

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Enroxil Max 100 mg/ml solution for injection for cattle (CZ, IE, RO, SK, UK (NI))
Floxacril Max 100 mg/ml solution for injection for cattle (IT)

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each ml contains:

Active substance:

Enrofloxacin 100 mg.

Excipients:

Qualitative composition of excipients and other constituents	Quantitative composition if that information is essential for proper administration of the veterinary medicinal product
Benzyl alcohol (E1519)	20 mg
Butyl alcohol	30 mg
L-Arginine	
Water for injection	

Clear, yellow solution.

3. CLINICAL INFORMATION

3.1 Target species

Cattle.

3.2 Indications for use for each target species

Treatment of bovine respiratory disease associated with *Mannheimia haemolytica*, *Pasteurella multocida*, *Histophilus somni* and *Mycoplasma* spp. where clinical experience, supported where possible by sensitivity testing of the causal organism, indicates enrofloxacin as the drug of choice.

Treatment of local signs (inflammation, milk quality and yield) associated with peracute/acute mastitis in lactating dairy cattle caused by *E. coli*, where herd history and previous sensitivity testing indicate enrofloxacin as the drug of choice.

3.3 Contraindications

Do not use for prophylaxis.

Do not use in cases of hypersensitivity to the active substance or to any of the excipients.

3.4 Special warnings

None.

3.5 Special precautions for use

Special precautions for safe use in the target species:

Normal sterile precautions should be taken.

The safety of the veterinary medicinal product has not been established in calves when administered by the intravenous route and use of this route of administration in calves is therefore not recommended.

Official and local antimicrobial policies should be taken into account when the veterinary medicinal product is used.

Fluoroquinolones should be reserved for the treatment of clinical conditions which have responded poorly, or are expected to respond poorly, to other classes of antimicrobials.

Whenever possible, fluoroquinolones should only be used based on susceptibility testing.

Use of the veterinary medicinal product deviating from the instructions given in the SPC may increase the prevalence of bacteria resistant to the fluoroquinolones and may decrease the effectiveness of treatment with other quinolones due to the potential for cross resistance.

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

The veterinary medicinal product is an alkaline solution. Any spillage onto the skin should be washed off immediately with water.

In the event of accidental splash into the eye, rinse with copious amounts of clean water. If irritation occurs, seek medical advice.

Do not eat, drink or smoke whilst using the veterinary medicinal product.

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

People with known hypersensitivity to (fluoro)quinolones should avoid contact with the veterinary medicinal product.

Special precautions for the protection of the environment:

Not applicable.

3.6 Adverse events

Cattle:

Undetermined frequency (cannot be estimated from the available data):	Injection site reaction ¹
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¹Transient.

Reporting adverse events is important. It allows continuous safety monitoring of a veterinary medicinal product. Reports should be sent, preferably via a veterinarian, to either the marketing authorisation holder or its local representative or the national competent authority via the national reporting system. See the package leaflet for respective contact details.

3.7 Use during pregnancy, lactation or lay

Pregnancy and lactation:

Can be used during pregnancy and lactation.

3.8 Interaction with other medicinal products and other forms of interaction

None known.

3.9 Administration routes and dosage

Subcutaneous or intravenous use.

To ensure a correct dosage, body weight should be determined as accurately as possible.

Dosage and duration of treatment:

For respiratory infections in cattle: administer by subcutaneous injection:

A single dose of 7.5 mg/kg bodyweight (7.5 ml per 100 kg bodyweight)

Not more than 15 ml should be administered at one subcutaneous injection site.

For *E. coli* mastitis in cattle: administer by slow intravenous injection.

5.0 ml per 100 kg body weight (5 mg enrofloxacin per kg bodyweight) daily for 2 days.

3.10 Symptoms of overdose (and where applicable, emergency procedures and antidotes)

A dose of 25 mg/kg bodyweight administered for 15 consecutive days is tolerated without any clinical symptoms.

