PARTICULARS TO APPEAR ON THE OUTER PACKAGE

{BOX}

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

AMFLEE combo 402 mg/ 361.8 mg spot-on solution for extra large dogs
Fipronil/S-Methoprene

2. STATEMENT OF ACTIVE SUBSTANCES

Each 4.02 ml pipette contains:
Active substance:

Fipronil	402 mg
S-Methoprene	361.8 mg

3. PHARMACEUTICAL FORM

Spot-on solution.

4. PACKAGE SIZE

1 x 4.02 ml
3 x 4.02 ml
6 x 4.02 ml
10 x 4.02 ml
30 x 4.02 ml

5. TARGET SPECIES

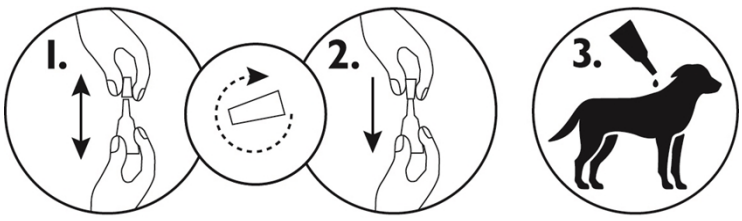
DOGS OVER 40 KG

6. INDICATION(S)

For use against flea infestations, alone or in association with ticks and/or biting lice.

7. METHOD AND ROUTE(S) OF ADMINISTRATION

Read the package leaflet before use.
External use only.
Spot-on use.



8. WITHDRAWAL PERIOD

9. SPECIAL WARNING(S), IF NECESSARY

Read the package leaflet before use.

10. EXPIRY DATE

EXP

11. SPECIAL STORAGE CONDITIONS

Store in the original package in order to protect from light and moisture.

12. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCTS OR WASTE MATERIALS, IF ANY

Disposal: read package leaflet.
Fipronil and S-methoprene may adversely affect aquatic organisms. Do not contaminate ponds, waterways or ditches with the product or empty container.

13. THE WORDS "FOR ANIMAL TREATMENT ONLY" AND CONDITIONS OR RESTRICTIONS REGARDING SUPPLY AND USE, IF APPLICABLE

For animal treatment only.
Only for those countries where the product is available on prescription:
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

KRKA, d.d., Novo mesto
Šmarješka cesta 6
8501 Novo mesto
Slovenia

16. MARKETING AUTHORISATION NUMBER

Vm 01656/4112

17. MANUFACTURER'S BATCH NUMBER

Lot

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

{ BAG }

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

AMFLEE combo 402 mg/ 361.8 mg spot-on solution for extra large dogs
Fipronil/S-Methoprene

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

Each 4.02 ml pipette contains:
Active substance:

Fipronil	402 mg
S-Methoprene	361.8 mg

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

1 x 4.02 ml

4. ROUTE(S) OF ADMINISTRATION

Spot-on use.
DOGS OVER 40 KG

5. WITHDRAWAL PERIOD

6. BATCH NUMBER

Lot:

7. EXPIRY DATE

EXP

8. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.

MINIMUM PARTICULARS TO APPEAR ON BLISTERS OR STRIPS

{ PIPETTE }

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

AMFLEE combo 402 mg/ 361.8 mg
Fipronil/S-Methoprene

2. NAME OF THE MARKETING AUTHORISATION HOLDER

KRKA

3. EXPIRY DATE

EXP

4. BATCH NUMBER

Lot

5. THE WORDS "FOR ANIMAL TREATMENT ONLY"

PACKAGE LEAFLET:
AMFLEE combo 67 mg/ 60.3 mg spot-on solution for small dogs
AMFLEE combo 134 mg/ 120.6 mg spot-on solution for medium dogs
AMFLEE combo 268 mg/ 241.2 mg spot-on solution for large dogs
AMFLEE combo 402 mg/ 361.8 mg spot-on solution for extra large 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:
KRKA, d.d., Novo mesto, Šmarješka cesta 6, 8501 Novo mesto, Slovenia

