

ANNEX III
LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

CARTON

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Zenalpha 0.5 mg/ml + 10 mg/ml solution for injection for dogs
medetomidine hydrochloride / vatinoxan hydrochloride

2. STATEMENT OF ACTIVE SUBSTANCES

1 ml contains:
0.5 mg medetomidine hydrochloride (equivalent to 0.425 mg medetomidine)
10 mg vatinoxan hydrochloride (equivalent to 9.2 mg vatinoxan)

3. PHARMACEUTICAL FORM

Solution for injection.

4. PACKAGE SIZE

10 ml
5 x 10 ml
10 x 10 ml

5. TARGET SPECIES

Dogs.

6. INDICATION(S)

7. METHOD AND ROUTE(S) OF ADMINISTRATION

For intramuscular use.
Read the package leaflet before use.

8. WITHDRAWAL PERIOD(S)

9. SPECIAL WARNING(S), IF NECESSARY

10. EXPIRY DATE

EXP:
Once broached use within 3 months.

11. SPECIAL STORAGE CONDITIONS

Keep the vial in the outer carton in order to protect from light.

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

Vetcare Oy
P. O. Box 99
FI-24101 Salo
Finland

16. MARKETING AUTHORISATION NUMBER(S)

Vm 42810/5000

17. MANUFACTURER'S BATCH NUMBER

Lot:

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

LABEL

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

Zenalpha 0.5 mg/ml + 10 mg/ml solution for injection for dogs
medetomidine hydrochloride / vatinoxan hydrochloride

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

0.5 mg/ml medetomidine hydrochloride
10 mg/ml vatinoxan hydrochloride

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

10 ml

4. ROUTE(S) OF ADMINISTRATION

IM

5. WITHDRAWAL PERIOD(S)

6. BATCH NUMBER

Lot:

7. EXPIRY DATE

EXP:
Once broached use within 3 months.

8. THE WORDS "FOR ANIMAL TREATMENT ONLY"

For animal treatment only.

B. PACKAGE LEAFLET

PACKAGE LEAFLET:
Zenalpha 0.5 mg/ml + 10 mg/ml solution for injection for dogs

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

Marketing authorisation holder:

Vetcare Oy, P.O. Box 99, 24101 Salo, Finland

Manufacturer responsible for batch release:

Apotek Produktion & Laboratorier AB, Formvägen 5B, SE-90621 Umeå, Sweden
Eurovet Animal Health BV, Handelsweg 25, NL-5531 AE Bladel, The Netherlands

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

Zenalpha 0.5 mg/ml +10 mg/ml solution for injection for dogs
medetomidine hydrochloride / vatinoxan hydrochloride

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

1 ml contains:

Active substances:

Medetomidine hydrochloride	0.5 mg (equivalent to 0.425 mg medetomidine)
Vatinoxan hydrochloride	10 mg (equivalent to 9.2 mg vatinoxan)

Excipients:

Methyl parahydroxybenzoate (E 218)	1.8 mg
Propyl parahydroxybenzoate	0.2 mg

Clear, slightly yellow to yellow or brownish yellow solution.

4. INDICATION(S)

To provide restraint, sedation and analgesia during conduct of non-invasive, non-painful or mildly painful procedures and examinations intended to last no more than 30 minutes.

5. CONTRAINDICATIONS

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

Do not use in animals with cardiovascular disease, respiratory disease or impaired liver or kidney function.

Do not use in animals that are in shock or severely debilitated.

Do not use in animals that have hypoglycaemia or are at risk of developing hypoglycaemia.

Do not use as pre-anaesthetic medicine.

Do not use in cats.

6. ADVERSE REACTIONS

Hypothermia, bradycardia and tachycardia were very commonly observed in safety and clinical studies. Diarrhoea/colitis and muscle tremor were commonly observed. Vomiting/nausea and involuntary defaecation were uncommonly observed. Cardiac arrhythmias, such as second-degree AV block and ventricular escape complexes, were seen very commonly in laboratory safety studies. Injected sclera were observed very rarely in laboratory safety studies.

All of the above adverse reactions were transient/resolved without treatment, although external warming was provided where required in cases of hypothermia.

The frequency of adverse reactions is defined using the following convention:

- very common (more than 1 in 10 animals treated displaying adverse reaction(s))
- common (more than 1 but less than 10 animals in 100 animals treated)
- uncommon (more than 1 but less than 10 animals in 1,000 animals treated)
- rare (more than 1 but less than 10 animals in 10,000 animals treated)
- very rare (less than 1 animal in 10,000 animals treated, including isolated reports).

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.

7. TARGET SPECIES

Dogs.

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

For intramuscular use.

The dose is based on body surface area. The dose will result in administration of 1 mg medetomidine and 20 mg vatinoxan per square metre of body surface area (m²).

Calculate the dose using 1 mg/m² medetomidine or use the dosing table below. Note that the mg/kg dosage decreases as body weight increases.

To ensure a correct dosage, bodyweight should be determined as accurately as possible.

Table 1. Dose volume based on bodyweight

Dog bodyweight kg	Dose volume ml
3.5 to 4	0.4
4.1 to 5	0.6
5.1 to 7	0.7
7.1 to 10	0.8
10.1 to 13	1.0
13.1 to 15	1.2
15.1 to 20	1.4
20.1 to 25	1.6
25.1 to 30	1.8
30.1 to 33	2.0
33.1 to 37	2.2
37.1 to 45	2.4
45.1 to 50	2.6
50.1 to 55	2.8
55.1 to 60	3.0
60.1 to 65	3.2
65.1 to 70	3.4
70.1 to 80	3.6
>80	3.8

Re-administration of the veterinary medicinal product during the same procedure has not been evaluated and therefore the veterinary medicinal product should not be re-administered during the same procedure.