Clinical signs seen in gross overdosage include lethargy, lameness, ataxia, slight salivation and muscle tremors. In accidental overdose there is no antidote and treatment should be symptomatic.

3.11 Special restrictions for use and special conditions for use, including restrictions on the use of antimicrobial and antiparasitic veterinary medicinal products in order to limit the risk of development of resistance

Not applicable.

3.12 Withdrawal periods

Subcutaneous use:

Meat and offal: 14 days.

Milk: 84 hours.

Intravenous use:

Meat and offal: 4 days.

Milk: 72 hours.

4. PHARMACOLOGICAL INFORMATION

4.1 ATCvet code : QJ01MA90

4.2 Pharmacodynamics

It is bactericidal in action with activity against many Gram positive and Gram negative bacteria and mycoplasmas. The mechanism of action of the quinolones is unique among antimicrobials – they act primarily to inhibit bacterial DNA gyrase, an enzyme responsible for controlling the supercoiling of bacterial DNA during replication. Resealing of the double stranded helix is inhibited resulting in irreversible degradation of the chromosomal DNA. The fluoroquinolones also possess activity against bacteria in the stationary phase by an alteration of the permeability of the outer membrane phospholipid cell wall but are inactive against strict anaerobes.

Molecular resistance to fluoroquinolones has been observed to arise from two principal sources, (i) alteration to DNA gyrase or topoisomerase IV and (ii) alterations in drug permeability of the bacterial cell. Both mechanisms lead to a reduced susceptibility of the bacteria to fluoroquinolones. Clinical resistance is dependent on several mutations accumulating in a step-wise manner.

4.3 Pharmacokinetics

The pharmacokinetics of enrofloxacin is such that oral and parenteral administration leads to similar serum levels. Enrofloxacin is lipid soluble and amphoteric and possesses a high distribution volume. Tissue levels 2-3 higher than that found in the serum, have been demonstrated in laboratory animals and target species. Organs in which high levels can be expected are the lungs, liver, kidney, skin, bone and lymphatic system. Enrofloxacin also distributes into the cerebrospinal fluid, the aqueous humour and the foetus in pregnant animals.

After subcutaneous administration of 7.5 mg/kg the mean peak plasma concentration is 0.8 µg/ml achieved within 6 hours. Enrofloxacin is partly metabolized in the liver. Approximately 45 per cent of the dose is excreted in the urine and 55 per cent in the faeces as active and metabolites.

After an intravenous dose of 5 mg enrofloxacin per kg body weight (bw) to lactating dairy cattle, the total systemic exposure over the dosing interval of 24 h was at 7.1 mg*h/L. In cattle serum, approximately 30% of drug exposure (2.31 mg*h/L) consisted of ciprofloxacin, the active metabolite of enrofloxacin. The drug was well distributed into the body compartments (V_{enro} = 1.5 L/kg, V_{cipro} = 8.51 L/kg). Total body clearance was 0.71 L/h/kg.

In milk, most of drug activity consisted of ciprofloxacin. Overall drug concentrations peaked at 4.1 mg/kg two hours after treatment. Overall drug exposure over 24 h was 22.1 mg*h/L. The actives were eliminated from milk with a mean exposure half-life of 2.8 h.

5. PHARMACEUTICAL PARTICULARS

5.1 Major incompatibilities

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

5.2 Shelf life

Shelf life of the veterinary medicinal product as packaged for sale: 5 years.

Shelf life after first opening the immediate packaging: 28 days.

5.3 Special precautions for storage

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

Do not freeze.

5.4 Nature and composition of immediate packaging

Amber glass Type 2 multi-dose vials of 100 ml with bromobutyl rubber stopper.

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

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

Use take-back schemes for the disposal of any unused veterinary medicinal product or waste materials derived thereof in accordance with local requirements and with any national collection systems applicable to the veterinary medicinal product concerned.

