

**PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGING WHERE THERE IS
NO PACKAGE LEAFLET, i.e. Combined label and package leaflet
COMPOSITE CAN, SECURITAINER AND BUCKET**

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Coldostin, 4800000 IU/g, powder for use in drinking water/milk.

2. COMPOSITION

Each gram contains:

Active substance:

Colistin sulfate 4 800 000 IU

White to off-white powder.

3. PACKAGE SIZE

100 gram, 1 kg

4. TARGET SPECIES

For cattle (calves), sheep (lambs), pigs, chickens and turkeys.

5. INDICATIONS FOR USE

Indications for use

Treatment and metaphylaxis of enteric infections caused by non-invasive *Escherichia coli* susceptible to colistin sulfate.

In the case of metaphylaxis, the presence of the disease in the group must be established before the veterinary medicinal product is used.

6. CONTRAINDICATIONS

Contraindications

Do not use in cases of hypersensitivity to colistin sulfate or to any of the excipients.

Do not use in cases of resistance to polymyxins.

Do not use in horses, particularly in foals, since colistin sulfate, due to a shift in the gastrointestinal microflora balance could lead to the development of antimicrobial associated colitis (Colitis X), typically associated with *Clostridium difficile*, which may be fatal.

7. SPECIAL WARNINGS

Special warnings

Special warnings:

As an adjunct to treatment, good management and hygiene practices should be introduced in

order to reduce the risk of infection and to control the potential build-up of resistance.

There is cross-resistance between colistin sulfate and polymyxin B.

Colistin sulfate exerts concentration-dependent activity against Gram-negative bacteria.

Following oral administration high concentrations are achieved in the gastrointestinal tract, i.e. the target site, due to the poor absorption of the substance. These factors indicate that a longer duration of treatment than the one indicated, leading to unnecessary exposure, is not recommended.

Special precautions for safe use in the target species:

Do not use colistin sulfate as a substitute for good management practices.

Colistin sulfate is a last resort drug in human medicine for treatment of infections caused by certain multi-drug resistant bacteria. In order to minimise any potential risk associated with widespread use of colistin sulfate, its use should be limited to treatment or treatment and metaphylaxis of diseases, and should not be used for prophylaxis.

Whenever possible, colistin sulfate should only be used based on susceptibility testing. An antibiotic with a lower risk of antimicrobial resistance selection (lower AMEG category) should be used for first line treatment where susceptibility testing suggests the likely efficacy of this approach.

Use of the veterinary medicinal product deviating from the instructions given in the leaflet may lead to treatment failures and increase the prevalence of bacteria resistant to colistin sulfate.

In the case of newborn animals and animals with severe gastrointestinal and renal disorders, systemic exposure to colistin sulfate may be increased. Neuro- and nephrotoxic alterations may occur.

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

People with known hypersensitivity to polymyxines, such as colistin sulfate, should avoid contact with the veterinary medicinal product.

If symptoms such as rash appear after exposure, seek medical advice immediately and show the package leaflet or the label to the physician. Swelling of the face, lips or eyes or respiratory difficulties are the most serious signs which require urgent medical attention.

The veterinary medicinal product may be irritating to the eyes, skin and mucous membranes.

When handling the veterinary medicinal product direct contact of the veterinary medicinal product with the skin, eyes and mucous membranes and inhalation of dust particles should be avoided.

Use the veterinary medicinal product in places with suitable ventilation.

The wearing of approved dust masks (either a disposable half-mask respirator conforming to European Standard EN 149 or a non-disposable respirator to European Standard EN 140 with a filter to EN 143), impervious gloves, overalls and safety glasses are recommended during the handling and mixing of this veterinary medicinal product.

Wash exposed skin after preparation. In case of accidental projection into the eyes, rinse abundantly with water.

Wash hands after use.

Wash your clothes daily after using the veterinary medicinal product.

Do not smoke, eat or drink when handling the veterinary medicinal product.

Pregnancy, lactation and lay:

The safety of the veterinary medicinal product has not been established during pregnancy, lactation or lay. Colistin sulfate is poorly absorbed after oral administration; therefore, the use of colistin sulfate during pregnancy, lactation or lay should not lead to particular problems. Use only accordingly to the benefit-risk assessment by the responsible veterinarian.

Interactions with other medicinal products and other forms of interaction:

After oral administration of colistin sulfate interaction with anaesthetics and myorelaxants may not be excluded in individual cases.

The combination with aminoglycosides and levamisole should be avoided.

The effects of colistin sulfate may be antagonized by binary cations (iron, calcium, magnesium) and by unsaturated fatty acids and polyphosphates.

Major incompatibilities:

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

8. ADVERSE EVENTS

Adverse events

None known.

