not required.

Tolerance in the Target Species of Animals

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of tolerance studies are not required.

Resistance

As this is a generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of resistance development are not required.

Adequate warnings and precautions appear on the product literature.

IV.B Clinical Studies

As this is generic application according to Article 13 (1) of Directive 2001/82/EC, as amended, and bioequivalence with reference products has been demonstrated, the results of clinical trials are not required.

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.

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 veterinary Heads of Agencies 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.

None