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MUTUAL RECOGNITION PROCEDURE

PUBLICLY AVAILABLE ASSESSMENT REPORT FOR A VETERINARY MEDICINAL PRODUCT

MENBUTIL 100 MG/ML SOLUTION FOR INJECTION FOR CATTLE, PIGS, HORSES, SHEEP, GOATS

JULY 2017

MODULE 1

PRODUCT SUMMARY

EU Procedure number	FR/V/0200/001/E/001
Name, strength and pharmaceutical form	MENBUTIL 100 MG/ML SOLUTION FOR INJECTION FOR CATTLE, PIGS, HORSES, SHEEP, GOATS
Applicant	ANIMEDICA GMBH IM SÜDFELD 9 D-48308 SENDEN-BÖSENSELL GERMANY
Active substance(s)	Menbutone
ATC Vetcode	QA05AX90
Target species	Cattle, pigs, horses, sheep and goats.
Indication for use	Stimulation of hepato-digestive activity in case of digestive disorders and hepatic insufficiency.

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	Generic application in accordance with Article 13.1 of Directive 2001/82/EC as amended.
Date of completion of the original decentralised procedure	27 May 2009
Date product first authorised in the Reference Member State (MRP only)	
Concerned Member States for original procedure	AT DE ES HU PL RO

I. SCIENTIFIC OVERVIEW

The product has been developed as a generic of GENABILINE. The reference product has been authorised in FR since July 2009.
The proposed and reference products were considered bioequivalent.

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. Composition

The product contains 100 mg/ml of menbutone and the following excipients: chlorocresol, sodium metabisulfite (E223), edetic acid, ethanolamine 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.

C. Control of Starting Materials

The active substance is menbutone, 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.

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.
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 13.1, and bioequivalence with a reference product has been demonstrated, results of pharmacological tests are not required.

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.

User Safety

The applicant has provided a user safety assessment in compliance with the relevant guideline which shows that the risk for the user when the product is used as recommended, is very low.
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. No special warnings are therefore 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

No residue depletion studies were conducted because:

- the product is a generic medicinal product,
- all the excipients are listed in Table 1 of MRL Regulation 470/2009,
- the product contains the same qualitative and quantitative composition, as the reference product GENABILINE and is administered by the same route at the same dose,
- the product is typically administered to individual animals not intended for immediate slaughter,
- Menbutone is rapidly eliminated (around 8 hours after parenteral administration),
- an ADI of 60 µg/kg (i.e. 3.6 mg/person) has been established.

Considering this, the EMEA Summary report goes on to state “the assumed intake after 48 hours constitutes an insignificant fraction of the established ADI.

MRLs

Menbutone is listed in Table 1 of MRL Regulation 37/2010.

MRLs are listed below:

Marker residue	Animal species	MRL	Target tissues	Other provisions	Therapeutic classification
Not applicable	Bovine Ovine Caprine Porcine Equidae	No MRL required	Not applicable	No entry	No entry

All excipients of the product were listed in Table I of MRL Regulation 37/2010

Withdrawal Periods

The withdrawal periods agreed for the reference product can be applied to the MENBUTIL as follows:

Species	Tissues	**Withdrawal periods**
All target species	Meat & offal	Zero days
	Milk	Zero days

IV. CLINICAL ASSESSMENT (EFFICACY)

IV.A Pre-Clinical Studies

Pharmacology

The product is parenterally administered as a solution and contains the same active substance and excipients in the same concentration as the reference product GENABILINE. It is the exemption 4b of guidelines EMEA/CVMP/016/00. Therefore no bioequivalence study was required.

Tolerance in the Target Species of Animals

The product is parenterally administered as a solution and contains the same active substance and excipients in the same concentration as the reference product GENABILINE. Therefore no tolerance study was required

IV.B Clinical Studies

The product is parenterally administered as a solution and contains the same active substance and excipients in the same concentration as the reference product GENABILINE 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 benefit/risk profile for the target species is favourable and the quality and safety of the product for humans and the environment is acceptable.