6. NAME OF THE MARKETING AUTHORISATION HOLDER

7. MARKETING AUTHORISATION NUMBER(S)

8. DATE OF FIRST AUTHORISATION

Date of first authorisation: {DD/MM/YYYY}

9. DATE OF THE LAST REVISION OF THE SUMMARY OF THE PRODUCT CHARACTERISTICS

{MM/YYYY}

10. CLASSIFICATION OF VETERINARY MEDICINAL PRODUCTS

Veterinary medicinal product subject to prescription.

Detailed information on this veterinary medicinal product is available in the [Union Product Database \(https://medicines.health.europa.eu/veterinary\)](https://medicines.health.europa.eu/veterinary).

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

{BOX}

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Enroxil Max 100 mg/ml solution for injection

2. STATEMENT OF ACTIVE SUBSTANCES

Each ml contains 100 mg of enrofloxacin.

3. PACKAGE SIZE

100 ml

4. TARGET SPECIES

Cattle



5. INDICATIONS

6. ROUTES OF ADMINISTRATION

s.c., i.v.

7. WITHDRAWAL PERIODS

Withdrawal periods:

s.c.: Meat and offal: 14 days. Milk: 84 hours.

i.v.: Meat and offal: 4 days. Milk: 72 hours.

8. EXPIRY DATE

Exp. {mm/yyyy}

Once broached use within 28 days.

9. SPECIAL STORAGE PRECAUTIONS

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

Do not freeze.

10. THE WORDS “READ THE PACKAGE LEAFLET BEFORE USE”

Read the package leaflet before use.

11. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.

12. THE WORDS “KEEP OUT OF THE SIGHT AND REACH OF CHILDREN”

Keep out of the sight and reach of children.

13. NAME OF THE MARKETING AUTHORISATION HOLDER

14. MARKETING AUTHORISATION NUMBERS

15. BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

{LABEL}

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Enroxil Max 100 mg/ml solution for injection

2. STATEMENT OF ACTIVE SUBSTANCES

Each ml contains 100 mg of enrofloxacin.

3. TARGET SPECIES

Cattle



4. ROUTES OF ADMINISTRATION

Read the package leaflet before use.

s.c., i.v.

5. WITHDRAWAL PERIODS

Withdrawal periods:

s.c.: Meat and offal: 14 days. Milk: 84 hours.

i.v.: Meat and offal: 4 days. Milk: 72 hours.

6. EXPIRY DATE

Exp. {mm/yyyy}

Once broached use within 28 days.

7. SPECIAL STORAGE PRECAUTIONS

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

Do not freeze.

8. NAME OF THE MARKETING AUTHORISATION HOLDER

9. BATCH NUMBER

Lot {number}

B. PACKAGE LEAFLET

PACKAGE LEAFLET

1. Name of the veterinary medicinal product

Enroxil Max 100 mg/ml solution for injection for cattle

2. Composition

Each ml contains:

Active substance:

Enrofloxacin 100 mg

Excipients:

Benzyl alcohol (E1519) 20 mg

Butyl alcohol 30 mg

Clear, yellow solution.

3. Target species

Cattle.



4. Indications for use

Treatment of bovine respiratory disease associated with *Mannheimia haemolytica*, *Pasteurella multocida*, *Histophilus somni* and *Mycoplasma* spp. where clinical experience, supported where possible by sensitivity testing of the causal organism, indicates enrofloxacin as the drug of choice.

Treatment of local signs (inflammation, milk quality and yield) associated with peracute/acute mastitis in lactating dairy cattle caused by *E. coli*, where herd history and previous sensitivity testing indicate enrofloxacin as the drug of choice.

5. Contraindications

Do not use for prophylaxis.

Do not use in cases of hypersensitivity to the active substance or to any of the excipients.

6. Special warnings

Special precautions for safe use in the target species:

Normal sterile precautions should be taken.

The safety of the veterinary medicinal product has not been established in calves when administered by the intravenous route and use of this route of administration in calves is therefore not recommended.

Official and local antimicrobial policies should be taken into account when the veterinary medicinal product is used.

