IPAR

Publicly Available Assessment Report for a
Veterinary Medicinal Product

DECTOMAX 10 mg/ml Solution for Injection for Cattle, Sheep and Pigs

PRODUCT SUMMARY

EU Procedure number	IE/V/0260/001/DC
Name, strength and pharmaceutical form	Dectomax 10 mg/ml Solution for Injection for Cattle, Sheep and Pigs
Active substance(s)	Doramectin
Applicant	Zoetis Belgium S.A. 2nd Floor, Building 10 Cherrywood Business Park Loughlinstown Co. Dublin
Legal basis of application	Generic application in accordance with Article 13(1) of Directive 2001/82/EC as amended.
Date of completion of procedure	18th April 2012
Target species	Cattle, Sheep and Pigs
Indication for use	Cattle: For the treatment and control of gastrointestinal nematodes, lungworms, eyeworms, warbles, lice, mange mites and ticks. Sheep: For treatment and control of gastrointestinal roundworms, mange mites and nasal bots Pigs: For treatment of mange mites, gastrointestinal roundworms, lungworms, kidney worms and sucking lice in pigs.
ATCvet code	QP 54AA03
Concerned Member States	AT, BG, CY, CZ, DK, EE, EL, ES, FI, FR, HU, IS, LT, LV, MT, NL, NO, PL, PT, RO, SE, SI, SK

PUBLIC ASSESSMENT REPORT

The public assessment report reflects the scientific conclusion reached by the HPRA at the end of the evaluation process and provides a summary of the grounds for approval of the marketing authorisation for the specific veterinary medicinal product. It is made available by the HPRA for information to the public, after the deletion of commercially confidential information. The legal basis for its creation and availability is contained in Article 25.4 of EC Directive 2001/82/EC as amended by Directive 2004/28/EC for veterinary medicinal products. It is a concise document which highlights the main parts of the documentation submitted by the applicant and the scientific evaluation carried out by the HPRA leading to the approval of the product for marketing in Ireland.

The Summary of Product Characteristics (SPC) for this product is available on the HPRA's website.

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 benefit/risk analysis is in favour of granting a marketing authorisation.

II. QUALITY ASPECTS

A.Qualitative and Quantitative Particulars

The product contains doramectin 10 mg/ml and the excipients ethyl oleate, sesame oil and butylhydroxyanisole.

The product is supplied in 50 ml, 200 ml and 500 ml multi-dose Type II or Type III amber glass vials with chlorobutyl rubber stoppers and aluminium overcaps or in 250 ml multi-dose Type II amber glass vials with chlorobutyl rubber stoppers and aluminium overcaps.
The choice of formulation 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 at 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 dormamectin, an established active substance. 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.

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.

D.Control on Intermediate Products

Not applicable.

E.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 has been provided.

Batch analytical data from the proposed production site has been provided demonstrating compliance with the specification.

F.Stability

Stability data on the active substance has 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.

G.Other Information

Not applicable.

III SAFETY AND RESIDUES ASSESSMENT (PHARMACO-TOXICOLOGICAL)

III.A Safety Testing

Pharmacological Studies

The application is made in accordance with Article 13(1) of Directive 2001/82/EC, as amended (a generic application). It was confirmed that the formulation and manufacturing process for the product is identical to that of the reference products. As a result it was accepted that the product was bioequivalent to the reference products, Zearl 10 mg/ml Solution for Injection for Cattle and Sheep (VPA 10019/018/001 transferred to 10047/027/001) and Zearl 10 mg/ml Solution for Injection for Pigs (VPA 10019/061/001 transferred to 10047/028/001).

As this is a generic application according to Article 13(1), and bioequivalence with a reference product has been accepted, results of pharmacological tests are not required.

The pharmacological aspects of this product reflect those of the reference products.

Toxicological Studies

As this is a generic application according to Article 13(1), and bioequivalence with a reference product has been claimed, results of toxicological tests are not provided.

User Safety

The applicant provided a user safety assessment which showed that when used in accordance with label recommendations the product will not pose any greater risk to the user than the risks associated with use of the reference products, Zearl 10 mg/ml Solution for Injection for Cattle and Sheep and Zearl 10 mg/ml Solution for Injection for Pigs.
Warnings and precautions as listed on the product literature are similar to those of the reference products and are adequate to ensure safety of the product to users.

