PARTICULARS TO APPEAR ON THE OUTER PACKAGE

Cardboard box (30 tablets)

Cardboard box (90 tablets)

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Isemid 2 mg chewable tablets for dogs (> 11.5-23 kg)

2. STATEMENT OF ACTIVE AND OTHER SUBSTANCES

Each chewable tablet contains:

2 mg of torasemide

3. PACKAGE SIZE

30 chewable tablets
90 chewable tablets

4. TARGET SPECIES

Dogs

5. INDICATION(S)

6. ROUTES OF ADMINISTRATION

Oral use

7. WITHDRAWAL PERIODS

8. EXPIRY DATE

Exp {mm/yyyy}

9. SPECIAL STORAGE PRECAUTIONS

Remaining tablet parts should be stored in the blister and be given at the next administration.

Store out of reach of animals.

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

Ceva Animal Health Ltd

14. MARKETING AUTHORISATION NUMBERS

Vm 15052/5009

15. BATCH NUMBER

Lot {number}

16. SPECIAL WARNING(S), IF NECESSARY

17. SPECIFIC PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCTS OR WASTE MATERIALS, IF ANY

Disposal: read package leaflet.

18. CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE, IF APPLICABLE

POM-V Veterinary medicinal product subject to prescription

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

Blister

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Isemid

2. QUALITATIVE AND QUANTITATIVE PARTICULARS OF THE ACTIVE SUBSTANCES

2 mg torasemide

3. BATCH NUMBER

Lot {number}

4. EXPIRY DATE

EXP {mm/yyyy}

5. ROUTE(S) OF ADMINISTRATION

6. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.

PARTICULARS TO APPEAR ON THE PACKAGE LEAFLET:

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Isemid 1 mg chewable tablets for dogs (2.5-11.5 kg)
Isemid 2 mg chewable tablets for dogs (> 11.5-23 kg)
Isemid 4 mg chewable tablets for dogs (> 23-60 kg)

2. COMPOSITION

Each chewable tablet contains:

Active substance:

Isemid 1 mg	1 mg of torasemide
Isemid 2 mg	2 mg of torasemide
Isemid 4 mg	4 mg of torasemide

The tablets are brown coloured, oblong shaped, chewable and can be divided into halves.

3. TARGET SPECIES

Dogs

4. INDICATIONS FOR USE

For treatment of clinical signs related to congestive heart failure in dogs, including pulmonary oedema.

5. CONTRAINDICATIONS

Do not use in cases of renal failure.
Do not use in cases of dehydration, hypovolaemia or hypotension.
Do not use concomitantly with other loop diuretics.
Do not use in cased of hypersensitivity to the active substance or to any of the excipients.

6. SPECIAL WARNINGS

The initial/maintenance dose may be temporarily increased when pulmonary oedema becomes more severe, i.e. reaching alveolar oedema state (see section 4.8).

Special precautions for use in animals
In dogs presenting with acute pulmonary oedema requiring emergency treatment, the use of injectable medicinal products should be considered first before commencing oral diuretic therapy.

Renal function (measurement of blood urea and creatinine as well as urine protein: creatinine (UPC) ratio), hydration status and serum electrolytes status should be monitored prior to and during treatment at very regular intervals according to the benefit-risk assessment performed by the responsible veterinarian (see sections 4.5 and 4.9 of the SPC). The diuretic response to torasemide may increase over time upon repeated dosing, particularly at doses greater than 0.2 mg/kg/day; therefore more frequent monitoring should be considered.

Torasemide should be used with caution in cases of diabetes mellitus. Monitoring of glycaemia in diabetic animals is recommended prior to and during treatment. In dogs with pre-existing electrolyte and/or water imbalance, this should be corrected prior to treatment with torasemide.

As torasemide increases thirst, dogs should have free access to fresh water.
In case of loss of appetite and/or vomiting and/or lethargy or in case of treatment adjustment, renal function (blood urea and creatinine as well as urine protein:creatinine (UPC) ratio) should be assessed.

In a clinical field trial, the efficacy of Isemid was demonstrated when it was used as a first line treatment. Transferring treatment from an alternative loop diuretic to this veterinary medicinal product has not been evaluated and such a change should only be implemented based on a benefit-risk assessment performed by the responsible veterinarian.