Manufacturer responsible for batch release:
KRKA, d.d., Novo mesto, Šmarješka cesta 6, 8501 Novo mesto, Slovenia
TAD Pharma GmbH, Heinz-Lohmann-Straße 5, 27472 Cuxhaven, Germany

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

AMFLEE combo 67 mg/ 60.3 mg spot-on solution for small dogs
AMFLEE combo 134 mg/ 120.6 mg spot-on solution for medium dogs
AMFLEE combo 268 mg/ 241.2 mg spot-on solution for large dogs
AMFLEE combo 402 mg/ 361.8 mg spot-on solution for extra large dogs

Fipronil/S-Methoprene

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

Each 0.67 ml pipette contains:
Active substance:

Fipronil	67 mg
S-Methoprene	60.30 mg

Excipients:

Butylhydroxyanisole (E320)	0.134 mg
Butylhydroxytoluene (E321)	0.067 mg

Each 1.34 ml pipette contains:
Active substance:

Fipronil	134 mg
S-Methoprene	120.60 mg

Excipients:

Butylhydroxyanisole (E320)	0.27 mg
Butylhydroxytoluene (E321)	0.13 mg

Each 2.68 ml pipette contains:
Active substance:

Fipronil	268 mg

S-Methoprene	241.20 mg

Excipients:

Butylhydroxyanisole (E320)	0.54 mg
Butylhydroxytoluene (E321)	0.27 mg

Each 4.02 ml pipette contains:
Active substance:

Fipronil	402 mg
S-Methoprene	361.80 mg

Excipients:

Butylhydroxyanisole (E320)	0.80 mg
Butylhydroxytoluene (E321)	0.40 mg

Clear yellow solution.

4. INDICATION(S)

For the treatment of dogs, dosage defined by bodyweight grouping (refer to section 8.):
- To be used against infestations 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 eight weeks after application.
- Treatment of tick infestations (*Ixodes ricinus, Dermacentor variabilis, Dermacentor reticulatus, Rhipicephalus sanguineus*). The product has persistent acaricidal efficacy for up to 4 weeks against ticks.
- 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), where this has been previously diagnosed by a veterinary surgeon.

5. CONTRAINDICATIONS

In the absence of supporting data, this product should not be used on puppies less than 8 weeks old.
Always establish the appropriate dosage according to bodyweight before using this product. In the absence of supporting data, the product for dogs of 2-10 kg bodyweight should not be used in dogs weighing less than 2 kg.
Do not use on sick (e.g. systemic diseases, fever) or convalescent animals.
Do not use in rabbits due to a risk of adverse reactions or even death.
This product is specifically developed for dogs. Do not use in cats and ferrets, as this could lead to overdosing.
Do not use in case of hypersensitivity to the active substance or to any of the excipients.

6. ADVERSE REACTIONS

Among the very rare suspected adverse reactions, 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 also been observed after use.

In the case of licking the administration site, a brief period of excessive salivation may be observed due mainly to the nature of the carrier.

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.

Alternatively you can report via your national reporting system {national system details}.

7. TARGET SPECIES

Dogs 2-10 kg
Dogs 10-20 kg
Dogs 20-40 kg
Dogs over 40 kg

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

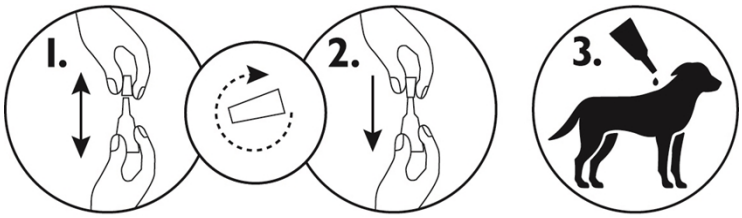
External use only, spot-on use.
Administer by topical application to the skin according to the bodyweight as follows:

Dogs	**Number of pipettes**	**Pipette volume**	**Potency (fipronil + (S)-methoprene**
over 2 kg and up to 10 kg	1 pipette	0.67 ml	67 mg + 60.3 mg
over 10 kg and up to 20 kg	1 pipette	1.34 ml	134 mg + 120.6 mg
over 20 kg and up to 40 kg	1 pipette	2.68 ml	268 mg + 241.2 mg
over 40 kg	1 pipette	4.02 ml	402 mg + 361.8 mg

This corresponds to a minimum recommended dose of 6.7 mg/kg for fipronil and 6 mg/kg for (S)-methoprene, by topical application to the skin.

