

FIPRONIL 9.8% w/v (S)-METHOPRENE 8.8% w/v FRONTLINE®

PLUS

Spot-on solution.

For Dogs
For external use only



Composition

Frontline Plus (Dogs) contains:
Fipronil9.8 %
(S)-methoprene.....8.8%

Keep out of reach of children.

Not for human use; for animal treatment only.

WARNING: To be sold by retail on the prescription of RMP only.

Indication

- To be used against infestation with fleas, alone or in association with ticks and/or biting lice.
 - Treatment of flea infestations (*Ctenocephalides* spp.). Insecticidal efficacy against new infestations with adult fleas persists for 8 weeks. Prevention of the multiplication of fleas by inhibiting the development of eggs (ovicidal activity) and larvae and pupae (larvicidal activity) originating from eggs laid by adult fleas for 8 weeks after application.
 - Treatment of tick infestations (*Ixodes ricinus*, *Dermacentor variabilis*, *Dermacentor reticulatus*, *Rhipicephalus sanguineus*). The product has a persistent acaricidal efficacy for up to 4 weeks against ticks (based on experimental data).
 - Treatment of infestations with biting lice (*Trichodectes canis*).
- The product can be used as part of a treatment strategy for the control of Flea Allergy Dermatitis (FAD). Fipronil kills fleas within 24 hours and ticks and lice within 48 hours post-exposure.

Dosage and method of administration

One pipette corresponds to the minimum recommended dose of 6.7mg/kg for fipronil and 6mg/kg for (S)-methoprene by topical application to the skin. Refer below table for dosage as per body weight dog.

Weight of dog	Size of animal	Each Pipette contains
2 – 10 kgs	Small (S)	0.67ml
10 – 20 kgs	Medium (M)	1.34ml
20 – 40 kgs	Large (L)	2.68ml
40 – 60 kgs	Extra large (XL)	4.02ml

Method of administration

Remove one pipette from the foil pack. Hold the pipette upright. Tap the narrow part of the pipette to ensure the contents remain within the main body of the pipette. Snap back the tip. Part the coat on the back of the animal at the base of the neck in front of the shoulder blades until the skin is visible. Place pipette tip on the skin. Squeeze pipette, applying entire contents in a single spot to the animal's skin. Avoid superficial application to the animal's hair. Only one pipette per treatment is needed. Temporary changes to the coat (clumped/greasy hair) may be noted at the site of application.

Contraindications

The product should not be used in puppies less than 8 week old and/or weighing less than 2 kgs. Do not use on sick, (e.g. systemic disease, fever) or convalescent animals.

Do not use in rabbits, as adverse reaction with even mortality could occur.

This product is specifically developed for dogs. Do not use in cats, as this could lead to overdosing. In the absence of safety studies, the minimum treatment interval is 4 weeks.

Adverse reaction

Among the very rare suspected adverse reaction, transient skin reactions on the application site (skin discoloration, local hair loss, itching, redness) and general itching or hair loss have been reported after use. Excessive salivation, reversible nervous signs (increased sensitivity to stimulation, depression, other nervous signs), vomiting or respiratory symptoms have been observed after use. If licking occurs a brief period of excessive salivation may be observed due mainly to the nature of the carrier. Do not overdose. If you notice any serious effects or other effects not mentioned here, please inform your veterinary surgeon.

Overdose

No adverse effects were observed in target animal safety studies in 8 week-old puppies, growing dogs and dogs weighing above 2kgs treated once at 5 times the recommended dose. The risk of experiencing adverse effects may however increase when overdosing, so animals always be treated with the correct pipette size according to the body weight.

Special warnings for target species

Bathing/immersion in water within 2 days after application of the product and more frequent bathing than once a week should be avoided, as no study has been performed to investigate how this affects the efficacy of the product. Emollient shampoos can be used prior to treatment, but reduce the duration of protection against fleas to approximately 5 weeks when used weekly after application of the product. Weekly bathing with a 2% chlorhexidine medicated shampoo did not affect efficacy against fleas during a 6 week long study.



927222

PACK OPS LYON GRAPHIC ARTWORK APPROVAL



Code: 0 053 927222	VERSION PTG : 0037 Version : A 180316	COLORS
PRODUCT: NOT FTL PLUS CHIEN X3 RECTO	INITIALS: CS - Composer RH VERSION - DATE: A 23/03/21 11h00 B 20/05/21 15h00	Noir/Black
COUNTRY: INDE (IN) Dimensions à plat (mm): 240 x 80 Dimensions plié(e)s: 120 x 80	CANCEL AND REPLACE: 0 053 920977	! Please use the official PANTONE® (P) matching system for an accurate color representation.
PRINTED BASIS CODE (if applicable): NA		

Marketing approval:

COMPLETE NAME:

DATE + SIGNATURE:

Regulatory approval:

COMPLETE NAME:

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Dogs should not be allowed to swim in watercourses for 2 days after application. There may be an attachment of a few ticks. For this reason a transmission of infectious disease cannot be completely excluded if conditions are unfavorable.

