[Version 9.1,11/2024]

ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Coccibal, 200 mg/mL solution for use in drinking water for chickens and turkeys
DE: Eimeryl, 200 mg/mL solution for use in drinking water for chickens and turkeys

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each mL contains:

Active substance:
Amprolium ..... 200 mg
(equivalent to 226.2 mg Amprolium Hydrochloride)

Excipients:

Qualitative composition of excipients and other constituents	**Quantitative composition if that information is essential for proper administration of the veterinary medicinal product**
Sodium methyl parahydroxybenzoate (E 219)	1 mg
Sodium propyl parahydroxybenzoate	0.2 mg
Propylene glycol	
Purified water	

Clear yellow solution.

3. CLINICAL INFORMATION

3.1 Target species

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

3.2 Indications for use for each target species

Treatment of intestinal coccidiosis caused by *Eimeria* spp susceptible to amprolium.

3.3 Contraindications

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

3.4 Special warnings

As with any antiparasiticide, frequent and repeated use of antiprotozoal agents of the same class can lead to resistance development.
In case of detection a lack of efficacy during treatment, communicate it to the national competent authorities.

3.5 Special precautions for use

Special precautions for safe use in the target species:

The product is not intended for preventive use.

This product should be reserved for use in case of coccidiosis outbreaks due to non-availability of vaccine, in case of lack of efficacy of vaccine and in vaccinated flocks if a severe coccidial challenge is diagnosed before immunity has fully developed.

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

This product is acidic and may cause irritation to, or corrosion of, the skin, eyes, throat and airways.
Avoid all physical contact with the product, including any vapours.
Do not eat, drink or smoke whilst handling this product.
Personal protective equipment consisting of impervious gloves and protective glasses should be worn when handling the veterinary medicinal product. The selected protective gloves should satisfy the specifications of EU Directive 89/686/EEC and the standard EN 374 derived from it.
In the case of contact with skin or eyes, wash the affected area with clean running water immediately and remove any contaminated clothes. If irritation persists, seek medical advice and show the label to the physician.
In case of accidental ingestion, rinse the mouth with fresh water, seek medical advice immediately and show the label to the physician
People with known hypersensitivity to amprolium or to any of the excipients should avoid contact with the product.
Wash hands and exposed skin after use.

Special precautions for the protection of the environment:

Not applicable.

3.6 Adverse events

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

None known.

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

The safety of the veterinary medicinal product has not been established during pregnancy and lay.

Pregnancy:
Studies in laboratory animals have not produced any evidence of teratogenic effects.
Laying birds:
The safety of amprolium has not been investigated in laying birds.

Use only according to the risk/benefit assessment by the responsible veterinarian.

3.8 Interaction with other medicinal products and other forms of interaction

Amprolium is a thiamine analogue. Therefore, the efficacy of amprolium may be reduced during a simultaneous administration of products containing vitamin B-complex.

3.9 Administration routes and dosage

In drinking water use.

Posology for each target species is 20 mg amprolium / kg b.w. a day (corresponding to 1 mL of oral solution / 10 kg of bodyweight / day) for 5-7 consecutive days.
For the preparation of medicated water and to ensure a correct dosage, body weight should be determined as accurately as possible ad their actual daily water consumption should be taken into account. Consumption may vary depending on factors like age, state of health, breed, husbandry system. To provide the required amount of veterinary medicinal product in mLmL per litre drinking water the following calculation should be made:

$$\frac{\text{0.1 mL of veterinary medicinal product / kg body weight per day} \times \text{average body weight (kg) of animals to be treated} \times \text{number of animals}}{\text{Average daily water intake (l/animal)}} = \text{mL veterinary medicinal product per litre of drinking water}$$

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. 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.

The veterinary medicinal product must not come into contact with metal water pipes or metal containers.

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

Prolonged uses can produce thiamine deficiencies
This deficiency can be compensated by a thiamine intake.

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

Chickens and turkeys:
- Meat and offal: zero days.
- Eggs: zero days.

4. PHARMACOLOGICAL INFORMATION

4.1 ATCvet code:

QP51BX02.

4.2 Pharmacodynamics

Amprolium is an anticoccidial which belongs to the thiamine analogues family. Amprolium acts by interfering as a competitive antagonist of thiamine within thiamine transport mechanisms. It interferes in the carbohydrate metabolism required for coccidian multiplication and survival.
In in-vitro studies it was shown that the uptake of thiamine by schizonts of Eimeria tenella and by host intestinal cells can occur through passive diffusion or by an active, energy-and ph-dependent process.

Amprolium competitively inhibited both systems, however, the parasite was shown to be more sensitive to amprolium than the host.

