LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

CARTON BOX 100 ml

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys
Colistin sulfate

2. STATEMENT OF ACTIVE SUBSTANCES

Colistin sulfate 5,000,000 IU/ml

3. PHARMACEUTICAL FORM

Concentrate for oral solution

4. PACKAGE SIZE

100 ml

5. TARGET SPECIES

Cattle (calves), pigs, sheep (lambs), chickens and turkeys

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

Oral use
In drinking water/milk use
Read the package leaflet before use.

8. WITHDRAWAL PERIOD

Withdrawal periods:
Cattle (calves), sheep (lambs) and pigs
Meat and offal: 1 day

Chickens and turkeys
Meat and offal: 1 day
Eggs: Zero days

9. SPECIAL WARNING(S), IF NECESSARY

10. EXPIRY DATE

EXP
Shelf-life after first opening the immediate container: 3 months.
Shelf life after reconstitution in water according to directions: 24 hours
Shelf life after reconstitution in milk according to directions: 6 hours
Once broached, use by…

11. SPECIAL STORAGE CONDITIONS

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

LIVISTO Int'l, S.L.
Av. Universitat Autònoma, 29
08290 Cerdanyola del Vallès
(Barcelona) Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 43173/4000

17. MANUFACTURER'S BATCH NUMBER

Batch

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE

LABEL 100 ml

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys
Colistin sulfate

2. STATEMENT OF ACTIVE SUBSTANCES

Colistin sulfate 5,000,000 IU/ml

3. PHARMACEUTICAL FORM

4. PACKAGE SIZE

100 ml

5. TARGET SPECIES

Cattle (calves), pigs, sheep (lambs), chickens and turkeys

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

Oral useIn drinking water/milk use
Read the package leaflet before use.

8. WITHDRAWAL PERIOD

Withdrawal periods:
Cattle (calves), sheep (lambs) and pigs
Meat and offal: 1 day

Chickens and turkeys
Meat and offal: 1 day
Eggs: Zero days

9. SPECIAL WARNING(S), IF NECESSARY

10. EXPIRY DATE

EXP
Shelf-life after first opening the immediate container: 3 months.
Shelf life after reconstitution in water according to directions: 24 hours
Shelf life after reconstitution in milk according to directions: 6 hours
Once broached, use by…

11. SPECIAL STORAGE CONDITIONS

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

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

LIVISTO Int'l, S.L.
Av. Universitat Autònoma, 29
08290 Cerdanyola del Vallès
(Barcelona) Spain

16. MARKETING AUTHORISATION NUMBER(S)

Vm 43173/4000

17. MANUFACTURER'S BATCH NUMBER

Batch

B. PACKAGE LEAFLET – 100 ml

PACKAGE LEAFLET:
Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys

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

Marketing authorisation holder:
LIVISTO Int'l, S.L.
Av. Universitat Autònoma, 29
08290 Cerdanyola del Vallès
(Barcelona) Spain

Manufacturer responsible for batch release:
Industrial Veterinaria S.A.
Esmeralda, 19
08950 Esplugues de Llobregat (Barcelona) Spain

aniMedica GmbH
Im Südfeld 9
48308 Senden-Bösensell Germany

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys
Colistin sulfate

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

Each ml contains:

Active substance:
Colistin sulfate 5,000,000IU

Excipients:
Benzyl alcohol (E1519) 10 mg

Clear orange-brown solution

4. INDICATION(S)

Calves, lambs, pigs, chickens, turkeys:
Treatment and metaphylaxis of entericinfections caused by non-invasive *E. coli* susceptible to colistin sulfate.The presence of the disease in the herd should be established before metaphylactic treatment.

5. CONTRAINDICATIONS

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

6. ADVERSE REACTIONS

None known.

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 inform your veterinary surgeon.

Alternatively you can report via your national reporting system {national system details}.

7. TARGET SPECIES

Cattle (calves), pigs, sheep (lambs), chickens and turkeys

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

Oral use.
In drinking water/milk use
Calves, lambs, pigs: 100 000 IU of colistin sulfate per kg body weight daily for 3-5 consecutive days in drinking water or milk (replacer) in calves, equivalent to 0.20 ml of the concentrate solution per 10 kg body weight per day for 3-5 days.

Chickens and turkeys: 75 000 IU of colistin sulfate per kg body weight daily for 3-5 consecutive days in drinking water, equivalent to 15 ml of the concentrate solution per Ton of body weight per day for 3-5 days.

Duration of treatment should be limited to the minimum time necessary for the treatment of the disease.
Any medicated water which is not consumed within 24 hours should be discarded.
Any medicated milk which is not consumed within 6 hours should be discarded.

9. ADVICE ON CORRECT ADMINISTRATION

Direct oral administration to individual animals
The recommended daily dose should be divided into two if the product is to be administered directly into the mouth of the animal.

Prior to direct oral administration, the product should be diluted with a volume of drinking water equivalent to 2.5 x the volume of product concentrate to be administered.

