ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

KETOPROCEN 100 mg/ml SOLUTION FOR INJECTION

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each ml contains:

Active substance:

Ketoprofen 100 mg

Excipients:

Benzyl alcohol (E1519) 10 mg

For the full list of excipients, see section 6.1.

3. PHARMACEUTICAL FORM

Solution for injection

Clear yellowish solution.

4. CLINICAL PARTICULARS

4.1 Target species

Horses, bovine and porcine.

4.2 Indications for use, specifying the target species

In horses: treatment of inflammatory and painful states of the osteoarticular and musculoskeletal systems of sport and race horses, in particular: limps of traumatic origin, arthritis, arthrosis, articular injuries (sprains, synovitis), fractures, tendonitis, peritendinitis, hoof complaints (navicular disease, shoe accidents, pododermatitis circumscripta, founder), post-surgical inflammations. Symptomatic treatment of colic.

In bovine: anti-inflammatory, analgesic and anti-pyretic treatment in:

Musculoskeletal inflammatory processes.
Mastitis.
Mammary oedema.
Inflammatory processes associated to respiratory diseases.
Colic.

In porcine: treatment of hyperthermia in acute diseases. In sows: treatment of Mastitis-Metritis-Agalactia syndrome.

4.3 Contraindications

As with all non-steroidal anti-inflammatory drugs, the administration of ketoprofen is counter-indicated: in cases of severe renal insufficiency, in association with other non-steroidal anti-inflammatory drugs, with diuretics or with anticoagulants.

Do not use in animals that have previously shown hypersensitivity to ketoprofen.

4.4 Special warnings for each target species

None.

4.5 Special precautions for use

Special precautions for use in animals

Do not mix with another substance in the same syringe.

Special precautions to be taken by the person administering the veterinary medicinal product to animals

People with known hypersensitivity to the active substance and/or benzyl alcohol should avoid contact with the veterinary medicinal product.

In case of accidental self-injection, seek medical advice immediately and show the package leaflet or the label to the physician.

Avoid splashes on the skin and eyes. Irrigate thoroughly with water should this occur. If irritation persists seek medical advice.

Wash hands after use.

4.6 Adverse reactions (frequency and seriousness)

Very rare, irritation or gastro-intestinal ulceration may appear.

4.7 Use during pregnancy, lactation or lay

The veterinary medicinal product can be used during pregnancy and lactation in cows and sows.

In the absence of specific data in pregnant mares and in very young foals, it is recommended that the product not be used in pregnant mares or in foals less than 15 days old.

4.8 Interaction with other medicinal products and other forms of interaction

Do not administer with other non-steroidal anti-inflammatory drugs, or with diuretics or anticoagulants.

4.9 Amounts to be administered and administration route

Horses:

Intravenous injection.

- For the treatment of musculoskeletal system and osteoarticular ailments, the dosage is 2.2 mg of ketoprofen per kg and day, i.e. 1 ml of product for each 45 kg of bodyweight, for 3 to 5 consecutive days.

- For the symptomatic treatment of colic, the dosage is 2.2 mg of ketoprofen per kg, i.e. 1 ml of product for each 45 kg of bodyweight, in a single injection.

Generally a single injection is sufficient: any additional injection should be preceded by a clinical re-assessment of the animal.

Bovine:

Dose of 3 mg/kg/day, for 1 to 3 consecutive days by intramuscular or intravenous injection for anti-inflammatory, analgesic and anti-pyretic treatment in:

Musculoskeletal inflammatory processes.
Mastitis.
Mammary oedema.
Inflammatory processes associated to respiratory diseases.
Colic.

Porcine:

Dose of 3 mg/kg/day by intramuscular injection for the treatment of hyperthermia in acute diseases and in sows for the treatment of Mastitis-Metritis-Agalactia syndrome.

4.10 Overdose (symptoms, emergency procedures, antidotes), if necessary

Anorexia, vomiting and diarrhoea may be identified.

4.11 Withdrawal period(s)

Horses: Meat: 4 days.

Milk: zero hours

Cattle: Meat: 4 days

Milk: zero hours

Pigs: Meat: 4 days.

5. PHARMACOLOGICAL PROPERTIES

Pharmacotherapeutic group: QM01AE03.

ATCvet code: antiinflammatory and antirheumatic products, non-steroids. Propionic acid derivatives.