Method of administration:

1. Remove the pipette from its packaging. Hold the pipette in an upright position, twist and pull the cap off.
2. Turn the cap around and place the other end of the cap back on the pipette. Push and twist the cap to break the seal, then remove the cap from the pipette.
3. 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 the tip of the pipette onto the skin and squeeze the pipette several times to empty its contents completely and directly onto the skin in one spot.



9. ADVICE ON CORRECT ADMINISTRATION

In the absence of safety studies the minimum treatment interval is 4 weeks.
Temporary changes to the coat (clumped/greasy hair) may be noted at the application site.

10. WITHDRAWAL PERIOD

Not applicable.

11. SPECIAL STORAGE PRECAUTIONS

Keep out of the sight and reach of children.

Store in the original package in order to protect from light and moisture.
This veterinary medicinal product does not require any special temperature storage conditions.

Do not use this veterinary medicinal product after the expiry date which is stated on the label after {EXP}.The expiry date refers to the last day of that month.

12. SPECIAL WARNING(S)

Special warnings for each 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.

For the treatment and control of flea allergy dermatitis it is recommended that allergic patients and all other animals in the household should be treated on a regular basis. 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.

Special precautions for use in animals:
The attachment of single ticks after treatment cannot be ruled out. Therefore the transmission of infectious disease cannot be completely excluded if conditions are unfavourable.
For external use only. Do not administer orally.
Avoid contact with the eyes of the animal. If the product is in contact with eyes, rinse immediately with plenty of water.
Do not apply the product on wounds or skin lesions.
It is important to make sure that the veterinary medicinal product is applied directly onto an area of dry skin where the animal cannot lick it off and to make sure that animals do not lick each other following treatment.
Dogs should not be allowed to swim in watercourses for 2 days after application.
Wait for the application site to dry before allowing the treated dog to come into contact with valuable fabrics or furnishings.

Pregnancy and lactation:
The product can be used during pregnancy and lactation.

Overdose (symptoms, emergency procedures, antidotes):
No adverse effects were observed in target animal safety studies in 8 week-old puppies, growing dogs and dogs weighing about 2 kg treated once at five times the recommended dose. The risk of experiencing adverse effects (see section 6) may however increase when overdosing, so animals should always be treated with the correct pipette size according to bodyweight.

Special precautions to be taken by the 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.
People with known hypersensitivity to fipronil or s-methoprene and/or any of the ingredients should avoid contact with the veterinary medicinal product.
Avoid contents coming into contact with the fingers. If this occurs, wash hands with soap and water.
After accidental ocular exposure the eye should be rinsed carefully in clean water.
Wash hands after use.
In case of accidental ingestion seek medical advice immediately and show the package leaflet or the label to the physician.
Do not smoke, drink or eat during application.
Treated animals should not 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.

Keep pipettes in the original packaging until ready to use, and dispose of used pipettes immediately.

13. SPECIAL PRECAUTIONS FOR THE DISPOSAL OF UNUSED PRODUCT OR WASTE MATERIALS, IF ANY

Any unused veterinary medicinal product or waste materials derived from such veterinary medicinal product should be disposed of in accordance with local requirements. The product should not enter water courses as this may be dangerous for fish and other aquatic organisms.

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

February 2021

15. OTHER INFORMATION

Cardboard box containing 1, 3, 6, 10, 30 pipettes.
Not all pack sizes may be marketed.

For any information about this veterinary medicinal product, please contact the local representative of the marketing authorisation holder.

UK only

POM-V For animal treatment only. To be supplied only on veterinary prescription.
Vm 01656/4114
Vm 01656/4110
Vm 01656/4111
Vm 01656/4112

Local representative:
KRKA UK Ltd
United Kingdom
Tel: 02071 646 156
pharmacovigilance.uk@krka.biz

Approved 22 April 2021