Ecotoxicity

The applicant provided a first phase environmental risk assessment (ERA) in compliance with the relevant guideline which showed that further assessment was required.
The applicant provided a targeted Phase II ERA. The outcome of the ERA indicates a potential risk to aquatic organisms (namely daphnids, following direct excretion scenario) and dung fauna. In order to address the identified risks for aquatic organisms and dung fauna the following risk mitigation measures are recommended:

The following text is proposed in section 4.5 of the SPC (special precautions for use):
Doramectin is very toxic to dung fauna and aquatic organisms and may accumulate in sediments.

The risk to aquatic ecosystems and dung fauna can be reduced by avoiding too frequent and repeated use of doramectin (and products of the same anthelmintic class) in cattle.
The risk to aquatic ecosystems will be reduced by keeping treated cattle away from water bodies for two to five weeks after treatment.

The following text should be included in section 5.3 of the SPC (environmental properties):
Like other macrocyclic lactones, doramectin has the potential to adversely affect non-target organisms. Following treatment, excretion of potentially toxic levels of doramectin may take place over a period of several weeks. Faeces containing doramectin excreted onto pasture by treated animals may reduce the abundance of dung feeding organisms which may impact on the dung degradation.
Doramectin is very toxic to aquatic organisms and may accumulate in sediments.
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

No residue depletion studies were conducted for sheep and pigs because the product that is the subject of the present application is identical in every respect (composition, manufacturing process) to the reference products. On this basis it was assumed that depletion of residues from target tissues will be identical. Consequently, exemption from the requirement to present confirmatory residue data was justified and the authorised withdrawal period for the reference product can be applied to the generic product for sheep and pigs.

In the case of cattle, a number of residue depletion studies were presented in support of the withdrawal period. Given the nature of the data it was considered appropriate to accept the meat withdrawal period at 70 days.

MRLs

Doramectin is listed in Table I of the Annex to Commission Regulation (EU) No 37/2010 as follows:

	All mammalian food producing species
Muscle	40 µg/kg
Liver	100 µg/kg
Kidney	60 µg/kg
Fat / skin	150 µg/kg

Withdrawal Periods

Cattle:
Based on the data provided above, a withdrawal period of 70 days for meat in cattle is justified. The product is not to be used in lactating cows used to produce milk for human consumption, or in dry cows or pregnant dairy heifers within 60 days prior to calving.

Sheep:

Based on the information provided above, a withdrawal period of 70 days for meat in sheep is justified. The product is not to be used in dry dairy ewes including pregnant dairy ewes within 70 days prior to calving.

Pigs:
Based on the information provided above, a withdrawal period of 77 days for meat in pigs is justified.

IV. CLINICAL ASSESSMENT

As this is a generic application according to Article 13(1), and bioequivalence with a reference product has been accepted, efficacy studies are not required. The efficacy claims for this product are equivalent to those of the reference product.

IV.A Pre-Clinical Studies

Tolerance in the Target Species of Animals

The product literature accurately reflects the type and incidence of adverse effects which might be expected.

Resistance

Adequate warnings and precautions in relation to the prudent use of anthelmintic products appear on the product literature.

IV.B Clinical Studies

As this is a generic application according to Article 13(1), and bioequivalence with a reference product has been claimed, efficacy studies are not provided. 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 benefit/risk profile for the target species is favourable and the quality and safety of the product for humans and the environment is acceptable.

VI. 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 HPRA website.

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.

Changes:

Quality Changes

Summary of change **(Application number)**	**Approval date**
A.2.B Change in invented name of the medicinal product IE/V/0260/001/IB/001 B.II.e.1.b.2 Change in the immediate packaging of the finished product IE/V/xxxx/WS/018	26th July 2012 4th August 2020

Safety/Efficacy Changes

Summary of change **(Application number)**	**Approval date**
C.I.1 Change(s) in the Summary of Product Characteristics, Labelling or Package Leaflet intended to implement the outcome of a Union referral procedure IE/V/0260/1/IA/006 This variation is submitted to amend SPC, PL and Labels in compliance to the CVMP opinion on the Article 35 Referral procedure EMEA/V/A/81. The European Commission adopted the implementing decision on September 9, 2013. In particular, the modifications affect the withdrawal period duration for pigs (increased from 56 to 77 days).	13th December 2013