ANNEX I

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

DAMTIX 200 mg/40 mg spot-on solution for dogs up to 4 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 200 mg/40 mg spot-on solution for dogs up to 4 kg (FI)
Permetrix biocanina 200 mg/40 mg spot-on solution for dogs up to 4 kg (FR)

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 0.4 ml pipette contains:
Active substances:
Permethrin (40:60) 200.0 mg
Imidacloprid 40.0 mg

Excipients:
Butylhydroxytoluene (E321) 0.4 mg
N-methylpyrrolidone 80.0 mg

For the full list of excipients, see section 6.1.

3. PHARMACEUTICAL FORM

Spot-on solution.
Clear yellowish to brownish solution.

4. CLINICAL PARTICULARS

4.1 Target species

Dogs

4.2 Indications for use, specifying the target species

For the treatment and prevention of flea (*Ctenocephalides felis*) infestation.

Fleas on dogs are killed within one day following treatment. One treatment prevents further flea infestation for four weeks. The product can be used as part of a treatment strategy for flea allergy dermatitis (FAD) where this has been previously diagnosed by a veterinary surgeon.

The product has persistent acaricidal efficacy against tick infestations (*Rhipicephalus sanguineus* and *Ixodes ricinus* for four weeks, and *Dermacentor reticulatus* for three weeks) and persistent repellent efficacy (*Ixodes ricinus*) for three weeks .

Ticks already on the dog may not be killed within two days after treatment and may remain attached and visible. Therefore the removal of ticks already on the dog at the time of treatment is recommended, in order to prevent them from attaching and having a blood meal.

One treatment provides repellent (anti-feeding) activity against the sand fly *Phlebotomus perniciosus* for three weeks and against the mosquito *Aedes aegypti* from 7 days up to 14 days after treatment.

Reduction of the risk of infection with *Leishmania infantum* via transmission by sandflies (*Phlebotomus perniciosus*) for up to 3 weeks. The effect is indirect due to product's activity against the vector.

4.3 Contraindications

In the absence of available data do not use the product on puppies of less than 7 weeks of age, or 1.5 kg of weight.
Do not use in known cases of hypersensitivity to the active substances or to any of the excipients.
Do not use on cats. (Refer to section 4.5 – Special precautions for use).

4.4 Special warnings for each target species

There may be an attachment of single ticks or bites by single sand flies or mosquitoes. For this reason, a transmission of infectious diseases cannot be excluded if conditions are unfavourable.
As the product exerts a repellent (anti-feeding) activity against *Aedes aegypti* mosquitoes 7 days after treatment, the product should preferably be applied 1 week before animals are likely to become exposed to these mosquitoes.
The product remains effective against fleas if the animal becomes wet. After weekly immersions in water for one minute the period of persistent insecticidal efficacy against fleas was not reduced. However, prolonged, intense exposure to water should be avoided. In cases of frequent and/or prolonged water exposure the persistent efficacy may be reduced. In these cases do not retreat more frequently than once weekly. If a dog requires a shampoo, it should be administered before applying the product or at least 2 weeks after application, to optimise efficacy of the product.
The effectiveness of the product against ticks, and the repellent effectiveness of the product following swimming or shampooing has not been investigated.
Immediate protection against sandflies bites is not documented. Treated dogs for the reduction of the risk of infection with *Leishmania infantum* via transmission by sandflies *P. perniciosus* should be kept in a protected environment during the first 24 hours after the initial treatment application.

4.5 Special precautions for use

Special precautions for use in animals

Care should be taken to avoid the content of the pipette coming into contact with the eyes or mouth of the recipient dogs.

Care should be taken to administer the product correctly as described under section 4.9. In particular oral uptake due to the licking of the application site by treated or in-contact animals should be avoided.

Do not use on cats.

This product is extremely poisonous to cats and could be fatal due to the unique physiology of cats which is unable to metabolise certain compounds including permethrin. To prevent cats from being accidentally exposed to the product, keep treated dogs away from cats after treatment until the application site is dry. It is important to ensure that cats do not groom the site of application on a dog, which has been treated with this product. Seek veterinary advice immediately if this occurs.

Consult your veterinary surgeon before using the product on sick and debilitated dogs.

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

Avoid contact between the product and skin, eyes or mouth.
Do not eat, drink or smoke during application.
Wash hands thoroughly after use.
In case of accidental spillage onto skin, wash off immediately with soap and water.
People with known skin sensitivity may be particularly sensitive to this product.
The predominant clinical symptoms that in extremely rare case may be shown are transient sensory irritations of the skin like tingling, burning sensation or numbness.
If the product gets accidentally into the eyes, they should be thoroughly flushed with water. If skin or if eye irritation persists, seek medical advice immediately and show the package leaflet or the label to the physician.
Do not ingest. In case of accidental ingestion seek medical advice immediately and show the package leaflet or the label to the physician.
Treated dogs should not be handled especially by children until the application site is dry. This may be ensured by treating the dogs. in the evening. Recently treated dogs should not be allowed to sleep together with their owner, especially children.
Keep the stored pipettes in the original blister. In order to prevent children from getting access to used pipettes, used pipettes should be disposed of immediately.

