

Summary of Product Characteristics

1 NAME OF THE VETERINARY MEDICINAL PRODUCT

Dinalgen 60 mg/ml solution for injection for pigs

2 QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 1 ml of the product contains:

Ketoprofen	60 mg
Benzyl alcohol (E1519)	10 mg

For a full list of excipients, see section 6.1.

3 PHARMACEUTICAL FORM

Solution for injection.

Clear colourless solution.

4 CLINICAL PARTICULARS

4.1 Target Species

Pigs.

4.2 Indications for use, specifying the target species

Pigs

Reduction of pyrexia in cases of respiratory disease and Postpartum Dysgalactia Syndrome/Mastitis, Metritis, Agalactiae (MMA syndrome) in sows, in combination with appropriate anti-infective therapy.

4.3 Contraindications

Do not use in animals where there is the possibility of gastro-intestinal ulceration or bleeding, in order not to aggravate their situation.

Do not use in animals suffering from cardiac, hepatic, or renal disease.

Do not use in dehydrated or hypovolemic or hypotensive animals due to the potential risk of increased renal toxicity.

Do not use in animals that have previously shown signs of hypersensitivity to ketoprofen or aspirin or any of the excipients.

Do not use where there is evidence of blood dyscrasia or blood coagulation disturbances

See also section 4.7

4.4 Special warnings for each target species

Since gastric ulceration is a common finding in PMWS (Post-weaning Multisystemic Wasting Syndrome), the use of ketoprofen in pigs affected by this pathology is not recommended, in order not to aggravate their situation.

4.5 Special precautions for use

Special precautions for use in animals

Do not exceed the recommended dose or duration of treatment

When administering to pigs of less than 6 weeks of age it is necessary to adjust the dose accurately as well as to perform a close clinical follow-up.

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

Avoid contact with the skin, eyes and mucous membranes.

In case of accidental skin, eye or mucous membrane contact, irrigate the affected area thoroughly with clean running water immediately. Seek medical advice if irritation persists.

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

Wash hands after use.

Hypersensitivity reactions (skin rash, urticaria) could occur. People with known hypersensitivity to the active substance should avoid contact with this product.

4.6 Adverse reactions (frequency and seriousness)

Intramuscular injection may be followed by transient irritation at the injection site.

The administration of ketoprofen in pigs at the recommended therapeutic dosage may cause superficial erosion and/or superficial ulceration of the gastrointestinal tract.

If side effects occur treatment must be stopped and the advice of a veterinarian should be sought.

4.7 Use during pregnancy, lactation or lay

The safety of ketoprofen has been investigated in pregnant laboratory animals (rats, mice, rabbits) and cattle. No adverse effects were noted. As the safety of ketoprofen has not been assessed in pregnant sows, the product should be used in this case only according to the benefit/risk assessment by the responsible veterinarian.

4.8 Interaction with other medicinal products and other forms of interaction

Interactions between Ketoprofen and the most commonly used antibiotics have not been investigated. Concurrent administration of diuretics or potentially nephrotoxic drugs should be avoided since there is an increase of renal disturbances. This is secondary to the diminished blood flow caused by the inhibition of prostaglandins.

This product must not be administered in conjunction with other NSAIDs or glucocorticosteroids since gastrointestinal ulceration may be exacerbated.

Pre-treatment with other anti-inflammatory substances may result in additional or increased adverse effects and accordingly a treatment-free period with such drugs should be observed for at least 24 hours before the commencement of treatment. The treatment-free period should, however, take into account the pharmacological properties of the products used previously.

Anticoagulants, particularly coumarin derivatives such as warfarin should not be used in combination with ketoprofen.

Ketoprofen is highly bound to plasma proteins and may compete with other highly bound drugs which can lead to toxic effects.

4.9 Amounts to be administered and administration route

Administer the product via the intramuscular route at a dose of 3mg/Kg bodyweight of ketoprofen, equivalent to 1ml/20Kg bodyweight of product, on a single occasion. Depending on the response observed and based on the benefit-risk analysis by the responsible veterinarian treatment may be repeated at intervals of 24 hours for a maximum of three treatments. Each injection should be given at a different site.