5.1 Pharmacodynamic properties

Ketoprofen is a non-steroidal anti-inflammatory drug with analgesic properties belonging to the propionic acid group.

The inflammatory stimuli that damage the cell activate phospholipases which release arachidonic acid. This is the substrate of the cyclooxygenase and lipoxygenase enzyme systems, the activity of which results in the production of important inflammation mediators: prostaglandins, thromboxanes and leucotrienes.

Ketoprofen acts as a double inhibitor of inflammation, by inhibiting both the cyclooxygenase and lipoxygenase pathways, thus preventing the production of prostaglandins and leucotrienes.

Ketoprofen is also a powerful analgesic with effects at central and peripheral level. Its action consists of directly inhibiting bradykinin, a vasodilator and pain mediator.

Bradykinin initiates the pain impulse by stimulating the nerve endings of nociceptors.

In addition to its anti-bradykinin activity, ketoprofen also acts at central nervous system level to inhibition pain perception.

Furthermore, in horses ketoprofen counters the effect of endotoxins and acts an antagonist to intestinal spasm induced by bradykinin.

5.2 Pharmacokinetic particulars

Ketoprofen is rapidly absorbed. Maximum plasma concentration is reached in less than one hour after parenteral administration. Bioavailability is almost complete. Ketoprofen binds strongly to plasma proteins, allowing their accumulation in exudates at the inflamed sites.

The duration of the action is longer than would be expected from its plasma half-life, which varies from one to four hours depending on the species.

Ketoprofen passes into synovial fluid and stays there at higher levels than in plasma, with a half-life two or three times longer than the plasma half-life.

Ketoprofen is metabolized in the liver and 90 per cent is excreted through urine.

Studies of ketoprofen in horses have made it possible to establish that toxic doses were 55 mg/kg. Taking into account the proposed dosage (2.2 mg/kg), the therapeutic index of ketoprofen in horses is high (approximately 25).

Trials carried out using 2, 3 and 5 times the therapeutic dose in a significant number of horses have demonstrated the good general tolerance of the product. Likewise, local tolerance of the product is good (absence of phlebitis in perivenous injection and of inflammatory reaction in intramuscular injection).

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

- Benzyl alcohol (E1519)
- Arginine
- Citric acid monohydrate (E330)
- Water for injections

6.2 Incompatibilities

In the absence of compatibility studies, this veterinary medicinal product must not be mixed with other veterinary medicinal products.

6.3 Shelf life

Shelf life of the veterinary medicinal product as packaged for sale: 30 months

Shelf life after first opening the immediate packaging: 28 days

6.4. Special precautions for storage

Keep the vial in the outer carton in order to protect from light.

6.5 Nature and composition of immediate packaging

50 ml, 100 ml and 250 ml amber glass type II vials, with bromobutyl stopper and aluminium cap with FLIP-OFF seal.

Pack sizes:

- Box with 1 vial of 50 ml
- Box with 1 vial of 100 ml
- Box with 1 vial of 250 ml
- Box with 10 vials of 50 ml
- Box with 10 vials of 100 ml
- Box with 10 vials of 250 ml

Not all pack sizes may be marketed.

6.6 Special precautions for the disposal of unused veterinary medicinal product or waste materials derived from the use of such products

Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal product should be disposed of in accordance with local requirements.

7. MARKETING AUTHORISATION HOLDER

CENAVISA, S.L.

Camí Pedra Estela s/n
43205 Reus (SPAIN)

8. MARKETING AUTHORISATION NUMBER(S)

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of first authorization:

10 DATE OF REVISION OF THE TEXT

PROHIBITION OF SALE, SUPPLY AND/OR USE

Dispensing conditions: To be supplied only on veterinary prescription.

Administration conditions: Administration by a veterinarian surgeon (in case of intravenous route) or under their direct responsibility.

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON <THE OUTER PACKAGE> <AND> <THE IMMEDIATE PACKAGE>

Labels for 100 ml and 250 ml vials

Individual boxes for 50 ml, 100 ml and 250 ml vials

Boxes with 10 vials of 50 ml, 100 ml or 250 ml

○

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

KETOPROCEN 100 mg/ml SOLUTION FOR INJECTION

Ketoprofen

2. STATEMENT OF ACTIVE AND OTHER SUBSTANCES

Each ml contains:

Ketoprofen 100 mg

Benzyl alcohol (E1519) 10 mg

Other excipients q.s.

