Simparica TRIO.

(sarolaner, moxidectin, and pyrantel chews for dogs)

**ORAL USE ONLY FOR DOGS ONLY**

**CAUTION**
Federal (USA) law restricts this drug for use by or on the order of a licensed veterinarian.

**DESCRIPTION**
Simparica TRIO contains the S-enantiomer of sarolaner. Sarolaner is a member of the isoxazoline class of parasiticides and the chemical name is 1,4,5,6-tetrahydro-23(S)-3-{[2-(3-hydroxy-2-naphthyl)]-1,1-dimethyl-2-buten-1-yl}-2,6-epoxy-23-(methylsulfonyliminobenzyl) B. Sarolaner is a semi-synthetic methoxime derivative of remdecanate which is a fermentation product of Strepognum cyanescens subsp. noncyanogenum. Moxidectin is a pentaacetyl 16-membered lactone macrolide. The chemical name for moxidectin is (3R,2S,5S)-5-Demethyl-3-deoxy-25-[1(E),3-dimethyl-1-buten-1-yl]-6,26-epoxy-25-methoxypyrrolomycin B.

Pyrantel belongs to a family chemically classified as tetrahydropyrindines and the chemical name is C9H19N2O2. Pyrantel pamoate is an anti-parasitic agent that can be administered to dogs with or without food.

**DOSAGE AND ADMINISTRATION**

**Tick Treatment and Control:**
Should be administered year-round at monthly intervals or started at least one month continuing until at least 1 month after the dog's last seasonal exposure. If a dose is missed, give it immediately and resume monthly dosing.

**Pet Care:**
Care should be taken to ensure that the dog consumes the complete dose and that SIMPARICA TRIO can be offered to the dog with or without food.

**ADVERSE REACTIONS**
In a well-controlled laboratory study, SIMPARICA TRIO was administered to dogs for the prevention of heartworm disease caused by Dirofilaria immitis and for the treatment and control of roundworm (immature adult and adult) Toxocara canis and hookworm (L4, immature adult, and adult Anclylostoma caninum and adult Uncinaria stenocephala) infections.

**MODE OF ACTION**

*The mode of action of simparica is described as follows:*

**TICKS**
Simparica TRIO is indicated for the prevention of *Ixodes scapularis* (blacklegged tick), *Dermacentor variabilis* (American dog tick), *Amblyomma maculatum* (Gulf Coast tick), *Amblyomma americanum* (dog tick), and *Ctenocephalides felis* (cat flea).

**FLEAS**
Simparica TRIO is indicated for the prevention of flea infestations and circulating microfilariae. Dogs in the control group received placebo. Clinical signs (claws, muscle fasciulations, mydriasis) associated with avermectin sensitivity were observed in the SX group. All dogs were completely recovered by the third day of the study.

**FLEA CONTROL**
In two well-controlled field studies, SIMPARICA TRIO was used concurrently with other medications such as vaccines, antimicrobials, antihelmintics, antiparasitics, steroids, and non-steroidal anti-inflammatory agents, antibiotics and analgesics. No adverse reaction was associated with the concurrent use of SIMPARICA TRIO and other medications.

**STORAGE CONDITIONS**
Store at or below 30°C (86°F).

**HOW SUPPLIED**
Simparica TRIO is available in six flavor tab sizes (see DOSAGE AND ADMINISTRATION). Each tablet size is available in packages of one, three, or six tablets.

**EFFECTIVENESS**

**Heartworm Prevention**
In two well-controlled laboratory studies, a single oral dose of SIMPARICA TRIO was 100% effective in preventing the development of adult D. immitis in dogs inoculated with infective larvae 30 days before treatment.

**Flea Treatment and Prevention**
In a well-controlled laboratory study, SIMPARICA TRIO began to kill fleas at 4 hours and demonstrated 100% effectiveness at 8 hours after initial administration. After weekly re-infestations, SIMPARICA TRIO reduced the number of live fleas by ≥97.9% effectiveness against weekly re-infestations for 35 days. In a study to explore flea egg production and viability, SIMPARICA TRIO killed fleas before they could lay eggs for 28 days.

**Intestinal Nematode Treatment and Control**

*For the treatment of roundworm (immature adult and adult) Toxocara canis and adult Toxocara canis* hookworm (L4, immature adult, and adult *Anclylostoma caninum* and adult *Uncinaria stenocephala* infections, SIMPARICA TRIO should be administered once as a single dose. Monthly use of SIMPARICA TRIO will control any subsequent infections.