As shown with Eimeria maxima inoculated chicken, the administration of Amprolium resulted in a proportion of morphologically abnormal macrogametes and oocysts which may be considered the reason for a reduced sporulation rate.

4.3 Pharmacokinetics

After oral administration absorption is low, reaching the maximum concentration 4 hours later. It is excreted mainly through faeces.

Environmental properties

Amprolium is persistent in soil.

5. PHARMACEUTICAL PARTICULARS

5.1 Major incompatibilities

No information is available on potential interactions or incompatibilities of this veterinary medicinal product administered orally by mixing into drinking water containing biocidal products, feed additives or other substances used in drinking water.

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

5.2 Shelf life

Shelf-life of the veterinary medicinal product as packaged for sale: 2 years.
Shelf-life after first opening the immediate packaging: 6 months.
Shelf-life after dilution or reconstitution according to directions: 24 hours.

5.3 Special precautions for storage

This veterinary medicinal product does not require any special storage conditions.

5.4 Nature and composition of immediate packaging

100 mL and 1 litre containers: white, opaque high density polyethylene bottles sealed by induction and with screw-on cap.
5 litres container: white, opaque high density polyethylene barrels sealed by induction and with screw-on cap.
Presentations: 1 L, 5 L, 12 x 1 L in cardboard box, 4 x 5 L in cardboard box, 10 x 100 mL in cardboard box with leaflet.

Not all pack sizes may be marketed.

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

S.P. VETERINARIA, S.A.

7. MARKETING AUTHORISATION NUMBER(S)

8. DATE OF FIRST AUTHORISATION

Date of first authorisation: {DD month 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).

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

Label for 1 L box, 5 L box, 100 mL box

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Coccibal, 200 mg/mL solution for use in drinking water

2. STATEMENT OF ACTIVE SUBSTANCES

Each mL contains:

Amprolium ..... 200 mg
(equivalent to 226.2 mg Amprolium Hydrochloride)

3. PACKAGE SIZE

1 L.
5 L.
100 mL.

4. TARGET SPECIES

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

5. INDICATIONS

6. ROUTES OF ADMINISTRATION

In drinking water use.

7. WITHDRAWAL PERIODS

Withdrawal periods:
Chickens and turkeys:
- Meat and offal: zero days.
- Eggs: zero days.

8. EXPIRY DATE

Exp. {mm/yyyy}

Once opened use within 6 months.
Once opened use by…

9. SPECIAL STORAGE PRECAUTIONS

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

S.P. VETERINARIA, S.A.

14. MARKETING AUTHORISATION NUMBERS

15. BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

Label for 1 L without box, 5 L without box

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Coccibal, 200 mg/mL solution for use in drinking water

2. STATEMENT OF ACTIVE SUBSTANCES

Each mL contains:

Amprolium ..... 200 mg
(equivalent to 226.2 mg Amprolium Hydrochloride)

3. PACKAGE SIZE

1 L.
5 L.

4. TARGET SPECIES

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

5. INDICATIONS

6. ROUTES OF ADMINISTRATION

In drinking water use.

7. WITHDRAWAL PERIODS

Withdrawal periods:
Chickens and turkeys:
- Meat and offal: zero days.
- Eggs: zero days.

8. EXPIRY DATE

Exp. {mm/yyyy}

Once opened use within 6 months.
Once opened use by…

9. SPECIAL STORAGE PRECAUTIONS

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

S.P. VETERINARIA, S.A.

14. MARKETING AUTHORISATION NUMBERS

15. BATCH NUMBER

Lot {number}

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

Label for 1 L with box, 5 L with box, 100 mL with box

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Coccibal, 200 mg/mL solution for use in drinking water

2. STATEMENT OF ACTIVE SUBSTANCES

Each mL contains:

Amprolium ..... 200 mg
(equivalent to 226.2 mg Amprolium Hydrochloride)

3. TARGET SPECIES

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

4. ROUTES OF ADMINISTRATION

In drinking water use.
Read the package leaflet before use.

5. WITHDRAWAL PERIODS

Withdrawal periods:
Chickens and turkeys:
- Meat and offal: zero days
- Eggs: zero days

6. EXPIRY DATE

Exp. {mm/yyyy}

Once opened use within 6 months.
Once opened use by…

7. SPECIAL STORAGE PRECAUTIONS

8. NAME OF THE MARKETING AUTHORISATION HOLDER

S.P.VETERINARIA, S.A.

