ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

HDPE bottle

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

AMPHEN 200 mg/ml suspension for use in drinking water.

2. STATEMENT OF ACTIVE SUBSTANCES

1 ml contains:
Active substance:
Florfenicol 200 mg

3. PACKAGE SIZE

1 L

4. TARGET SPECIES

Pigs

5. INDICATIONS

6. ROUTES OF ADMINISTRATION

Oral use in drinking water.

7. WITHDRAWAL PERIODS

Withdrawal period: Meat and offal: 20 days

8. EXPIRY DATE

Exp {mm/yyyy}:

Once opened, use within 3 months.
Once diluted, use within 24 hours.

9. SPECIAL STORAGE PRECAUTIONS

Do not store above 25°C. Do not freeze. Protect from frost.

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

Huvepharma NV

14. MARKETING AUTHORISATION NUMBERS

Vm 30282/3006

15. BATCH NUMBER

Lot {number}:

B. PACKAGE LEAFLET

PACKAGE LEAFLET

1. Name of the veterinary medicinal product

AMPHEN 200 mg/ml suspension for use in drinking water for pigs

2. Composition

Each mL contains:

Active substance:

Florfenicol 200 mg

Excipient(s):

Qualitative composition of excipients and other constituents	Quantitative composition if that information is essential for proper administration of the veterinary medicinal product
Sodium benzoate	3.0 mg

White to almost white suspension for use in drinking water.

3. Target species

Pigs.

4. Indications for use

Treatment and metaphylaxis at the group level of swine respiratory diseases associated with *Actinobacillus pleuropneumoniae* and *Pasteurella multocida* .
The presence of the disease in the group must be established before the product is used.

5. Contraindications

Do not use in known cases of hypersensitivity to the active substance or to any of the excipients.
See also section Pregnancy and lactation for further information.

6. Special warnings

Special warnings:
Do not use the product with chlorinated water

The uptake of medication by animals may be altered as a consequence of illness. In case of insufficient water uptake, animals should be treated parenterally using a suitable injectable product prescribed by the veterinarian.

Resistance to florfenicol has been detected in *Salmonella typhimurium* and other foodborne pathogens.

There is cross-resistance between substances of the phenicol class. Additionally, other resistance genes have been identified that can be located on plasmids or transposons, such as the cfr gene, which confers cross-resistance between pleuromutilins, oxazolidinones, phenicols, streptogramin A, and lincosamides.

Special precautions for safe use in the target species:
In addition to medication, it is important to ensure proper husbandry conditions, including good hygiene, proper ventilation and avoiding crowded conditions.

Use of the product should be based on identification and susceptibility testing of the target pathogen(s). If this is not possible, therapy should be based on epidemiological information and knowledge of susceptibility of the target pathogens at farm level, or at local/regional level.

Use of the product should be in accordance with official, national and regional antimicrobial policies.

An antibiotic with a lower risk of antimicrobial resistance selection (lower AMEG[1]-category) should be used for first line treatment where susceptibility testing suggests the likely efficacy of this approach.

Not for use for prophylaxis.

The duration of treatment should not exceed 5 days.

Special precautions to be taken by the person administering the veterinary medicinal product to animals:
This veterinary medicinal product may cause hypersensitivity reactions.
People with known hypersensitivity to florfenicol or sodium benzoate should avoid contact with this veterinary medicinal product.

This veterinary medicinal product may be slightly irritating to the skin and eyes.
Avoid skin or eye contact, including hand-to-eye-contact.

This veterinary medicinal product may be harmful when ingested, including effects on male fertility. Avoid oral ingestion, including hand-to-mouth-contact when preparing the product. Do not eat, drink or smoke while handling this product.

Personal protective equipment consisting of gloves, clothing and safety glasses should be worn when handling and mixing the veterinary medicinal product.

In case of accidental spillage onto eyes, wash them immediately with water. In case of accidental spillage onto skin, wash immediately the affected area and take the contaminated clothes off.

Wash hands after use.

[1] Antimicrobial Advice Ad Hoc Expert Group category

If you develop symptoms following exposure such as skin rash, seek medical advice immediately and show the package leaflet or the label to the physician.

Special precautions for the protection of the environment:
The use of the veterinary medicinal product poses a risk to terrestrial organisms (plants) and to aquatic organisms (cyanobacteria), including groundwater organisms.

In order to prevent any adverse effects on terrestrial plants and algae and to prevent possible contamination of groundwater, manure from treated pigs must not be spread onto land without dilution with manure from untreated pigs. Manure from treated pigs must be diluted with at least 5 times the weight of manure from untreated pigs before it can be spread onto arable land or before the manure is traded.

Pregnancy and lactation:
The safety of the veterinary medicinal product has not been established during pregnancy or lactation.
The use in sows is not recommended during pregnancy and lactation
Laboratory studies in rats and mice have not revealed any evidence of potential embryotoxic or foetotoxic effect of florfenicol.

Fertility:
Do not administer to boars intended for breeding.