Other precautions

The solvent in the product may stain certain materials including leather, fabrics, plastics and finished surfaces. Allow the application site to dry before permitting contact with such materials.
The product is toxic for aquatic organisms. Treated dogs must not under any circumstances be allowed into any type of surface water for at least 48 hours after treatment.

4.6 Adverse reactions (frequency and seriousness)

Pruritus, hair loss, erythema, oedema, and erosions may occur in very rare cases at the application site and are generally self-resolving.

Behaviour changes (agitation, restlessness, whining or rolling), gastro-intestinal symptoms (vomiting, diarrhoea, hypersalivation, diminished appetite) and neurological signs such as unsteady movement and twitching, or lethargy in dogs susceptible to the ingredient permethrin may be observed in very rare cases. These signs are generally transient and self-resolving.

Accidental oral uptake may result in transient vomiting and neurological signs such as tremor and incoordination. Treatment should be symptomatic. There is no known specific antidote.

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).

4.7 Use during pregnancy, lactation or lay

The safety of the veterinary medicinal product has not been established in dogs during pregnancy, lactation or in animals intended for breeding. Laboratory studies in rabbits and rats with the excipient N-methyl pyrrolidone have shown evidence of foetotoxic effects. Use only according to the benefit-risk assessment by the responsible veterinarian.

4.8 Interaction with other medicinal products and other forms of interaction

None known.

4.9 Amounts to be administered and administration route

Route of administration and dosage:

Spot-on use only. Apply only to undamaged skin.

The recommended minimum dose is:
10 mg/kg body weight (bw) imidacloprid and 50 mg/kg body weight (bw) permethrin.

Administer by topical application to the skin according to the bodyweight as follows:

Dogs (kg body weight)	**Strength**	**Volume (ml)**	**Imidacloprid (mg/kg body weight)**	**Permethrin (mg/kg body weight)**
≤ 4 kg	200 mg/40 mg spot-on solution for dogs up to 4 kg	0.4 ml	minimum of 10	minimum of 50
>4 kg ≤ 10 kg	500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg	1.0 ml	10 - 25	50 - 125
>10 kg ≤ 25 kg	1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg	2.5 ml	10 - 25	50 - 125
>25 kg ≤ 40 kg	2000 mg/400 mg spot-on solution for dogs over 25 kg	4.0 ml	10 - 16	50 - 80

For dogs > 40 kg the appropriate combination of pipettes should be used.

To ensure correct dosage, body weight should be determined as accurately as possible.

To reduce re-infestation from emergence of new fleas, it is recommended to treat all dogs in a household. Other animals living in the same household should also be treated with a suitable product. To aid further in reducing environmental challenge, the additional use of a suitable environmental treatment against adult fleas and their developmental stages is recommended.

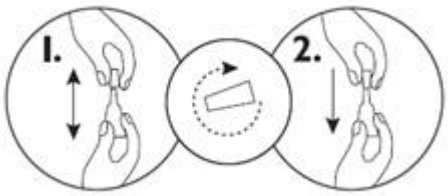
Depending on the ectoparasite challenge, it may be necessary to repeat the treatment. The interval between two treatments should be 4 weeks. However, in cases of frequent and/or prolonged water exposure the persistent efficacy may be reduced. In these cases do not retreat more frequently than once weekly.

To protect a dog over the whole sand fly season, treatment should be compliantly continued throughout.

Transient cosmetic changes (e.g. skin scaling, white deposits and spiking of the hair) may be observed at application sites.

Method of administration:

Remove one pipette from the package. Hold applicator pipette in an upright position. Tap the narrow part of pipette to ensure the contents are within the main body of the pipette, twist and pull cap off. Turn the cap around and place the other end of cap back on pipette. Push and twist the cap to break seal, and then remove the cap from the pipette.



For dogs 10 kg body weight or less:

With the dog standing still, part the coat between the shoulder blades until the skin is visible. Place the tip of the pipette on the skin and squeeze firmly several times to empty the contents directly onto the skin.

For dogs of more than 10 kg body weight:

With the dog standing still, the entire contents of the pipette should be applied evenly to four spots on the top of the back from the shoulder to the base of the tail. At each spot, part the hair until the skin is visible. Place the tip of the pipette on the skin and gently squeeze to expel a portion of the solution on the skin. Do not apply an excessive amount of solution at any one spot that could cause some of the solution to run off the side of the dog.