Administration via drinking water
The intake of medicated water depends on the clinical condition of the animals. In order to obtain the correct dosage the concentration of colistin sulfate has to be adjusted accordingly. Carefully calculate the average body weight to be treated and the average daily water consumption before each treatment.
Medicated water should be made every day, immediately prior to provision.
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.

With the following formula, we can calculate an exact dosage:

$$\frac{\text{…ml of the product per kg body weight and day} \times \text{Average body weight (kg)}}{\text{Average daily water intake (l/animal)}} = \text{…ml of the product per litre of drinking water}$$

• Administration without a dosing pump:
The treatment is distributed in a tank over a period of 24 hours, for 3-5 consecutive days.
The productis added to a volume of the drinking water corresponding to the volume consumed by the animals over the treatment period (24 hours) to achieve a dose of 100 000 IU of colistin sulfate per kg body weight for pigs, lambs and calves and 75 000 IU of colistin sulfate per kg body weight for chickens and turkeys.
• Administration via a dosing pump
The treatment is distributed over a period of 24 hours, for 3-5 consecutive days.
A dosing pump is used to add a stock solution at a pre-determined concentration to the drinking water.

10. WITHDRAWAL PERIOD

Cattle (calves), sheep (lambs) and pigs
Meat and offal: 1 day

Chickens and turkeys
Meat and offal: 1 day
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 container: 3 months.
Shelf life after reconstitution in water according to directions: 24 hours
Shelf life after reconstitution in milk according to directions: 6 hours

12. SPECIAL WARNING(S)

Special warnings for each target species:
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.
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 treatmentthan the one indicated in section 8 of this package leaflet, leading to unnecessary exposure, is not recommended.

Special precautions for use in animals:
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-drugresistant 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.
Use of the product deviating from the instructions given in the leaflet may leadto treatment failures and increase the prevalence of bacteria resistant to colistin sulfate. There is cross-resistance between colistin sulfate and polymyxin B.
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 polymyxins, such as colistin sulfate, should avoid contact with the veterinary medicinal product.
Avoid direct contact with skin and eyes while handling the product. Personal protective equipment consisting of gloves and protective goggles should be worn when handling and dosing the veterinary medicinal product.
Wash splashes from skin immediately with soap and plenty of water.
In case of accidental eye exposure, wash with plenty of water and seek medical attention immediately and show the label to the physician.
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 or eyes or difficulty in breathing are more serious symptoms and require urgent medical attention.
Wash hands after use.

Use during pregnancy, lactation or lay:
The safety of the veterinary medicinal product has not been established during pregnancy, lactation or lay. However, 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 according to the benefit-risk assessment by the responsible veterinarian.

Interaction with other medicinal products and other forms of interaction:
After oral administration of colistin sulfate interaction with anaesthetics (curarimimetic agents) 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.

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

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

Medicines should not be disposed of via wastewater or household waste. Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal product should be disposed of in accordance with local requirements.
Ask your veterinary surgeon or pharmacist how to dispose of medicines no longer required. These measures should help to protect the environment.

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

January 2021

15. OTHER INFORMATION

The active ingredient colistin sulfate is very persistent in soils.

Package sizes:
Box with a bottle of 100 ml
Bottle of 1 l
Bottle of 5 l
Not all pack sizes may be marketed.

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE OF 1 l and 5 l LABEL

All the information required is conveyed on the container

PARTICULARS TO APPEAR ON THE IMMEDIATE PACKAGE – COMBINED LABEL AND PACKAGE LEAFLET

LABEL 1 l and 5 l

Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys

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

Marketing authorisation holder:
LIVISTO Int'l, S.L.
Av. Universitat Autònoma, 29
08290 Cerdanyola del Vallès
(Barcelona) Spain

Manufacturer responsible for batch release:
Industrial Veterinaria S.A.
Esmeralda, 19
08950 Esplugues de Llobregat (Barcelona) Spain

aniMedica GmbH
Im Südfeld 9
48308 Senden-Bösensell Germany

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

Colfive 5,000,000 IU/ml concentrate for oral solution for calves, pigs, lambs, chickens and turkeys
Colistin sulfate

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

Each ml contains:

Active substance:
Colistin sulfate 5,000,000IU

Excipients:
Benzyl alcohol (E1519) 10 mg

Clear orange-brown solution

4. PHARMACEUTICAL FORM

Concentrate for oral solution

5. PACKAGE SIZE

1 l
5 l

6. INDICATION(S)

Calves, lambs, pigs, chickens, turkeys:
Treatment and metaphylaxis of enteric infections caused by non-invasive *E. coli* susceptible to colistin sulfate.The presence of the disease in the herd should be established before metaphylactic treatment.

7. CONTRAINDICATIONS

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

8. ADVERSE REACTIONS

None known.

If you notice any side effects, even those not already listed in this label or you think that the medicine has not worked, please inform your veterinary surgeon.

Alternatively you can report via your national reporting system {national system details}.