Reporting adverse events is important. It allows continuous safety monitoring of a product. If you notice any side effects, even those not already listed on this label, 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 using the contact details at the end of this label, or via your national reporting system at:

Website: <https://www.gov.uk/report-veterinary-medicine-problem/animal-reacts-medicine>

e-mail: adverse.events@vmd.gov.uk

9. DOSAGE FOR EACH TARGET SPECIES, ROUTES AND METHOD OF ADMINISTRATION

Dosage for each species, routes and method of administration

Administration route: In drinking water/milk use.

Calves, lambs and pigs:

100 000 IU of colistin sulfate per kilogram body weight i.e. 1 g of veterinary medicinal product per 48 kg body weight daily for 3-5 consecutive days.

Chickens and turkeys:

75 000 IU of colistin sulfate per kilogram body weight i.e. 1 g of veterinary medicinal product per 64 kg body weight daily for 3-5 consecutive days.

Duration of treatment should be limited to the minimum time necessary for the treatment of the disease.

10. ADVICE ON CORRECT ADMINISTRATION

Advice on correct administration

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

Based on the recommended dose and the number and weight of animals to be treated, the exact daily concentration of the veterinary medicinal product should be calculated according to the following formula:

$$\frac{\text{..... mg veterinary medicinal product per kg body weight per day} \times \text{mean body weight (kg) of animals to be treated}}{\text{mean daily water consumption (litre per animal)}} = \text{..... mg veterinary medicinal product per litre of drinking water}$$

The uptake of medicated water depends on the physiological and clinical conditions of the animals. In order to obtain the correct dosage, the concentration of colistin sulfate has to be adjusted accordingly.

The use of suitably calibrated measuring equipment is recommended.

The veterinary medicinal product may be introduced via a water proportioner pump. Select the treatment dosage. Set the proportioner at the desired delivery rate. To prepare the stock solution, place the indicated quantity of veterinary medicinal product in a 10-litres container, fill with water and stir until dissolved. The maximum recommended concentration is 250 grams of veterinary medicinal product per 10 litres of drinking water and 500 mg of veterinary medicinal product per litre of milk(replacer).

The medicated water should be the only source of drinking water for the animals for the entire duration of the treatment period.

Water uptake should be monitored at frequent intervals.

After the end of the medication period, the water supply system should be cleaned appropriately to avoid intake of sub-therapeutic amounts of the active substance.

Medicated drinking water should be freshly prepared every 24 hours.

The medicated milk (replacer) should be used within 4 hours.

A levelled measuring spoon contains 3 grams of veterinary medicinal product.

11. WITHDRAWAL PERIODS

Withdrawal periods

Cattle (calves) and sheep (lambs):

Meat and offal: 1 day.

Pigs:

Meat and offal: 1 day.

Chickens and turkeys:

Meat and offal: 1 day.

Eggs: zero days.

12. SPECIAL STORAGE PRECAUTIONS

Special storage precautions

Keep out of the sight and reach of children.

Keep the original container tightly closed, in order to protect from light.

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

13. SPECIAL PRECAUTIONS FOR DISPOSAL

Special precautions for disposal

Medicines should not be disposed of via wastewater.

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 how to dispose of medicines no longer required.

14. CLASSIFICATION OF VETERINARY MEDICINAL PRODUCTS

Classification of veterinary medicinal products

Veterinary medicinal product subject to prescription.

15. MARKETING AUTHORISATION NUMBERS AND PACK SIZES

Vm 28365/4010

Pack sizes

- Can: 1 kg
- Securitainer: 100 g, 1 kg
- Bucket: 1 kg

Not all pack sizes may be marketed.

16 . PID LINK (Do not print heading)

[The following statement must be included where reference to the European Union Product Database is included on the product information. This statement is relevant to both UK(GB) and UK(NI) products:]

Find more product information by searching for the 'Product Information Database' on www.gov.uk.

17. CONTACT DETAILS

Contact details

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

Dopharma Research B.V.
Zalmweg 24
NL-4941 VX Raamsdonksveer
Tel. +31-162-582000
pharmacovigilance@dopharma.com

Manufacturer responsible for the batch release:

Dopharma B.V.
Zalmweg 24
NL-4941 VX Raamsdonksveer

18. OTHER INFORMATION

Other information

POM-V

19. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.

20. EXPIRY DATE

Exp. {mm/yyyy}

Once opened, use by ...

Shelf life after first opening the immediate packaging: 3 months.

Shelf life after dissolution in drinking water according to directions: 24 hours.

Shelf life after dissolution in milk/milk replacer according to directions: 4 hours.

21. BATCH NUMBER

Lot {number}

Gavin Hall
Approved: 11 February 2025