MEB agency / Veterinary Medicinal Products Unit
The Netherlands

College ter Beoordeling van Geneesmiddelen (CBG)
Medicines Evaluation Board (MEB)

Graadt van Roggenweg 500
3531 AH Utrecht The Netherlands

P.O. Box 8275
3503 RG Utrecht

DECENTRALISED PROCEDURE

PUBLICLY AVAILABLE ASSESSMENT REPORT FOR A VETERINARY MEDICINAL PRODUCT

Tralieve 50 mg/ml solution for injection for dogs

January 2018

MODULE 1

PRODUCT SUMMARY

Marketing Authorisation number (Dutch)	REG NL 120432
EU Procedure number	NL/V/0226/001/DC
Name, strength and pharmaceutical form	Tralieve 50 mg/ml, solution for injection.
Marketing Authorisation Holder (MAH)	Le Vet Beheer B.V. Wilgenweg 7 3421 TV Oudewater The Netherlands
Active substance(s)	Tramadol hydrochloride
ATC Vetcode	QN02AX02.
Target species	Dogs
Indication for use	For the reduction of mild postoperative pain.

MODULE 2

The Summary of Product Characteristics (SPC) for this product is available on the website:

http://www.cbg-meb.nl/CBG/en/veterinary-medicines/database-veterinary-medicines/default.htm

MODULE 3

PUBLIC ASSESSMENT REPORT

Legal basis of original application	Application in accordance with Article 13(1) of Directive 2001/82/EC as amended.
Date of completion of the original Decentralised Procedure	22 November 2017.
Concerned Member States for original procedure	AT, BE, BG, CY, CZ, DE, DK, EE, EL, ES, FI, FR, HR, HU, IE, IS, IT, LT, LU, LV, NO, PL, PT, RO, SE, SI, SK, UK.

1. SCIENTIFIC OVERVIEW

Tralieve 50 mg/ml, solution for injection for dogs 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 specie.
Tralieve 50 mg/ml, solution for injection for dogs is safe for the user 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.

2. QUALITY ASPECTS

A. Composition

The proposed product is an aqueous veterinary medicinal product for parenteral administration. The product contains 50 mg/mL tramadol hydrochloride as active substance and the following excipients: sodium acetate trihydrate, benzyl alcohol and water for injections. Hydrochloric acid (diluted) and sodium hydroxide are used for pH adjustment.

The product is packed in clear type I glass vials of 10, 20 and 50 mL, closed with dark grey coated rubber stoppers (Omniflex Plus) and aluminium caps. The glass vials and stoppers are in conformity with Ph. Eur. requirements.

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.

The product is manufactured using conventional manufacturing techniques. Process validation results for two 50 L small scale production batches, including all fill volumes, have been provided. Process validation results for one full scale production batch (500 L) will be made available for verification post authorisation by the supervisory authority. The tests performed during production are described.

C. Control of Starting Materials

The active substance is tramadol hydrochloride, an established active substance described in the European Pharmacopoeia (monograph 1681). For the active substance the CEP procedure is followed. 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.

No materials of animal origin are contained or used in the manufacturing process of the veterinary medicinal 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 the corresponding acceptance criteria are considered acceptable.

Satisfactory validation data for the analytical methods have been provided.

Batch analytical data from the proposed production site have been submitted, demonstrating compliance with the finished product specification.

F. Stability

No re-test period of the active substance is indicated on the CEP. The applicant has submitted stability data, demonstrating the stability of the active substance for 5 years (i.e. 60 months) when stored in double LDPE bags enclosed with trilaminated aluminium pouch and kept in HDPE drums.

Stability data on the finished product have been provided up to 18 months of storage. Based on the submitted stability data for the drug product, the proposed shelf-life of 30 months and the proposed in-use shelf-life of 56 days (i.e. 8 weeks) can be granted.

G. Other Information

Not applicable.

3. SAFETY AND RESIDUES ASSESSMENT (PHARMACO-TOXICOLOGICAL)

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

Warning statements and precautions as listed in the product literature are based on those of the reference product and supplemented with additional statements, based on increased knowledge and the current state of science. This information is considered adequate to ensure safety of the product to users and the environment.

3.A User Safety

User Safety

Being a generic procedure the applicant refers to the reference product for information on this section. Additionally, the applicant has provided a user safety assessment. Combined with increased knowledge and the current state of science, warning statements and precautions have been added to the product literature, ensuring safety to users of the product.

Ecotoxicity

Phase I
The environmental risk assessment can stop in Phase I, because the product will be used only in non-food animals.

Conclusion
Based on the data provided, the ERA can stop at Phase I. The product is not expected to pose an unacceptable risk for the environment when used according to the SPC.

4. CLINICAL ASSESSMENT (EFFICACY)

As this is a generic application according to Article 13, and bioequivalence with a reference product has been demonstrated, efficacy studies are not required. The efficacy claims for this product are based on increased knowledge and the current state of science.

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 specie is favourable and the quality and safety of the product for humans and the environment is acceptable.