**CONTRAINDICATIONS**

*There are no known contraindications for the use of SIMPARICA TRIO.*

**Warnings**
Not for use in humans. Keep this and all drugs out of reach of children.

**Precautions**

*Sarolaner, one of the ingredients in SIMPARICA TRIO, is a member of the isoxazoline class. This class has been associated with neurologic adverse reactions including tremors, ataxia, and seizures. Seizures have been reported in dogs receiving isoxazoline class drugs, even in dogs without a history of seizures. Use with caution in dogs with a history of seizures or neurologic disorders.*

**Toxocara canis** infections, SIMPARICA TRIO should be administered orally once at 1, 3, and 5X the maximum labeled dose (2.4 mg/kg sarolaner, 4.4 mg/kg moxidectin, and 2.0 mg/kg pyrantel) at 28 day intervals for 7 treatments. Dogs in the control group received placebo. Clinical signs (claws, muscle fasciulations, mydriasis) associated with avermectin sensitivity were observed in the SX group. All dogs were completely recovered by the third day of the study.

**Heartworm Prevention:**
SIMPARICA TRIO should be administered at monthly intervals year-round at or at least within one month of the animal's first seasonal exposure to mosquitos and continuing until at least one month after the last dose. If a dose is missed, give it immediately and resume monthly dosing.

**Heartworm Prevention:**
SIMPARICA TRIO should be administered at monthly intervals year-round at or at least within one month of the animal's first seasonal exposure to mosquitos and continuing until at least one month after the last dose. If a dose is missed, give it immediately and resume monthly dosing. When a monthly heartworm preventive product, SIMPARICA TRIO should be given within one week of the last dose of the former medication.

**Flea Treatment and Prevention:**
Treatment with SIMPARICA TRIO may begin at any time of the year. SIMPARICA TRIO should be administered year-round at monthly intervals or started at least one month before fleas become active.

**To minimize the likelihood of flea re-infestation, it is important to treat all dogs and cats with a flea control product.**

**Tick Treatment and Control:**
Treatment with SIMPARICA TRIO can begin at any time of the year. SIMPARICA TRIO should be administered year-round at monthly intervals or started at least one month before fleas become active.

**Intestinal Nematode Treatment and Control:**

*For the treatment of roundworm (immature adult and adult) Toxocara canis and adult Toxocara canis* hookworm (L4, immature adult, and adult *Anclylostoma caninum* and adult *Uncinaria stenocephala* infections, SIMPARICA TRIO should be administered once as a single dose. Monthly use of SIMPARICA TRIO will control any subsequent infections.*

**CONTRAINDICATIONS**

*There are no known contraindications for the use of SIMPARICA TRIO.*

**Warnings**
Not for use in humans. Keep this and all drugs out of reach of children.

**Precautions**

*Sarolaner, one of the ingredients in SIMPARICA TRIO, is a member of the isoxazoline class. This class has been associated with neurologic adverse reactions including tremors, ataxia, and seizures. Seizures have been reported in dogs receiving isoxazoline class drugs, even in dogs without a history of seizures. Use with caution in dogs with a history of seizures or neurologic disorders.*

**Toxocara canis** infections, SIMPARICA TRIO should be administered orally once at 1, 3, and 5X the maximum labeled dose (2.4 mg/kg sarolaner, 4.4 mg/kg moxidectin, and 2.0 mg/kg pyrantel) at 28 day intervals for 7 treatments. Dogs in the control group received placebo. Clinical signs (claws, muscle fasciulations, mydriasis) associated with avermectin sensitivity were observed in the SX group. All dogs were completely recovered by the third day of the study.

**Heartworm Prevention:**
SIMPARICA TRIO should be administered at monthly intervals year-round at or at least within one month of the animal's first seasonal exposure to mosquitos and continuing until at least one month after the last dose. If a dose is missed, give it immediately and resume monthly dosing. When a monthly heartworm preventive product, SIMPARICA TRIO should be given within one week of the last dose of the former medication.

**Flea Treatment and Prevention:**
Treatment with SIMPARICA TRIO may begin at any time of the year. SIMPARICA TRIO should be administered year-round at monthly intervals or started at least one month before fleas become active.

**To minimize the likelihood of flea re-infestation, it is important to treat all dogs and cats with a flea control product.**

**Tick Treatment and Control:**
Treatment with SIMPARICA TRIO can begin at any time of the year. SIMPARICA TRIO should be administered year-round at monthly intervals or started at least one month before fleas become active.