3. PHARMACEUTICAL FORM

Solution for injection

4. PACKAGE SIZE

Box with 1 vial of 50 ml

Box with 1 vial of 100 ml

Box with 1 vial of 250 ml

Box with 10 vials of 50 ml

Box with 10 vials of 100 ml

Box with 10 vials of 250 ml

5. TARGET SPECIES

Horses, bovine and porcine.

6. INDICATION(S)

Read the package leaflet before use.

7. METHOD AND ROUTE(S) OF ADMINISTRATION

Read the package leaflet before use.

8. WITHDRAWAL PERIOD

Horses: Meat: 4 days.

Milk: zero hours.

Cattle: Meat: 4 days

Milk: zero hours

Pigs: Meat: 4 days.

9. SPECIAL WARNING(S), IF NECESSARY

Read the package leaflet before use.

10. EXPIRY DATE

EXP {month/year}

Once opened, use by 28 days

11. SPECIAL STORAGE CONDITIONS

Keep the vial in the outer carton in order to protect from light.

12. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCTS OR WASTE MATERIALS, IF ANY

Dispose of waste material in accordance with local requirements.

13. THE WORDS "FOR ANIMAL TREATMENT ONLY" AND CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE, IF APPLICABLE

For animal treatment only. To be supplied only on veterinary prescription.

14. THE WORDS "KEEP OUT OF THE SIGHT AND REACH OF CHILDREN"

Keep out of the sight and reach of children.

15. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

CENAVISA, S.L.
Camí Pedra Estela s/n
43205 Reus (SPAIN)
Tel: 34 977 / 757 273 / Fax: 34 977 / 751 398
www.cenavisa.com / e-mail: cenavisa@cenavisa.com

16. MARKETING AUTHORISATION NUMBER(S)

17. MANUFACTURER'S BATCH NUMBER

Batch

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

Label for 50 ml vials

○

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

KETOPROCEN 100 mg/ml SOLUTION FOR INJECTION

Ketoprofen

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

100 mg/ml.

3. CONTENTS BY WEIGHT, BY VOLUME OR BY NUMBER OF DOSES

50 ml.

4. ROUTE(S) OF ADMINISTRATION

Intramuscular (bovine and porcine) or intravenous (horses and bovine).

5. WITHDRAWAL PERIOD

Horses: Meat: 4 days.

Milk: zero hours.

Cattle: Meat: 4 days

Milk: zero hours

Pigs: Meat: 4 days.

6. BATCH NUMBER

Batch

7. EXPIRY DATE

EXP {month/year}

Once opened, use by 28 days.

8. THE WORDS "FOR ANIMAL TREATMENT ONLY"

For animal treatment only.

B. PACKAGE LEAFLET

PACKAGE LEAFLET FOR:

KETOPROCEN 100 mg/ml SOLUTION FOR INJECTION

1. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER AND OF THE MANUFACTURING AUTHORISATION HOLDER RESPONSIBLE FOR BATCH RELEASE, IF DIFFERENT

Marketing authorisation holder and manufacturer responsible for batch release:

CENAVISA S.L.
Camí Pedra Estela s/n
43205 Reus (SPAIN)
Tel. 34 977 757 273 / Fax. 34 977 751 398

www.cenavisa.com / e-mail: cenavisa@cenavisa.com

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

KETOPROCEN 100 mg/ml SOLUTION FOR INJECTION

Ketoprofen

3. STATEMENT OF THE ACTIVE SUBSTANCE(S) AND OTHER INGREDIENT(S)

Each ml contains:

Active substance:

Ketoprofen 100 mg

Excipients:

Benzyl alcohol (E1519) 10 mg

Other excipients q.s.

4. INDICATION(S)

In horses: treatment of inflammatory and painful states of the osteoarticular and musculoskeletal systems of sport and race horses, in particular: limps of traumatic origin, arthritis, arthrosis, articular injuries (sprains, synovitis), fractures, tendonitis, peritendinitis, hoof complaints (navicular disease, shoe accidents, pododermatitis circumscripta, founder), post-surgical inflammations. Symptomatic treatment of colic.

In bovine: anti-inflammatory, analgesic and anti-pyretic treatment in:

Musculoskeletal inflammatory processes.
Mastitis.
Mammary oedema.
Inflammatory processes associated to respiratory diseases.
Colic.

