LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

Resealable 1.1 kg PE-Alu-PET bag

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pharmasin 100 % w/w Granules for Use in Drinking Water
for Pigs, Chickens, Turkeys and Calves

Tylosin (as Tylosin tartrate)

2. STATEMENT OF ACTIVE AND OTHER SUBSTANCES

Each 1.1 g of granules contains

1000 mg of tylosin (corresponding to 1100 mg of tylosin tartrate)

3. PHARMACEUTICAL FORM

Granules for use in drinking water

4. PACKAGE SIZE

1.1 kg

5. TARGET SPECIES

Pigs, chickens, turkeys and calves

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

Calves (meat and offal): 12 days
Pigs (meat and offal): 1 day
Turkeys (meat and offal): 2 days
Turkey (eggs): Zero days
Chickens (meat and offal): 1 day
Chicken (eggs): Zero days

9. SPECIAL WARNING(S), IF NECESSARY

Read the package leaflet for full instructions and user warnings before use

10. EXPIRY DATE

EXP {month/year}
Shelf life after dilution or reconstitution in water, milk or milk replacer: 24 hours
Shelf life after first opening: 3 months
Once broached, use by:

11. SPECIAL STORAGE CODITIONS

Store in the original container in order to protect from light.
Store below 30° C

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

Disposal: read package leaflet.

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

Huvepharma NV
Uitbreidingstraat 80
2600 Antwerpen
Belgium

16. MARKETING AUTHORISATION NUMBER(S)

17. MANUFACTURER'S BATCH NUMBER

Lot:

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

HDPE pot of 110 g

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pharmasin 100 % w/w Granules for Use in Drinking Water
for Pigs, Chickens, Turkeys and Calves

Tylosin (as Tylosin tartrate)

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

Each 1.1 g of granules contains

1000 mg of tylosin (corresponding to 1100 mg of tylosin tartrate)

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

110 g

4. ROUTE(S) OF ADMINISTRATION

Oral use.

5. WITHDRAWAL PERIOD

Calves (meat and offal): 12 days
Pigs (meat and offal): 1 day
Turkeys (meat and offal): 2 days
Turkey (eggs): Zero days
Chickens (meat and offal): 1 day
Chicken (eggs): Zero days

6. BATCH NUMBER

Lot:

7. EXPIRY DATE

EXP {month/year
Shelf life after dilution or reconstitution in water, milk or milk replacer: 24 hours
Shelf life after first opening: 3 months
Once broached, use by:

8. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.

B. PACKAGE LEAFLET

PACKAGE LEAFLET

Pharmasin 100 % w/w Granules for Use in Drinking Water
for Pigs, Chickens, Turkeys and Calves

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

Marketing authorisation

Huvepharma NV, Uitbreidingstraat 80, 2600 Antwerpen, Belgium

Manufacturer responsible for batch release

Biovet JSC, 39 Petar Rakov Str, 4550 Peshtera - Bulgaria

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

Pharmasin 100 % w/w Granules for Use in Drinking Water
for Pigs, Chickens, Turkeys and Calves
Tylosin (as Tylosin tartrate)

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

Each 1.1 g of granules contains

1000 mg of tylosin (corresponding to 1100 mg of tylosin tartrate)

White to light yellow coloured granules.

4. INDICATION(S)

Calves: Treatment and prevention of pneumonia caused by *Mycoplasma spp* when the disease has been established in the herd

Pigs:

- Treatment and prevention of enzootic pneumonia caused by *Mycoplasma hyopneumoniae* and *Mycoplasma hyorhinis* when the when the disease has been established in the herd.
- Treatment and prevention of Porcine Intestinal Adenomatosis (Ileitis) associated with *Lawsonia intracellularis* when the disease has been established in the herd.

Chickens:

- Treatment and prevention of chronic respiratory diseases (CRD) caused by *Mycoplasma gallisepticum* and *Mycoplasma synoviae* when the disease has been established in the flock

- Treatment and prevention of necrotic enteritis caused by *Clostridium perfringens* when the disease has been established in the flock

Turkeys: Treatment and prevention of infectious sinusitis caused by *Mycoplasma gallisepticum.* when the disease has been established in the flock.

5. CONTRAINDICATIONS

Do not use in animals with known hypersensitivity to tylosin or other macrolides,
Do not use in cases with known resistance to tylosin or cross-resistance to other macrolides (MLS-resistance).
Do not use in animals vaccinated with tylosin-sensitive vaccines either at the same time or within 1 week previously.
Do not use in animals with hepatic disorders.
Do not use in horses because of danger for inflammation of the caecum.

6. ADVERSE REACTIONS

In pigs, adverse reactions have been observed, including diarrhoea, pruritus, erythema of the skin, swelling of the vulva, rectal edema and prolapse. These reversible signs appeared 48-72 hours after start of treatment.
If you notice any serious effects or other effects not mentioned in this leaflet, please inform your veterinary surgeon.

7. TARGET SPECIES

Calves, pigs, chickens, turkeys

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

Oral administration through the drinking water
In calves the product can also be administered through milk or milk replacer.

1.1 gram of the veterinary medicinal product corresponds to 1 gram of tylosin. The dosages are as follows:

Calves:
10 – 20 mg tylosin per kg BW (corresponding to 11 – 22 mg of the veterinary medicinal product per kg BW), twice daily (corresponding to a daily dose of 20 – 40 mg tylosin per kg BW), for 7 - 14 days.