9. TARGET SPECIES

Cattle (calves), pigs, sheep (lambs), chickens and turkeys.

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

Oral use.
In drinking water/milk use
Calves, lambs, pigs: 100 000 IU of colistin sulfate per kg body weight daily for 3-5 consecutive days in drinking water or milk (replacer) in calves, equivalent to 0.20 ml of the concentrate solution per 10 kg body weight per day for 3-5 days.

Chickens and turkeys: 75 000 IU of colistin sulfate per kg body weight daily for 3-5 consecutive days in drinking water, equivalent to 15 ml of the concentrate solution per Ton of body weight per day for 3-5 days.

Duration of treatment should be limited to the minimum time necessary for the treatment of the disease.
Any medicated water which is not consumed within 24 hours should be discarded.
Any medicated milk which is not consumed within 6 hours should be discarded.

11. ADVICE ON CORRECT ADMINISTRATION

Direct oral administration to individual animals
The recommended daily dose should be divided into two if the product is to be administered directly into the mouth of the animal.
Prior to direct oral administration, the product should be diluted with a volume of drinking water equivalent to 2.5 x the volume of product concentrate to be administered.

Administration via drinking water
The intake of medicated water depends on the clinical condition of the animals. In order to obtain the correct dosage the concentration of colistin sulfate has to be adjusted accordingly. Carefully calculate the average body weight to be treated and the average daily water consumption before each treatment.
Medicated water should be made every day, immediately prior to provision.
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Water uptake should be monitored at frequent intervals.

With the following formula, we can calculate an exact dosage:

$$\frac{\text{…ml of the product per kg body weight and day} \times \text{Average body weight (kg)}}{\text{Average daily water intake (l/animal)}} = \text{…ml of the product per litre of drinking water}$$

• Administration without a dosing pump:
The treatment is distributed in a tank over a period of 24 hours, for 3-5 consecutive days.
The productis added to a volume of the drinking water corresponding to the volume consumed by the animals over the treatment period (24 hours) to achieve a dose of 100 000 IU of colistin sulfate per kg body weight for pigs, lambs and calves and 75 000 IU of colistin sulfate per kg body weight for chickens and turkeys.

• Administration via a dosing pump
The treatment is distributed over a period of 24 hours, for 3-5 consecutive days.
A dosing pump is used to add a stock solution at a pre-determined concentration to the drinking water.

12. WITHDRAWAL PERIOD

Withdrawal period
Cattle (calves), sheep (lambs) and pigs

Meat and offal: 1 day

Chickens and turkeys
Meat and offal: 1 day
Eggs: Zero days

13. 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.

14. SPECIAL WARNING(S)

Special warnings for each target species:
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.
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 in section 8 of this package leaflet, leading to unnecessary exposure, is notrecommended.

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Whenever possible, colistin sulfate should only be used based on susceptibility testing.
Use of the product deviating from the instructions given in the leaflet may leadto treatment failures and increase the prevalence of bacteria resistant to colistin sulfate. There is cross-resistance between colistin and polymyxin B.
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 polymyxins, such as colistin sulfate, should avoid contact with the veterinary medicinal product.
Avoid direct contact with skin and eyes while handling the product. Personal protective equipment consisting of gloves and protective goggles should be worn when handling and dosing the veterinary medicinal product.
Wash splashes from skin immediately with soap and plenty of water.

In case of accidental eye exposure, wash with plenty of water and seek medical attention immediately and show the label to the physician.
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 or eyes or difficulty in breathing are more serious symptoms and require urgent medical attention.
Wash hands after use.

Use during pregnancy, lactation or lay:
The safety of the veterinary medicinal product has not been established during pregnancy, lactation or lay. However, 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 according to the benefit-risk assessment by the responsible veterinarian.

Interaction with other medicinal products and other forms of interaction:
After oral administration of colistin sulfate interaction with anaesthetics (curarimimetic agents) 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.

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

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

Medicines should not be disposed of via wastewater or household waste. Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal product should be disposed of in accordance with local requirements.
Ask your veterinary surgeon or pharmacist how to dispose of medicines no longer required. These measures should help to protect the environment.

16. DATE ON WHICH THE LABEL WAS LAST APPROVED

January 2021

17. OTHER INFORMATION

The active ingredient colistin sulfate is very persistent in soils.

Package sizes:
Box with a bottle of 100 ml
Bottle of 1 l
Bottle of 5 l
Not all pack sizes may be marketed.

18. 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.

19. THE WORDS "KEEP OUT OF THE SIGHT AND REACH OF CHILDREN"

Keep out of the sight and reach of children.

20. EXPIRY DATE

EXP
Once broached, use by…
Shelf-life after first opening the immediate container: 3 months.
Shelf life after reconstitution in water according to directions: 24 hours
Shelf life after reconstitution in milk according to directions: 6 hours

21. MARKETING AUTHORISATION NUMBER

Vm 43173/4000

22. MANUFACTURER'S BATCH NUMBER

Batch

Approved 29 January 2021