In porcine: treatment of hyperthermia in acute diseases. In sows: treatment of Mastitis-Metritis-Agalactia syndrome.

5. CONTRAINDICATIONS

As with all non-steroidal anti-inflammatory drugs, the administration of ketoprofen is counter-indicated: in cases of severe renal insufficiency, in association with other non-steroidal anti-inflammatory drugs, with diuretics or with anticoagulants.

Do not use in animals that have previously shown hypersensitivity to ketoprofen.

6. ADVERSE REACTIONS

None known.

7. TARGET SPECIES

Horses, bovine and porcine.

8. DOSAGE FOR EACH SPECIES, ROUTE(S) AND METHOD OF ADMINISTRATION

Horses: Intravenous injection.

- For the treatment of musculoskeletal system and osteoarticular ailments, the dosage is 2.2 mg of ketoprofen per kg and day, i.e. 1 ml of product for each 45 kg of bodyweight, for 3 to 5 consecutive days.

- For the symptomatic treatment of colic, the dosage is 2.2 mg of ketoprofen per kg, i.e. 1 ml of product for each 45 kg of bodyweight, in a single injection.

Generally a single injection is sufficient: any additional injection should be preceded by a clinical re-assessment of the animal.

Bovine:

Dose of 3 mg/kg/day, for 1 to 3 consecutive days by intramuscular or intravenous injection for anti-inflammatory, analgesic and anti-pyretic treatment in:

Musculoskeletal inflammatory processes.
Mastitis.
Mammary oedema.
Inflammatory processes associated to respiratory diseases.
Colic.

Porcine:

Dose of 3 mg/kg/day, by intramuscular injection, for the treatment of hyperthermia in acute diseases and in sows for the treatment of Mastitis-Metritis-Agalactia syndrome.

9. ADVICE ON CORRECT ADMINISTRATION

Do not mix with another substance in the same syringe.

10. WITHDRAWAL PERIOD

Horses: Meat: 4 days.

Milk: zero hours.

Cattle: Meat: 4 days

Milk: zero hours

Pigs: Meat: 4 days.

11. SPECIAL STORAGE PRECAUTIONS

Keep out of the sight and reach of children.

Keep the vial in the outer carton in order to protect from light.

Do not use this veterinary medicinal product after the expiry date which is stated on the label.

Shelf life after first opening the container: 28 days.

12. SPECIAL WARNING(S)

Special warnings for each target species:

None.

Special precautions for use in animals:

Do not mix with another substance in the same syringe.

Special precautions to be taken by the person administering the veterinary medicinal product to animals:

People with known hypersensitivity to the active substance and/or benzyl alcohol should avoid contact with the veterinary medicinal product.

In case of accidental self-injection, seek medical advice immediately and show the package leaflet or the label to the physician.

Avoid splashes on the skin and eyes. Irrigate thoroughly with water should this occur. If irritation persists seek medical advice.

Wash hands after use.

Pregnancy and lactation:

The veterinary medicinal product can be used during pregnancy and lactation in cows and sows.

In the absence of specific data in pregnant mares and in very young foals, it is recommended that the product not be used in pregnant mares or in foals less than 15 days old.

Interaction with other medicinal products and other forms of interaction:

Do not administer with other non-steroidal anti-inflammatory drugs, or with diuretics or anticoagulants.

Overdose (symptoms, emergency procedures, antidotes):

Anorexia, vomiting and diarrhoea may be identified.

Incompatibilities:

In the absence of compatibility studies, this veterinary medicinal product must not be mixed with other veterinary medicinal products.

13. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCT OR WASTE MATERIALS, IF ANY

Medicines should not be disposed of via wastewater or household waste.

Ask your veterinary surgeon or pharmacist how to dispose of medicines no longer required. These measures should help to protect the environment.

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

15. OTHER INFORMATION

Pack sizes:

- Box with 1 vial of 50 ml

- Box with 1 vial of 100 ml
- Box with 1 vial of 250 ml
- Box with 10 vials of 50 ml
- Box with 10 vials of 100 ml
- Box with 10 vials of 250 ml

Not all pack sizes may be marketed.

For animal treatment only. To be supplied only on veterinary prescription.

Administration conditions: Administration by a veterinarian surgeon (in case of intravenous route) or under their direct responsibility.