Turkeys:

75 – 100 mg tylosin per kg BW per day (corresponding to 82.5 – 110 mg of the veterinary medicinal product per kg BW) for 3 – 5 days.

Chickens:
For the treatment of chronic respiratory disease:
75 – 100 mg tylosin per kg BW per day (corresponding to 82.5 – 110 mg of the veterinary medicinal product per kg BW) for 3 – 5 days.

For the treatment of necrotic enteritis:
20 mg tylosin per kg BW per day (corresponding to 22 mg of the veterinary medicinal product) for 3 days.

Pigs:
For the treatment of enzootic pneumonia:
20 mg tylosin per kg BW per day (corresponding to 22 mg of the veterinary medicinal product per kg BW) for 10 days.
For the treatment of ileitis or PIA:
5 – 10 mg tylosin per kg BW per day (corresponding to 5.5 - 11 mg of the veterinary medicinal product per kg BW) for 7 days.

For the preparation of the medicated water/milk/milk-replacer the body weight of the animals to be treated and their actual daily water/milk/milk-replacer consumption should be taken into due account. Consumption may vary depending on factors like age, state of health, breed, husbandry system. To provide the required amount of active substance in mg per litre drinking water/milk/milk-replacer the following calculation should be made:

$$\frac{\text{…… mg tylosin per kg bodyweight per day} \times \text{Average bodyweight (kg) of the animals to be treated}}{\text{Average amount of drinking water or milk / animal (l)}} = \text{…...mg tylosin / l of drinking water}$$

If individual animals show signs of a serious infection such as a reduced water or feed intake, then they should be treated individually, such as by injection.

9. ADVICE ON CORRECT ADMINISTRATION

Sufficient access to the system of water supply should be available for the animals to be treated to ensure adequate water consumption. No other source of drinking water should be available during the medication period.

Should there be no clear response to treatment within 3 days the treatment approach should be reconsidered. 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 which might support development of resistance.

Medicated water, milk or milk replacer should be replaced every 24 hours.

10. WITHDRAWAL PERIOD

Calves (meat and offal): 12 days
Pigs (meat and offal): 1 day
Turkeys (meat and offal): 2 days
Turkey (eggs): Zero days

Chickens (meat and offal): 1 day
Chicken (eggs): Zero days

11. SPECIAL STORAGE PRECAUTIONS

Store in the original container in order to protect from light. Store below 30°C.

Shelf-life after dilution or reconstitution according to directions:

medicated water:24 hours
medicated milk or milk replacer:24 hours

Shelf-life after first opening the immediate packaging: 3 months
Do not use this veterinary medicinal product after the expiry date which is stated on the label.
"When the container is broached (opened) for the first time, using the in-use shelf-life which is specified on this label, the date on which any product remaining in the container should be discarded should be worked out. This discard date should be written in the space provided on the label."

Keep out of the sight and reach of children.

12. SPECIAL WARNING(S)

Special warnings for each target species
Due to likely variability (time, geographical) in susceptibility of bacteria to tylosin, bacteriological sampling and susceptibility testing are recommended.
Under-dosing and/or treating for an insufficient length of time are considered to promote the development of resistance in bacteria and should be avoided.

Interactions with other medicinal products and other forms of interaction
Lincosamides and aminoglycoside antibiotics antagonize the activity of tylosin.

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

Special precautions for use in animals
Animals with acute infections may have a reduced water and feed intake and should be treated with a suitable injectable veterinary medicinal product first.
Do not leave or dispose of water containing tylosin tartrate where it may be accessible to either animals not under treatment or wildlife.

User warnings
Tylosin may induce irritation. Macrolides, such as tylosin, may also cause hypersensitivity (allergy) following injection, inhalation, ingestion or contact with skin or eye. Hypersensitivity to tylosin may lead to cross reactions to other macrolides and vice versa. Allergic reactions to these substances may occasionally be serious and therefore direct contact should be avoided.
To avoid exposure during preparation of the medicated Tylosin may induce irritation. Macrolides, such as tylosin, may also cause hypersensitivity (allergy) following drinking water, wear overalls, safety glasses, impervious gloves and wear either a disposable half mask respirator conforming to European Standard EN149 or a non-disposable respirator to European Standard EN140 with a filter to EN143.

Wash hands after use.
In the event of accidental skin contact, wash thoroughly with soap and water. In case of accidental eye contact, flush the eyes with plenty of clean, running water. Do not handle the product if you are allergic to ingredients in the product.
If you develop symptoms following exposure, such as skin rash, you should seek medical advice and show the physician this warning. Swelling of the face, lips and eyes or difficulty in breathing are more serious symptoms and require urgent medical attention.

Use during pregnancy, lactation or lay
Laboratory studies in mice and rats have not produced any evidence of teratogenic, foetotoxic or maternotoxic effects. No studies have been conducted in the target species. Use only according to the benefit/risk assessment by the responsible veterinarian.

Overdose (symptoms, emergency procedures, antidotes)
There is no evidence of tylosin toxicity in rats, at dose rates of up to 1000 mg/kg by the oral route.There is no evidence of tylosin toxicity in chickens, turkeys, pigs or calves when administered orally at up to three times the recommended dose. Lincosamides and aminoglycoside antibiotics antagonize the activity of tylosin.

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

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

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

15. OTHER INFORMATION

1.1 kg resealable block bottom zipped sachet made of polyethylene /aluminium/polyethylene terephthalate laminate

110 g high density polyethylene pot with polypropylene cap

Not all pack sizes may be marketed.

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