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

Overdose with NSAIDS can lead to gastro-intestinal ulceration, loss of proteins, hepatic and renal impairment. In tolerance studies performed in pigs with the product up to 25% of the animals treated at three times the maximum recommended dose (9 mg/kg) for three days or at the recommended dose (3 mg/kg) for triple the maximum recommended time (9 days) showed erosive and/or ulcerative lesions in both the aglandular (*pars oesophagica*) and glandular parts of the stomach. Early signs of toxicity include loss of appetite and pasty faeces or diarrhoea. If overdose symptoms are observed, symptomatic treatment should be initiated. The occurrence of ulcers is dose dependent to a limited extent.

4.11 Withdrawal Period(s)

Meat & offal: 3 days

5 PHARMACOLOGICAL or IMMUNOLOGICAL PROPERTIES

Pharmacotherapeutic group: Anti-inflammatory and Antirheumatic Products, Non-Steroids, Propionic acid derivatives, ATCvet code: QM01 AE 03

5.1 Pharmacodynamic properties

Ketoprofen, 2-(phenyl 3-benzoyl) propionic acid is a non-steroidal anti-inflammatory drug belonging to the arylpropionic acid group. Ketoprofen inhibits the biosynthesis of prostaglandins (PGE₂ and PGF₂ α) without affecting the ratio of PGE₂/PGF₂ α and thromboxanes. This mechanism of action results in its anti-inflammatory, anti-pyretic and analgesic activity. These properties are also attributed to its inhibiting effect on bradykinin and superoxide anions together with its stabilizing action on lysosomal membranes.

The anti-inflammatory effect is enhanced by the conversion of the (R)-enantiomer to (S)-enantiomer. It is known that the (S)-enantiomer supports the anti-inflammatory effect of ketoprofen.

5.2 Pharmacokinetic properties

After intramuscular administration ketoprofen is rapidly absorbed, having a high bioavailability and binding extensively to plasma proteins (>90%). Its elimination from plasma is rapid, although in the inflammatory exudate, it is more persistent. Ketoprofen is metabolized in liver and it is excreted mainly in urine and, to a lesser extent, in faeces. In pigs, following the intramuscular injection of a single dose of 3 mg/ketoprofen/kg body weight, the active drug substance is rapidly absorbed, reaching its average C_{max} in plasma (13 μ g/ml) between 0,5 and 1 hour (T_{max}) after initiation of the treatment. The bioavailability is high, of approximately 96%. Mean distribution volume is low (V_d=0.2 l/kg), and the average elimination half-life is short (T_{1/2}=2 h).

6 PHARMACEUTICAL PARTICULARS

6.1 List of excipients

L-arginine
Benzyl alcohol (E1519)
Citric acid anhydrous for pH adjustment
Water for injection

6.2 Incompatibilities

In the absence of compatibility studies, this veterinary medicinal product must not be mixed with other substances in the same syringe.

6.3 Shelf-life

Shelf-life of the veterinary medicinal product as packaged for sale: 60 months
Shelf-life after first opening the immediate packaging: 28 days

6.4 Special precautions for storage

This veterinary medicinal product does not require any special storage conditions.

6.5 Nature and composition of immediate packaging

Nature of the container:

Amber type II glass vials, closed with bromobutyl rubber stoppers and flip-off aluminium caps (100 ml) or aluminium caps (250 ml).

Presentations:

Vial containing 100 ml
Vial containing 250 ml
10 x vial containing 100 ml
10 x vial containing 250 ml

Not all pack sizes may be marketed.

6.6 Special precautions for the disposal of unused veterinary medicinal products or waste materials

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

7 MARKETING AUTHORISATION HOLDER

Laboratorios Dr. ESTEVE, S.A.
Avda. Mare de Déu de Montserrat, 221
08041 – Barcelona
Spain

8 MARKETING AUTHORISATION NUMBER(S)

VPA 10546/002/001

9 DATE OF THE FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

8th April 2009

10 DATE OF REVISION OF THE TEXT

December 2011