

VETERINARY USE ONLY
NERFASIN 20™
 Xylazine Injection, Mfr Std
 20 mg/mL Sterile Injectable Solution
Sedative and Analgesic
For Use In Dogs, Cats and Cattle Only

DESCRIPTION:

Nerfasin 20™ (xylazine) is supplied in 10, 25, and 50 mL multiple-dose vials as a sterile solution.

Active Ingredients - Each mL contains:

Xylazine (as hydrochloride).....20 mg

Preservatives - Each mL contains:

Methyl parahydroxybenzoate.....1 mg

Inactive Ingredients - Each mL contains:

Sodium chloride.....3 mg

Sodium hydrogen carbonate solution.....q.s.

Hydrochloric acid.....q.s.

Water for injection.....q.s.

PHARMACOLOGY:

Xylazine is a potent sedative and analgesic as well as muscle relaxant. Its sedative and analgesic activity is related to central nervous system depression. Its muscle-relaxant effect is based on inhibition of the intraneural transmission of impulses in the central nervous system. The principal pharmacological activities develop within 10 to 15 minutes after intramuscular injection and within 3 to 5 minutes following intravenous administration.

A sleeplike state, the depth of which is dose-dependent, is usually maintained for 1 to 2 hours, while analgesia lasts from 15 to 30 minutes. The centrally-acting muscle relaxant effect causes relaxation of the skeletal musculature, complementing sedation and analgesia.

In animals under the influence of xylazine, the respiratory rate is reduced as in natural sleep. Following treatment with xylazine, the heart rate is decreased and a transient change in the conductivity of the cardiac muscle may occur, as evidenced by a partial atrioventricular block. This resembles the atrioventricular block often observed in normal animals^{1,2,3,4}. Intravenous administration of xylazine causes a transient rise in blood pressure, followed by a slight decrease.

Xylazine has no effect on blood clotting time or other hematologic parameters.

In limited tests, xylazine has been tolerated in dogs and cats at 10 times the recommended dose. However, doses of this magnitude produce muscle tremors, emesis and long periods of sedation.

INDICATIONS: DOGS AND CATS

Nerfasin 20™ should be used in dogs and cats when it is desirable to produce a state of sedation accompanied by a shorter period of analgesia. Xylazine has been used successfully as follows:

1. Diagnostic procedures - examination of the mouth and ears, abdominal palpation, rectal palpation, vaginal examination, catheterization of the bladder and radiographic examinations.
2. Orthopedic procedures, such as application of casting materials and splints.
3. Dental procedures.
4. Minor surgical procedures of short duration such as debridement, removal of cutaneous neoplasms and suturing of lacerations.
5. To calm and facilitate handling of fractious animals.
6. Therapeutic medication for sedation and relief of pain following injury or surgery.
7. Major surgical procedures:
 - a. When used as a preanesthetic to general anesthesia.
 - b. When used in conjunction with local anesthetics.

CONTRAINDICATIONS: DOGS AND CATS

Nerfasin 20™ is not recommended for use in pregnant cats.

Do not use Nerfasin 20™ in conjunction with tranquilizers.

DOSAGE AND ADMINISTRATION: DOGS AND CATS

1. Intravenously - 0.5 mL/9 kg of body weight (1.1 mg/kg or 0.5 mg/lb)

Intramuscularly or subcutaneously - 1.0 mL/9 kg body weight (2.2 mg/kg or 1 mg/lb). In large dogs (over 22.7 kg/50 lbs.) a dosage of 1.1 mg/kg (0.5 mg/lb) administered intramuscularly may provide sufficient sedation and/or analgesia for most procedures.

Since vomiting may occur (see ADVERSE REACTIONS), fasting for 6-24 hours prior to the use of Nerfasin 20™ may reduce the incidence; the I.V. route results in the least vomiting.

Following injection of xylazine, the animal should be allowed to rest quietly until the full effect has been reached.

These dosages produce sedation which is usually maintained for 1 to 2 hours and analgesia which lasts for 15 to 30 minutes.

2. Preanesthetic to Local Anaesthesia:

Nerfasin 20™ at the recommended dosages can be used in conjunction with local anesthetics such as procaine or lidocaine.

3. Preanesthetic to General Anaesthesia:

Nerfasin 20™, at the recommended dosage rates, produces an additive effect to central nervous system depressants such as pentobarbital sodium, thiopental sodium and thiamylal sodium. Therefore, the dosage of such compounds should be reduced and administered to the desired effect. In general, only 1/3 to 1/2 of the calculated dosage of the barbiturates will be needed to produce a surgical plane of anesthesia. Post anesthetic or emergence excitement has not been observed in animals preanesthetized with xylazine.

