ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

{OUTSIDE OF THE TEAR OPEN LEAFLET / 1L BOTTLE or 5L CONTAINER}

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Methoxasol 20/100 mg/ml solution for use in drinking water.

2. STATEMENT OF ACTIVE SUBSTANCES

Each ml contains:
Trimethoprim 20 mg
Sulfamethoxazole 100 mg

3. PACKAGE SIZE

1 litre 5 litres

4. TARGET SPECIES

Pigs and chickens (broilers).

5. INDICATIONS

6. ROUTES OF ADMINISTRATION

In drinking water use.

7. WITHDRAWAL PERIODS

Withdrawal period:
Meat and offal: Pigs: 5 days
Broilers: 6 days
Not for use in birds producing or intended to produce eggs for human consumption.

8. EXPIRY DATE

Exp. {mm/yyyy}
Once opened use within 1 year. Use by: __/__/__
Shelf life after dilution or reconstitution according to directions: 24 hours.

9. SPECIAL STORAGE PRECAUTIONS

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

Eurovet Animal Health B.V.(or logo)

14. MARKETING AUTHORISATION NUMBERS

Vm 16849/3007

15. BATCH NUMBER

Lot {number}

B. PACKAGE LEAFLET

PACKAGE LEAFLET (INSIDE OF TEAR OPEN LEAFLET)

1. Name of the veterinary medicinal product

Methoxasol 20/100 mg/ml solution for use in drinking water for pigs and chickens.

2. Composition

Each ml contains:
Active substances:

Trimethoprim	20 mg
Sulfamethoxazole	100 mg

Excipients:

N-methyl pyrrolidone	690,8 mg

Clear yellow solution.

3. Target species

Pigs and chickens (broilers).

4. Indications for use

Pigs: Treatment and metaphylaxis of respiratory infections caused by *Actinobacillus pleuropneumoniae* susceptible to trimethoprim and sulfamethoxazole where the disease has been diagnosed in the herd.
Broilers: Treatment and metaphylaxis of respiratory infections caused by *Escherichia coli* susceptible to trimethoprim and sulfamethoxazole where the disease has been diagnosed in the flock.

5. Contraindications

Do not use in animals suffering from severe liver or kidney disease, oliguria or anuria.
Do not use in animals with impaired haematopoietic systems.
Do not use in cases of hypersensitivity to sulfonamides or trimethoprim or to any of the excipients.

6. Special warnings

Special warnings:
Severely diseased animals can have a decreased appetite and water consumption. If necessary the concentration of the VMP in the drinking water may need to be adjusted to make sure that the recommended dosage is being consumed. However if the concentration of the veterinary medicinal product is increased too much, the intake of the medicated drinking water decreases for palatability reasons. Therefore water intake should be monitored regularly, especially in broilers. In case of insufficient intake of water, pigs should be treated parenterally.

Special precautions for safe use in the target species:
Due to the likely variability (time, geographical) in susceptibility of bacteria for potentiated sulphonamides, occurrence of resistance of bacteria may differ from country to country and even from farm to farm, and therefore bacteriological sampling and susceptibility testing are recommended. Use of the veterinary medicinal product should be based on culture and sensitivity of micro-organisms from diseased cases on farm or from recent previous experience on the farm.
Use of the veterinary medicinal product deviating from the instructions given in the SPC may increase the prevalence of bacteria resistant to sulfamethoxazole and trimethoprim and may also decrease the effectiveness of combinations of trimethoprim with other sulphonamides due to the potential for cross resistance.
Official and local antimicrobial policies should be taken into account when the veterinary medicinal product is used.

Special precautions to be taken by the person administering the veterinary medicinal product to animals:
This veterinary medicinal product contains sulfamethoxazole, which can cause allergic reactions in some people.
People with known hypersensitivity (allergy) to sulphonamides or trimethoprim should avoid contact with the veterinary medicinal product.

During preparation and administration of the medicated drinking water, skin contact with the drug should be avoided. Personal protective equipment consisting of impermeable gloves e.g. rubber or latex should be worn when handling the veterinary medicinal product.
Do not handle this veterinary medicinal product if you know you are allergic to trimethoprim or sulphonamides.
Do not smoke, drink or eat when handling the veterinary medicinal product.
If you develop symptoms following exposure to the veterinary medicinal product such as skin rash, you should seek medical advice and show the package leaflet or the label to the physician. Swelling of the face, lips or eyes or difficulty with breathing are more serious symptoms and require urgent medical attention.

This veterinary medicinal product can cause eye-irritation.
Personal protective equipment consisting of safety glasses should be worn when mixing the veterinary medicinal product with drinking water. In the event of eye contact, rinse the eye with copious amounts of clean water and if irritation occurs, seek medical advice.

In the event of accidental ingestion, seek medical advice. Wash hands and contaminated skin immediately after handling the veterinary medicinal product.

Laboratory studies in rabbits and rats conducted with the excipient N-methyl pyrrolidone have shown evidence of foetotoxic effects. The veterinary medicinal product should not be administered by pregnant women and women suspected of being pregnant. Personal protective equipment consisting of gloves should be worn when handling the veterinary medicinal product by women of childbearing age.