The number of permissible stopper broachings is limited to a maximum of 15.

9. ADVICE ON CORRECT ADMINISTRATION

Use of an appropriately graduated syringe is recommended to ensure accurate dosing when administering small volumes.

10. WITHDRAWAL PERIOD(S)

Not applicable.

11. SPECIAL STORAGE PRECAUTIONS

Keep out of the sight and reach of children.

Keep the vial in the outer carton in order to protect from light.

Do not use this veterinary medicinal product after the expiry date which is stated on the carton and label after EXP.

Shelf life after first opening the immediate packaging: 3 months.

12. SPECIAL WARNING(S)

Special warnings for each target species:

Nervous or excited dogs with high levels of endogenous catecholamines may exhibit a reduced pharmacological response to alpha-2 adrenoceptor agonists like medetomidine (ineffectiveness). In agitated animals, the onset of sedative/analgesic effects could be slowed, or the depth and duration of effects could be diminished or non-existent. Therefore, the dog should be given the possibility to calm down before initiation of the treatment and rest quietly after administration of the product until evidence of sedation has occurred.

Special precautions for use in animals:

In the absence of available data, treatment of puppies less than 4.5 months of age should be based on a benefit-risk assessment by the responsible veterinarian.

It is recommended that dogs should be fasted in accordance with currently recommended best practice (e.g. 4 – 6 hours for healthy dogs), prior to treatment with this veterinary medicinal product. Water can be given.

Animals should be frequently monitored for cardiovascular function and body temperature during sedation and recovery.

Some cardiovascular effects (e.g. bradycardia, cardiac arrhythmias such as second degree AV block or ventricular escape complexes) may be observed after treatment.

Over the period of 15–45 minutes post treatment, blood pressure is likely to decrease by approximately 30–50 % from pre-treatment levels. Tachycardia with normal blood pressure may be observed from approximately one hour post-treatment and lasting up to six hours. Therefore, frequent monitoring of cardiovascular function should preferably be performed until tachycardia has resolved.

A decrease in body temperature of approximately 1–2 °C is likely to occur after administration.

Once established, hypothermia may persist longer than sedation and analgesia. To prevent hypothermia, treated animals should be kept warm and at a constant temperature during the procedure and until fully recovered.

Medetomidine can cause apnoea and/or hypoxaemia. This effect is likely to be potentiated if used in combination with opioid medicines. Frequent monitoring of respiratory function should be performed in all cases. It is also advisable to have oxygen readily available, should hypoxaemia be detected or suspected.

Analgesia provided by the veterinary medicinal product may be shorter than the sedative effect. Additional pain management should be provided as needed.

Spontaneous muscle trembling or twitching can be expected in some dogs.

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

Accidental exposure may cause sedation and changes in blood pressure. Caution is required during treatment administration to avoid accidental self-injection, or skin, eye or mucosal contact. Adequate restraint of the animal is recommended, as some animals may react to the injection (e.g., defence reaction).

Pregnant women should administer the veterinary medicinal product with special caution to avoid self-injection since uterine contractions and decreased foetal blood pressure may occur after accidental systemic exposure.

People with known hypersensitivity to the active substance or any of the excipients should administer the veterinary medicinal product with caution.

In case of accidental self-injection or ingestion, seek medical advice immediately and show the package leaflet to the physician but DO NOT DRIVE.

In case of skin or mucosal contact, wash the exposed skin immediately after exposure with large amounts of water and remove contaminated clothes that are in direct contact with skin. In case of eye contact, rinse abundantly with fresh water. If symptoms occur, seek the advice of a physician.

To the physician: The veterinary medicinal product contains medetomidine, an alpha-2 adrenoceptor agonist, in combination with vatinoxan, a peripherally selective alpha-2 adrenoceptor antagonist. Symptoms after absorption may involve clinical effects including dose-dependent sedation, respiratory depression, bradycardia, hypotension, a dry mouth, and hyperglycaemia. Ventricular arrhythmias have also been reported. Respiratory and haemodynamic symptoms should be treated symptomatically.

Pregnancy and lactation:

The safety of the veterinary medicinal product has not been established in dogs during pregnancy or lactation or in dogs intended for breeding. No data are available on the use of vatinoxan in breeding animals. Published data in laboratory animals do not show direct reproductive or developmental toxicity of medetomidine. The use of the veterinary medicinal product is therefore not recommended in pregnant or lactating animals.

Interaction with other medicinal products and other forms of interaction:

The use of other central nervous system depressants and/or vasodilating medicines are expected to potentiate the effects of the veterinary medicinal product and an appropriate dose reduction should be made after benefit-risk assessment by the veterinarian.

Due to the rapid recovery from sedation expected with the veterinary medicinal product, routine administration of atipamezole is not indicated after the veterinary medicinal product. The intramuscular administration of atipamezole (30 minutes after the administration of the veterinary medicinal product) has been investigated in a study involving a limited number of animals. As tachycardia was observed in 50% of the animals following the administration of atipamezole, close monitoring of the heart

rate during recovery is therefore advised in those cases where administration of atipamezole is deemed clinically necessary.

Overdose (symptoms, emergency procedures, antidotes):

The veterinary medicinal product given 3 and 5 times the recommended dose, showed a slightly prolonged sedation and greater degree of reduction in mean arterial pressure and rectal temperature. Overdose can increase incidence of sinus tachycardia during recovery.

Atipamezole can be administered to reverse the central nervous system effects and most of the cardiovascular effects of medetomidine, excluding hypotension. Appropriate cardiopulmonary support should be initiated if required.

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. 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

August 2022

15. OTHER INFORMATION

Pack sizes: 10 ml, 5 x 10 ml, 10 x 10 ml
Not all pack sizes may be marketed.

Approved 26 August 2022