Fleas from pets often infest the animal's basket, bedding and regular resting areas such as carpets and soft furnishings which should be treated, in case of massive infestation and at the beginning of the control measures, with a suitable insecticide and vacuumed regularly.

Avoid contact with animal eyes.

It is important to make sure that the product is applied to an area where the animal cannot lick it off and to make sure that animals do not lick each other following treatment.

Special precautions to be taken by person administering the veterinary medicinal product to animals
This product can cause mucous membrane, skin and eye irritation. Therefore, contact of the product with mouth, skin and eyes should be avoided.

Animals or operators with a known sensitivity to insecticides or alcohol should avoid contact with Frontline spot-on for dogs. Avoids contents coming in contact with fingers. If this occurs, wash hands with soap and water.

After accidental ocular exposure the eye should be rinsed with pure water.

Wash hands after use.

Treated animals not to be handled until the application site is dry, and children should not be allowed to play with treated animals until the application site is dry. It is therefore recommended that animals are not treated during the day, but should be treated during the early evening, and that recently treated animals are not allowed to sleep with owners especially children.

Do not smoke, drink or eat during application.

Pharmacological properties
The product is an insecticidal and acaricidal solution for topical use, containing an association of an adulticidal active ingredient, fipronil, in combination with an ovicidal and larvicidal active ingredient, (S)-methoprene.

Pharmacodynamic properties
Fipronil is an insecticide and acaricide belonging to the phenylpyrazole family. It acts by interacting with ligand-gated chloride channels, in particular those gated by the neurotransmitter gamma-aminobutyric acid (GABA), thereby blocking pre- and post-synaptic transfer of chloride ions across cell membranes. This results in uncontrolled activity of the central nervous system and death of insects or acarines. Fipronil kills fleas within 24 hours and ticks (*Dermacentor reticulatus*, *Dermacentor variabilis*, *Rhipicephalus sanguineus*, *Ixodes scapularis*, *Ixodes ricinus*, *Haemaphysalis longicornis*, *Haemaphysalis flava*, *Haemaphysalis campanulata*) and lice within 48 hours post-exposure.

(S)-Methoprene is an insect growth regulator (IGR) of the class of compounds known as juvenile hormone analogues that inhibit the development of immature stages of insects. This compound mimics the action of juvenile hormone and causes impaired development and death of the developing stages of fleas. The on-animal ovicidal activity of (S)-methoprene results from either direct penetration of the eggshell of newly laid eggs or from absorption through the cuticle of the adult fleas. (S)-methoprene is also effective in preventing flea larvae and pupae from developing, which prevents contamination of the environment of treated animals with the immature stages of fleas.

Pharmacokinetic particulars
Studies of metabolism of fipronil have demonstrated that the major metabolite is the sulfone derivate of fipronil.

(S)-methoprene is extensively degraded into carbon dioxide and acetate that are subsequently incorporated into endogenous materials.

The pharmacokinetic profiles after topical application of fipronil and (S)-methoprene in combination were studied in dogs in comparison to intravenous dosing of fipronil or (S)-methoprene alone. This established absorption and other pharmacokinetic parameters. The topical application resulted in low systemic absorption of fipronil (11%) with a mean maximum concentration (C_{max}) of approximately 35ng/ml fipronil and 55ng/ml of fipronil sulfone in plasma.

Peak fipronil plasma concentrations are slowly attained (mean t_{max} approximately 101 h), and decline slowly (mean terminal half-life approximately 154 h, highest values are observed for males).

Fipronil is extensively metabolized to fipronil sulfone after topical administration.

Plasma concentration of (S)-methoprene were below the limit of quantitation (20 ng/ml) in dogs after topical application.

Both (S)-methoprene and fipronil, together with its major metabolite are well-distributed in the haircoat of a dog within one day after application. The concentration of fipronil, fipronil sulfone and (S)-methoprene in the hair coat decrease with time and are detectable for at least 60 days after dosing. Parasites are killed through contact rather than systemic exposure.

No pharmacological interaction between fipronil and (S)-methoprene was noted.

Use during pregnancy and lactation
The product can be used during pregnancy and lactation.

Interaction with other medicinal products and incompatibilities: None known

Storage
Do not store above 30°C. Store in the original package. Do not use after the expiry date stated on the package after EXP. The expiry date refers to the last day of the month.

Special precautions for the disposal of unused veterinary medicinal product or waste materials derived from the use of such products.
Any unused product or waste material should be disposed of in accordance with national requirement. Fipronil and (S)-methoprene may adversely affect aquatic organisms. Do not contaminate ponds, waterways or ditches with the product or empty containers.

Shelf life: 36 months

Withdrawal period: Not applicable

Manufactured by:
M/s. Boehringer Ingelheim Animal Health France SCS, 4 chemin du calquet, 31000 Toulouse, France

Importer:
M/s. Boehringer Ingelheim India Pvt Ltd, Bld No.D-5, Gala No: 1 to 5, Shree Arihant Compound Ground Floor, Reti Bunder Road, Kalher, Bhiwandi, Tal: Bhiwandi - 15 (Thane-Zone5), Pin: 421302, India

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