9. BATCH NUMBER

Lot {number}

B. PACKAGE LEAFLET

PACKAGE LEAFLET

1. Name of the veterinary medicinal product

Coccibal, 200 mg/mL solution for use in drinking water for chickens and turkeys

2. Composition

Each mL contains:

Active substance:
Amprolium ..... 200 mg
(equivalent to 226.2 mg Amprolium Hydrochloride)

Excipients:

Sodium methyl parahydroxybenzoate (E 219).....1 mg
Sodium propyl parahydroxybenzoate.....0.2 mg

Clear yellow solution.

3. Target species

Chickens (broilers, pullets, layers, breeder hens) and turkeys.

4. Indications for use

Treatment of intestinal coccidiosis caused by *Eimeria* spp susceptible to amprolium.

5. Contraindications

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

6. Special warnings

Special warnings:

As with any antiparasiticide, frequent and repeated use of antiprotozoal agents of the same class can lead to resistance development.
In case of detection a lack of efficacy during treatment, communicate it to the national competent authorities.

Special precautions for safe use in the target species:

The product is not intended for preventive use.
This product should be reserved for use in case of coccidiosis outbreaks due to non-availability of vaccine, in case of lack of efficacy of vaccine and in vaccinated flocks if a severe coccidial challenge is diagnosed before immunity has fully developed.

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

This product is acidic and may cause irritation to, or corrosion of, the skin, eyes, throat and airways.
Avoid all physical contact with the product, including any vapours.
Do not eat, drink or smoke whilst handling this product.
Personal protective equipment consisting of impervious gloves and protective glasses should be worn when handling the veterinary medicinal product.The selected protective gloves should satisfy the specifications of EU Directive 89/686/EEC and the standard EN 374 derived from it.
In the case of contact with skin or eyes, wash the affected area with clean running water immediately and remove any contaminated clothes. If irritation persists, seek medical advice and show the label to the physician.
In case of accidental ingestion, rinse the mouth with fresh water, seek medical advice immediately and show the label to the physician
People with known hypersensitivity to amprolium or to any of the excipients should avoid contact with the product.
Wash hands and exposed skin after use.

Pregnancy:

The safety of the veterinary medicinal product has not been established during pregnancy and lay.
Studies in laboratory animals have not produced any evidence of teratogenic effects.

Laying birds:

The safety of amprolium has not been investigated in laying birds.
Use only according to the risk/benefit assessment by the responsible veterinarian.

Interaction with other medicinal products and other forms of interaction:

Amprolium is a thiamine analogue. Therefore, the efficacy of amprolium may be reduced during a simultaneous administration of products containing vitamin B-complex.

Overdose:

Prolonged uses can produce thiamine deficiencies.
This deficiency can be compensated by a thiamine intake.

<Special restrictions for use and special conditions for use:>

Major incompatibilities:

No information is available on potential interactions or incompatibilities of this veterinary medicinal product administered orally by mixing into drinking water containing biocidal products, feed additives or other substances used in drinking water.

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

7. Adverse events

Chickens (broilers, pullets, layers, breeder hens) and turkeys

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 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

In drinking water use.
Posology for each target species is 20 mg amprolium / kg b.w. a day (corresponding to 1 mL of oral solution / 10 kg of bodyweight / day) for 5-7 consecutive days.
For the preparation of medicated water and to ensure a correct dosage, body weight should be determined as accurately as possible ad their actual daily water consumption should be taken into account.Consumption may vary depending on factors like age, state of health, breed, husbandry system. To provide the required amount of veterinary medicinal product in mL per litre drinking water the following calculation should be made:

$$\frac{\text{mLmL0.1 mL of veterinary medicinal product / kg body weight per day} \times \text{average body weight (kg) of animals to be treated} \times \text{number of animals}}{\text{Average daily water intake (l/animal)}} = \text{mL veterinary medicinal product per litre of drinking water}$$

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. 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.

The veterinary medicinal product must not come into contact with metal water pipes or metal containers.

10. Withdrawal periods

Chickens and turkeys:
- Meat and offal: zero days.
- Eggs: zero days.

11. Special storage precautions

Keep out of the sight and reach of children.

This veterinary medicinal product does not require any special storage conditions

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: 6 months.
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

Package sizes: 100 ml, 1 L and 5 L
Presentations: 1 L, 5 L, 12 x 1 L in cardboard box, 4 x 5 L in cardboard box, 10 x 100 mL in cardboard box with leaflet.
Not all pack sizes may be marketed.

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).

16. Contact details

Marketing authorisation holder andmanufacturer responsible for batch release:

S.P. VETERINARIA, S.A.
Ctra. Reus-Vinyols, km 4,1
43330 Riudoms (Spain)
Tel. +34 977 850 170

<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.>

17. Other information

Environmental properties
Amprolium is persistent in soil.