Fluoroquinolones should be reserved for the treatment of clinical conditions which have responded poorly, or are expected to respond poorly, to other classes of antimicrobials. Whenever possible, fluoroquinolones should only be used based on susceptibility testing. Use of the veterinary medicinal product deviating from the instructions given in the SPC may increase the prevalence of bacteria resistant to the fluoroquinolones and may decrease the effectiveness of treatment with other quinolones due to the potential for cross resistance.

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

The veterinary medicinal product is an alkaline solution. Any spillage onto the skin should be washed off immediately with water.

In the event of accidental splash into the eye, rinse with copious amounts of clean water. If irritation occurs, seek medical advice.

Do not eat, drink or smoke whilst using the veterinary medicinal product.

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

People with known hypersensitivity to (fluoro)quinolones should avoid contact with the veterinary medicinal product.

Pregnancy and lactation:

Can be used during pregnancy and lactation.

Overdose:

A dose of 25 mg/kg bodyweight administered for 15 consecutive days is tolerated without any clinical symptoms.

Clinical signs seen in gross overdosage include lethargy, lameness, ataxia, slight salivation and muscle tremors. In accidental overdose there is no antidote and treatment should be symptomatic.

Major incompatibilities:

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

7. Adverse events

Cattle:

Undetermined frequency (cannot be estimated from the available data):	Injection site reaction ¹
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¹Transient.

Reporting adverse events is important. It allows continuous safety monitoring of a product. If you notice any side effects, even those not already listed in this package leaflet, or you think that the medicine has not worked, please contact, in the first instance, your veterinarian. You can also report any adverse events to the marketing authorisation holder or its local representative using the contact details at the end of this leaflet, or via your national reporting system: {national system details}

8. Dosage for each species, routes and method of administration

Subcutaneous or intravenous use.

To ensure a correct dosage, body weight should be determined as accurately as possible.

Dosage and duration of treatment:

For respiratory infections in cattle: administer by subcutaneous injection (s.c.):
A single dose of 7.5 mg/kg bodyweight (7.5 ml per 100 kg bodyweight).

For *E. coli* mastitis in cattle: administer by slow intravenous injection (i.v.).
5.0 ml per 100 kg body weight (5 mg enrofloxacin per kg bodyweight) daily for 2 days.

9. Advice on correct administration

Not more than 15 ml should be administered at one subcutaneous injection site.

10. Withdrawal periods

Subcutaneous use:

Meat and offal: 14 days.
Milk: 84 hours.

Intravenous use:

Meat and offal: 4 days.
Milk: 72 hours.

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 freeze.

Do not use this veterinary medicinal product after the expiry date which is stated on the label and carton after Exp. The expiry date refers to the last day of that month.

Shelf life after first opening the immediate packaging: 28 days.

12. Special precautions for disposal

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

Use take-back schemes for the disposal of any unused veterinary medicinal product or waste materials derived thereof in accordance with local requirements and with any applicable national collection systems. These measures should help to protect the environment.

Ask your veterinary surgeon or pharmacist how to dispose of medicines no longer required.

13. Classification of veterinary medicinal products

Veterinary medicinal product subject to prescription.

14. Marketing authorisation numbers and pack sizes

Cardboard box with one 100 ml vial.

15. Date on which the package leaflet was last revised

{MM/YYYY}

Detailed information on this veterinary medicinal product is available in the [Union Product Database \(https://medicines.health.europa.eu/veterinary\)](https://medicines.health.europa.eu/veterinary).

16. Contact details

Marketing authorisation holder and contact details to report suspected adverse events:

Manufacturer responsible for batch release:

KRKA, d.d., Novo mesto, Šmarješka cesta 6, 8501 Novo mesto, Slovenia

VIRBAC

1^{ère} avenue 2065 m LID

06516 Carros

France

Local representatives and contact details to report suspected adverse events:

For any information about this veterinary medicinal product, please contact the local representative of the marketing authorisation holder.