Special precautions for the protection of the environment:
Not applicable.

Other precautions:
Not applicable.

Pregnancy and lactation or lay:
The safety of the veterinary medicinal product has not been established in pigs and chickens during pregnancy, lactation, lay or in animals intended for breeding.
Laboratory studies in rats conducted with trimethoprim have shown evidence of teratogenicity at higher doses than recommended therapeutic ones.
Laboratory studies in rabbits and rats conducted with the excipient N-methyl pyrrolidone have shown evidence of foetotoxic effects.
The use of the veterinary medicinal product is not recommended during pregnancy and lactation.

Interaction with other medicinal products and other forms of interaction:
Do not combine with other veterinary medicinal products

Overdose:
In pigs, a 2 ½ fold overdose induces no adverse reactions.
In chicken an acute overdose will not occur because the birds will be reluctant to drink the strongly concentrated drinking water (too bitter taste if above 2 litres of the veterinary medicinal product per 1000 litre drinking water). Chronic overdose in chicken will result in a strongly diminished water- and feed intake and retarded growth.

Special restrictions for use and special conditions for use:
Not applicable.

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

7. Adverse events

Pigs:

Rare (1 to 10 animals / 10,000 animals treated):	Hypersensitivity reaction.

Chickens:

Rare (1 to 10 animals / 10,000 animals treated):	Hypersensitivity reaction, decreased drinking

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

Route of administration: in drinking water use.

Pigs: 25 mg TMPS per kg bodyweight per day for 3-4 days, corresponding to 1 ml of the veterinary medicinal product per 4,8 kg bodyweight per day. This corresponds to approximately 1 litre of the veterinary medicinal product in 500 L drinking water.
Based on the recommended dose, daily water consumption, and the number and weight of the pigs to be treated, the exact daily concentration of the veterinary medicinal product should be calculated according to the following formula:

$$\frac{\text{Mean body weight (kg) of pigs to be treated}}{\text{Mean daily water consumption (l) per pig} \times 4.8} = \text{xx ml veterinary medicinal product per l drinking water}$$

Broilers: 33 mg TMPS per kg bodyweight per day for 3-4 days, corresponding to 1 ml of the veterinary medicinal product per 3,64 kg bodyweight per day. This corresponds to approximately 1 litre of the veterinary medicinal product in 750 L drinking water.
Based on the recommended dose, daily water consumption, and the number and weight of the birds to be treated, the exact daily concentration of the veterinary medicinal product should be calculated according to the following formula:

$$\frac{\text{Mean body weight (kg) of broilers to be treated}}{\text{Mean daily water consumption (l) per bird} \times 3.64} = \text{xx ml veterinary medicinal product per l drinking water}$$

One litre of the veterinary medicinal product weighs 1079 gram; therefore weight can also be used to measure the veterinary medicinal product quantity to be added in drinking water, according to the following formula:

Amount to be added in drinking water (g/L) = calculated ml/L x 1.079.

The daily amount is to be added to the drinking water such that all medication will be consumed in 24 hours.

9. Advice on correct administration

The veterinary medicinal product can be added directly to the drinking water to prepare a therapeutic solution at the calculated concentration, but can also be used in a concentrated stock solution by adding 200 ml of the veterinary medicinal product per litre water and diluting this further if required. Do not use this veterinary medicinal product undiluted or in higher concentrated stock solutions.
Medicated drinking water and stock solutions should be freshly prepared every 24 hours. During the treatment period animals should not have access to other water sources than the medicated water the dilution of which should be calculated to ensure that animals always have sufficient water available. To ensure a correct dosage body weight should be determined as accurately as possible to avoid underdosing. The intake of medicated water depends on the clinical condition of the animals. In order to obtain the correct dosage the concentration of the veterinary medicinal product may need to be adjusted accordingly.

10. Withdrawal periods

Meat and offal:
Pigs: 5 days
Broilers: 6 days
Eggs:
Not for use in birds producing or intended to produce eggs for human consumption.

11. Special storage precautions

Keep out of the sight and reach of children.

Do not freeze.

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.

Shelf life after first opening the immediate packaging: 1 year.
Shelf life after dilution or reconstitution according to directions: 24 hours.

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

Vm 16849/3007

Pack sizes:
1 litre bottle closed with a tamper-proof screw cap.
5 litre bottle closed with a tamper-proof screw cap.
Not all pack sizes may be marketed.

15. Date on which the package leaflet was last revised

{DD/MM/YYYY}

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

16. Contact details

Marketing authorisation holder and contact details to report suspected adverse reactions:
Eurovet Animal Health B.V.
Handelsweg 25
5531 AE Bladel
The Netherlands
Tel: +31 348 563434

Manufacturer responsible for batch release:
Genera Inc.
Svetonedeljska cesta 2
Kalinovica
10436 Rakov Potok
Croatia

Local representatives and contact details to report suspected adverse reactions:

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

17. Other information

Environmental properties:
Trimethoprim is persistent in soils.

Approved 15 December 2023