Interaction with other medicinal products and other forms of interaction:
See sub-section 'Special warnings' for further information.

Overdose:
In case of overdosing, a decrease in weight gain, food and water consumption, peri-anal erythema and oedema and modification of some haematological and biochemical parameters indicative of dehydration may be observed.

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:

Very common (>1 animal / 10 animals treated)	Diarrhoea Erythema[1] Oedema[1]
Undetermined frequency (cannot be estimated from the available data)	Decreased drinking Constipation Abnormal stool colouration[2] Rectal prolapse[3]

[1]perianal or rectal
[2]dark brown
[3]retroceding without treatment

Reporting adverse events is important. It allows continuous safety monitoring of a veterinary medicinal 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

first instance, your veterinarian. You can also report any adverse events to the marketing authorisation holder or the local representative of the marketing authorisation holder using the contact details at the end of this leaflet, or via your national reporting system.

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

In drinking water use.

The recommended dose is 10 mg florfenicol per kg bodyweight per day (equivalent to 5 mL of the product/100 kg bw) for 5 consecutive days.

9. Advice on correct administration

To ensure a correct dosage body weight should be determined as accurately as possible. In order to avoid under- and over-dosing, treated animals should be divided into groups of similar bodyweight and the dose should be calculated for each group individually.
Medicated water intake depends on many factors including the clinical condition of the animal as well as local conditions such as ambient temperature and humidity. All animals to be treated should have sufficient access to the water supply system to ensure adequate consumption of the medicated drinking water. In order to ensure consumption of the medicated water, animals should not have access to other water supplies whilst being treated. If, however, it is not possible to obtain a sufficient intake of medicated water, the animals should be treated parenterally.

The appropriate quantity of medicated water should be prepared based on the daily water consumption. In order to obtain the correct dosage, the water uptake has to be monitored and the concentration of florfenicol has to be adjusted accordingly.

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

$$\frac{\text{X mL veterinary medicinal product/ kg body weight/day} \times \text{Average body weight (kg) of animals to be treated}}{\text{Average daily water consumption (litre) per animal}} = \text{X ml veterinary product per litre drinking water}$$

If a weighing scale is used the required volume can be converted in grams as follows: quantity in g of product required per day = number of ml of product required per day x 1.075.
Accuracy of the dosing device should be thoroughly checked.

Shake the bottle vigorously for 60 seconds before use.
The product should be added to the water.
Prepare the solution with fresh potable water.

For water tanks:
The maximal solubility is reached at concentrations of 2 mL/L (0.4 g of florfenicol/L), 2.5 mL/L (0.5 g of florfenicol/L) and 3 ml/L (0.6 g of florfenicol/L) at respectively 4°C, 10°C and 20°C.
Solutions should be checked visually for complete dissolution.

To treat pigs drinking 10% of their bodyweight, at the dose of 10 mg/kg, add the veterinary medicinal product to the drinking water in the tank.
Use 1L of product for every 2000 L of water. This corresponds to a concentration of 0.10 g/L in drinking water.
Mix thoroughly, to achieve full dissolution the solution should be stirred vigorously using a manual whisk for 10 minutes. When using a magnetic stirrer at 100 rpm, the mixing time is 5 min.

For dosing pumps:
The veterinary medicinal product can only be used at the concentration of 50 mL/L i.e. 10 g of florfenicol per litre of water of stock solution.
To treat pigs drinking 10% of their bodyweight, at the dose of 10 mg/kg, add the veterinary medicinal product to the water in the reservoir of the dosing pump.
Add 1L of product to 20 L of unmedicated water. This corresponds to a concentration of 10 g/L in the stock solution.
Mix thoroughly using a manual whisk for 10 minutes until a homogeneous white milky suspension is obtained.
Set the dosing pump on 1 % and turn on the dosing pump.

Medicated drinking water should be replaced every 24 hours.

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.

10. Withdrawal periods

Meat and offal: 20 days.

11. Special storage precautions

Keep out of the sight and reach of children.
Do not store above 25 °C. Do not freeze. Protect from frost
Shelf-life after first opening the container: 3 months.
Shelf life after dilution according to directions: 24 hours.
Do not use the 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.

12. Special precautions for disposal

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

The veterinary medicinal product should not enter water courses as florfenicol may be dangerous for aquatic organisms (cyanobacteria), including groundwater organisms.

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 for the veterinary medicinal product concerned.

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 30282/3006

White rectangular HDPE bottle of 1 litre closed with a white PP tamper-evident screw cap with a LDPE-lined multiple-layer insert.

15. 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.

16. Contact details

Marketing authorisation holder <and contact details to report suspected adverse reactions>:
Huvepharma NV
Uitbreidingstraat 80
2600 Antwerp
Belgium

Manufacturer responsible for batch release:
Biovet JSC
39 Petar Rakov Str
4550 Pesthera
Bulgaria

<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

Florfenicol is toxic for terrestrial plants, cyanobacteria and groundwater organisms.

Approved: 07 June 2024

Gavin Hall