The safety and efficacy of the product has not been assessed for dogs weighing less than 2.5 kg. For these animals use only according to the benefit/risk assessment by the responsible veterinarian.

The tablets are flavoured.

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

This veterinary medicinal product may cause increased urination, thirst and/or gastrointestinal disturbances and/or hypotension and/or dehydration if ingested. Any part-used tablets should be returned to the blister pack and then to the original carton to help prevent access by children. In case of accidental ingestion, particularly by a child, seek medical advice immediately and show the package leaflet or the label to the physician.
This veterinary medicinal product may cause hypersensitivity (allergic) reactions in persons that are sensitized to torasemide. People with known hypersensitivity to torasemide, to sulfonamides or to any of the excipients should avoid contact with the veterinary medicinal product. If symptoms of allergy occur, seek medical advice and show the product packaging to the physician.

Wash hands after use.

Special precautions for the protection of the environment

Not applicable.

Pregnancy and lactation:
Not applicable.

Interactions with other medicinal products and other forms of interaction:
Co-administration of loop diuretics and NSAIDs can result in a decreased natriuretic response.

Concomitant use with NSAIDs, aminoglycosides or cephalosporins may increase the risk of nephrotoxicity and/or ototoxicity of those medicinal products.
Torasemide may antagonise the action of oral hypoglycaemic agents.
Torasemide may increase the risk of sulfonamide allergy.

In cases of co-administration with corticosteroids, the effects of loss of potassium may be potentiated.
In cases of co-administration with amphotericin B, increased potential for nephrotoxicity and intensification of electrolyte imbalance can be observed.

No pharmacokinetic interactions have been reported following co-administration of torasemide with digoxin; however hypokalaemia can enhance digoxin-induced arrhythmias.

Torasemide can reduce the renal excretion of salicylates, leading to an increased risk of toxicity.

Care should be exercised when administering torasemide with other highly plasma protein bound drugs. Since protein binding facilitates the renal secretion of torasemide, a decrease in binding due to displacement by another drug may be a cause of diuretic resistance.

Concomitant administration of torasemide with other substances metabolised by cytochrome P450 families 3A4 (e.g.: enalapril, buprenorphine, doxycycline, cyclosporine) and 2E1 (isoflurane, sevoflurane, theophylline) may decrease their clearance from the systemic circulation.
The effect of antihypertensive veterinary medicinal products, especially angiotensin converting enzyme (ACE)-inhibitors, may be potentiated when co-administered with torasemide.

Overdose (symptoms, emergency procedures, antidotes):
After administration to healthy dogs of 3 times and 5 times the maximum dose for 5 consecutive days followed by 177 daily administrations of 3 times and 5 times the highest therapeutic recommended dose for maintenance, histopathological changes in the kidneys (interstitial inflammation, dilatation of renal tubules and subscapular cysts) were noted in addition to the effects observed after the administration of the recommended dose (see section 7). The renal lesions were still present at 28 days after the end of treatment. Microscopic characteristics of the lesions suggest an ongoing repair process. These lesions may most likely be considered as a result of the pharmacodynamic effect (diuresis) and were not associated with evidence of glomerulosclerosis or interstitial fibrosis. Transient dose response alterations in the

adrenal glands, consisting of minimal to moderate reactive hypertrophy/hyperplasia, presumably related to high production of aldosterone, were observed in the dogs treated with up to 5 times the highest therapeutic recommended dose. An increase in albumin serum concentration was observed. ECG alterations without any clinical signs (increase in P wave and/or QT interval) were observed in some animals after the administration of 5 times the highest recommended dose. The causative role of changes in plasma electrolyte values cannot be excluded.

After administration of 3 and 5 times the highest therapeutic recommended dose to healthy dogs, a decrease of appetite was observed which led to a weight loss in some cases.

In case of overdose, treatment would be at the discretion of the responsible veterinarian, based on the presenting signs.

7. ADVERSE EVENTS

Very common (>1 animal / 10 animals treated):	Renal insufficiency Elevated renal Parameters Electrolyte disorder[1] Haemoconcentration.
Common (1 to 10 animals / 100 animals treated):	Vomiting, Diarrhoea[2] Polyuria, Urinary incontinence[2] Anorexia, Dehydration, Weight loss, Lethargy, Polydipsia.[2]

[1] Chloride, sodium, potassium, phosphorus, magnesium and calcium levels
[2] These signs are episodic.