Xylazine has been used successfully as a preanesthetic agent for pentobarbital sodium, thiopental sodium, thiamylal sodium, nitrous oxide, ether, halothane and methoxyflurane anesthesia.

ADVERSE REACTIONS: DOGS AND CATS

Emesis occurs occasionally in dogs and frequently in cats, soon after the administration of Nerfasin 20™. When observed, emesis usually occurs only a single time, after which there is no further emetic effect.

The use of antiemetics may delay this phenomenon.

The occurrence of emesis may be considered a desirable effect when Nerfasin 20™ is administered as a preanesthetic to general anesthesia.

Nerfasin 20™ used at recommended dosage levels may occasionally cause slight muscle tremors, bradycardia with partial A-V heart block and a reduced respiratory rate. Should excessive respiratory depression occur following the Nerfasin 20™, administer respiratory stimulants and provide artificial respiration.

Movement in response to sharp auditory stimuli may be observed.

Increased urination may occur in cats following the use of Nerfasin 20™.

CAUTIONS: DOGS AND CATS

Until more definitive studies are completed, Nerfasin 20™ is not recommended for use in pregnant cats.

Careful consideration should be given before administering to dogs or cats with significantly depressed respiration, severe pathologic heart disease, advanced liver and kidney disease, severe endotoxic or traumatic shock.

Since an additive effect results from the use of Nerfasin 20™ and the barbiturate compounds, it should be used with caution with these central nervous system depressants. Products known to produce respiratory depression or apnea, such as thiamylal sodium, should be given at a reduced dosage and, when injected intravenously, should be administered SLOWLY.

When intravenous administration is desired, avoid perivascular injection in order to achieve the desired effect. Studies have shown negligible evidence of tissue irritation, however, following perivascular injection of xylazine. Following the use of Nerfasin 20™, veterinarians and attendants should continue to use care and appropriate animal handling techniques, since conscious animals, although sedated, are arousable and capable of inflicting personal injury. Bradycardia and an arrhythmia in the form of incomplete atrioventricular block have been reported following xylazine administration. Although clinically the importance of this effect is questioned^{1,2,3,4}, a standard dose of atropine given prior to or following Nerfasin 20™ will greatly decrease the incidence.

INDICATIONS: CATTLE

Nerfasin 20™ is indicated in cattle to produce a state of sedation accompanied by a shorter period of analgesia. Xylazine has been used successfully as follows:

1. Diagnostic procedures - oral, vaginal and rectal examinations, as an aid in the collection of biopsies or blood samples and radiographic examinations.
2. Orthopedic procedures, such as application of casting materials and splints.
3. Dental procedures.
4. Minor surgical procedures of short duration such as debridement of wounds, dehorning, castration and suturing of skin lacerations.
5. Major surgical procedures when used in conjunction with local and epidural anesthetics - suturing of lacerations of the test and udder, surgery of the penis and sheath, caesarean sections, hernia repairs, digital amputations and eye enucleations.
6. Hoof trimming and handling of fractious animals.

CONTRAINDICATIONS: CATTLE

Do not use in pregnant animals as studies have not been completed to show its safety in all stages of pregnancy. Premature parturition and retained placenta have been reported in a limited number of cases where xylazine was administered during the last trimester of pregnancy.

Do not use Nerfasin 20™ in conjunction with tranquilizers.

DOSAGE AND ADMINISTRATION: CATTLE

Intramuscularly - Range of 0.25 to 0.75 mL/45 kg body weight (Equivalent to 0.11 to 0.33 mg/kg or 0.05 to 0.15 mg/lb).

Ruminants are more sensitive to Nerfasin 20™ than are other species in which the drug is indicated, and thus a much smaller dose is required per unit body weight to produce the desired effect.

The dosage of Nerfasin 20™ in the bovine species needed to achieve the desired effect varies between animals, depending largely upon the temperament of the individual animal. Quieter or more docile cattle will require a smaller dose to achieve the same effect. Nerfasin 20™ will often make the animal recumbent especially at the higher dose rates. Following injection of Nerfasin 20™ the animal should be allowed to rest quietly until the full effect has been reached.

Within the recommended dosage range, a range of effects can be achieved depending on the dose given. Low doses of Nerfasin 20™ produce a sedation and limited dermal analgesia while larger doses produce sedation, muscle relaxation described, produce recumbency and a true anaesthesia like condition under which many procedures may be carried out with or without local anaesthesia. Even high doses will not eliminate pain in the claws and lower limbs.

After intramuscular injection of Nerfasin 20™ the onset of sedation and analgesia follows in less than 10 minutes along with some incoordination. The duration of sedation and analgesia along with the ability to stand depends on the dose given. Duration of sedation and analgesia will vary from 30 minutes with low doses to 2 to 3 hours with higher doses.

Within the recommended dosage range Nerfasin 20™ can be used in conjunction with local anesthetics such as procaine and lidocaine. Many procedures may be carried out using Nerfasin 20™ alone especially at the higher dose rate.

ADVERSE REACTIONS: CATTLE

Nerfasin 20™ used at recommended dosage levels may occasionally cause slight muscle tremors, bradycardia and a reduced respiratory rate. Temporary salivation, diuresis and ruminal stasis may be observed during the period of sedation. A transient, self-limiting diarrhea may occur 24 to 48 hours following administration.

CAUTIONS: CATTLE

Careful consideration should be given before administering to cattle with significantly depressed respiration, severe pathologic heart disease, advanced liver or kidney disease, severe endotoxic or traumatic shock.

Special precautions should be taken when administering during warm environmental conditions as HYPERTHERMIA may occur. Proper aftercare must be provided in those cases. Always provide cool shade during the recovery period.

Lateral recumbency is to be avoided during recovery due to increasing the possibilities of bloat, regurgitation and/or aspiration. Sternal recumbency is the appropriate recovery position. A 24-hour fast prior to injection will also reduce the incidence of bloat.

Following the use of Nerfasin 20™, veterinarians and attendants should continue to use care and appropriate animal handling techniques, since conscious animals, although sedated, are capable of inflicting personal injury.

SAFETY:

Xylazine has been tolerated in cattle at 10 times the recommended dose. However, doses of this magnitude produced muscle tremors and long periods of sedation with careful surveillance necessary during the recovery period.

WARNINGS:

TREATED CATTLE MUST NOT BE SLAUGHTERED FOR USE IN FOOD FOR AT LEAST 3 DAYS AFTER THE LATEST TREATMENT WITH THIS DRUG. MILK TAKEN FROM TREATED ANIMALS DURING TREATMENT AND WITHIN 48 HOURS AFTER THE LATEST TREATMENT MUST NOT BE USED AS FOOD. This drug is for use in Dogs, Cats, and Cattle only.

Keep out of reach of children. Do not eat, drink, or smoke while handling veterinary drug product. Avoid skin and eye contact. For veterinary use only.

Xylazine is an alpha2-adrenergic agonist with sedative, some analgesic and muscle relaxant properties. Symptoms after absorption may include dose-dependent respiratory depression, bradycardia, hypotension, a dry mouth, and hyperglycemia. Ventricular arrhythmias have also been reported.

Strictly avoid self-injection, oral intake and any contact with the skin, eyes or mucosa. In the case of accidental contact, was exposed skin or eyes abundantly with water. If symptoms occur, seek medical advice. In the case of accidental oral intake or self-injection, seek the advice of a physician and show the package insert but DO NOT DRIVE.

If pregnant women handle the product, special caution should be observed not to self-inject as uterine contractions and decreased fetal blood pressure may occur after accidental systemic exposure.

Dispose unused drug or waste materials in accordance with the Provincial/Municipal guidelines.

STORAGE: Store at room temperature, 15-30°C; Do not freeze. Store the vial in the upright position. Discard any remainder 28 days after broaching. The stopper should not be punctured more than 20 times after broaching.

REFERENCES:

1. Detweiler, D.K.: The Diagnosis and Significance of Cardiac Arrhythmias in Progress in Equine Practice. Edited by E.J. Catcott and J.F. Smithcoors. American Veterinary Publications, Inc., Santa Barbara, California and Wheaton, Illinois, (1966), 280-281.
2. Glazier, D.B.: Atrioventricular Heart Block. Irish Vet. J., Vol. 12 (1958): 194-198.
3. Holmes, J.R., Alps, B.J.: Observations on Partial Atrioventricular Heart Block in the Horse. Can. Vet. J., Vol. 7, No. 12 (1966): 280-290.
4. Detweiler, D.K., Patterson, D.F., Luginuh, H., Rhodes, W.H., Buchanan, J.W., Knight, D.H., Hill, J.D.: Diseases of the Cardiovascular System in Canine Medicine. Edited by E.J. Catcott. American Veterinary Publication, Inc., Santa Barbara, California and Wheaton, Illinois, (1968), 589-679.

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