Other effects consistent with the pharmacological activity of torasemide were observed in pre-clinical studies in healthy dogs at the recommended dose, i.e. dry mucosa of the oral cavity, reversible increases in glucose and aldosterone serum concentrations, decrease in urine specific gravity and increase in urine pH.

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 the local representative of the marketing authorisation holder using the contact details at the end of this leaflet, or via your national reporting system
Website: https://www.gov.uk/report-veterinary-medicine-problem/animal-reacts-medicine
e-mail: adverse.events@vmd.gov.uk

8. DOSAGE FOR EACH SPECIES, ROUTES AND METHOD OF ADMINISTRATION

Oral use.

The recommended initial/maintenance dose is 0.13 to 0.25 mg torasemide/kg bodyweight/day, once daily.
In case of moderate or severe pulmonary oedema, this dose can be increased if necessary up to a maximum dose of 0.4 mg/kg bodyweight/day once daily.

Doses of 0.26 mg/kg and higher should only be administered for a maximum period of 5 days. After this period, the dose should be reduced to the maintenance dose and the dog should be evaluated by the veterinarian in a few days.
The following table shows the dose adjustment scheme within the recommended dose range of 0.13 to 0.4 mg/kg/day:

Dog Bodyweight (kg)	**Number and strength of Isemid tablets to be administered**	
	Initial/Maintenance dose (0.13 to 0.25 mg/kg/day)	**Temporary high dose** (0.26 to 0.40 mg/kg/day)
	1 mg	
2.5 to 4	½	1
> 4 to 6	1	1 + ½
> 6 to 8	From 1 to 1 + ½	From 2 to 2 + ½
> 8 to 11.5	From 1 + ½ to 2	From 2 + ½ to 3
	2 mg	
> 11.5 to 15	From 1 to 1 + ½	2
> 15 to 23	From 1 + ½ to 2	From 2 + ½ to 3
	4 mg	
> 23 to 30	From 1 to 1 + ½	2
> 30 to 40	From 1 + ½ to 2	From 2 + ½ to 3
> 40 to 60	From 2 to 2 + ½	From 3 to 4

The dose should be adjusted to maintain patient comfort with attention to renal function and electrolytes status. Once signs of congestive heart failure have been controlled and the patient is stable, it should be continued at the lowest effective dose, if long term diuretic therapy with this product is required.

9. ADVICE ON CORRECT ADMINISTRATION

If the tablet is not spontaneously taken by the dog, it can also be given with food or directly into the mouth.

10. WITHDRAWAL PERIODS

Not applicable.

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 carton or blister pack after "EXP". The expiry date refers to the last day of that month.

Remaining tablet parts should be stored in the blister and be given at the next administration. Keep the tablets out of the reach of animals in order to avoid any accidental ingestion.

12. SPECIAL PRECAUTIONS FOR DISPOSAL

Medicines should not be disposed of via wastewater.

Any unused veterinary medicinal product or waste material 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.

13. CLASSIFICATION OF VETERINARY MEDICINAL PRODUCTS

Veterinary medicinal product subject to prescription

14. MARKETING AUTHORISATION NUMBERS AND PACK SIZES

Vm 15052/5008

Vm 15052/5009

Vm 15052/5010

Pack sizes
Each blister contains 10 tablets and is packed in a cardboard box.
All strengths are available in pack sizes of 30 or 90 tablets.
Not all pack sizes may be marketed.

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:
Ceva Animal Health Ltd
Explorer House
Mercury Park
Wycombe Lane
Wooburn Green
High Wycombe
Buckinghamshire
HP10 0HH
United Kingdom
Tel. 01628 334056
Email technicalandpvuk-group@ceva.com

Manufacturer responsible for batch release:
Ceva Santé Animale
10 av. de La Ballastière
33500 Libourne
FRANCE
Tel: +800 35 22 11 51
E-mail: pharmacovigilance@ceva.com

Ceva Santé Animale
Boulevard de la Communication,
Zone autoroutière
53950 Louverne
France

17. OTHER INFORMATION

Gavin Hall

Approved 07 October 2024