4.10 Overdose (symptoms, emergency procedures, antidotes), if necessary

No adverse clinical signs were noted in healthy puppies or adult dogs exposed to 5x overdosage or for puppies whose mothers were treated with 3x overdosage of the combination of imidacloprid and permethrin. The severity of skin erythema, which sometimes occurs at the application site, increases with overdose.

4.11 Withdrawal period(s)

Not applicable.

5. PHARMACOLOGICAL PROPERTIES

Pharmacotherapeutic group: Ectoparasiticides for topical use, incl. Insecticides, permethrin, combinations.
ATCvet code: QP53AC54.

5.1 Pharmacodynamic properties

The product is an ectoparasiticide for topical use containing imidacloprid and permethrin. This combination acts as an insecticide and acaricide.

Imidacloprid is an ectoparasiticide belonging to the chloronicotinyl group of compounds. Chemically, it can be classified as a chloronicotinyl nitroguanidine. Imidacloprid is effective against adult fleas and larval flea stages. In addition to the adulticide flea efficacy of imidacloprid, a larvicidal flea efficacy in the surroundings of the treated pet has been demonstrated. Larval stages in the dog`s immediate surroundings are killed following contact with a treated animal. It has a high affinity for the nicotinergic acetylcholine receptors in the post-synaptic region of the central nervous system (CNS) in insects. The ensuing inhibition of cholinergic transmission in insects results in paralysis and death of the parasite.

Permethrin belongs to the type I class of pyrethroid acaricides and insecticides. Pyrethroids affect the voltage-gated sodium channels in vertebrates and non-vertebrates. Pyrethroids are so called “open channel blockers” affecting the sodium channel by slowing both the activation and the inactivation properties thus leading to hyperexcitability and death of the parasite.

In the combination of both substances, it has been shown that imidacloprid functions as the activator of arthropod ganglion and therefore increases the efficacy of permethrin.

5.2 Pharmacokinetic particulars

Following topical application in dogs, the solution distributes over the body surface of the animal. Both active substances remain on the skin and hair of the treated animal for at least 4 weeks. Systemic absorption of the product is sufficiently low so as not to affect efficacy or target species tolerance.

Environmental properties

The product should not be allowed to enter water courses as this may be dangerous for fish and aquatic organisms. For treated dogs, please see section 4.5.

Imidacloprid and/or permethrin containing products are toxic to honey bees.

6. PHARMACEUTICAL PARTICULARS

6.1 List of excipients

Butylhydroxytoluene (E321)
Triglycerides, Medium-Chain

N-methylpyrrolidone
Citric acid (E330)
Dimethyl sulfoxide

6.2 Major incompatibilities

None known.

6.3 Shelf life

Shelf life of the veterinary medicinal product as packaged for sale: 3 years.

6.4. Special precautions for storage

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

6.5 Nature and composition of immediate packaging

White polypropylene pipette closed with a polyethylene (HDPE) cap. Each pipette is packed in a polyethylene terephthalate/aluminium/low density polyethylene triplex bag.
1 ml pipette containing 0.4 ml of solution
Box containing 1, 3, 4, 6, 10 pipettes

Not all pack sizes may be marketed.

6.6 Special precautions for the disposal of unused veterinary medicinal product or waste materials derived from the use of such products, if appropriate

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

7. MARKETING AUTHORISATION HOLDER

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

8. MARKETING AUTHORISATION NUMBER(S)

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of first authorisation:
Date of last renewal:

10 DATE OF REVISION OF THE TEXT

PROHIBITION OF SALE, SUPPLY AND/OR USE

Not applicable.

ANNEX III

LABELLING AND PACKAGE LEAFLET

A. LABELLING

PARTICULARS TO APPEAR ON THE OUTER PACKAGE

BOX

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

DAMTIX 200 mg/40 mg spot-on solution for dogs up to 4 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 200 mg/40 mg spot-on solution for dogs up to 4 kg (FI)
Permetrix biocanina 200 mg/40 mg spot-on solution for dogs up to 4 kg (FR)
Permethrin (40:60)/Imidacloprid

2. STATEMENT OF ACTIVE SUBSTANCES

Each 0.4 ml pipette contains:
Active substances:
Permethrin (40:60) 200.0 mg
Imidacloprid 40.0 mg

3. PHARMACEUTICAL FORM

Spot-on solution

4. PACKAGE SIZE

1 unit-dose pipette of 0.4 ml
3 unit-dose pipettes of 0.4 ml
4 unit-dose pipettes of 0.4 ml
6 unit-dose pipettes of 0.4 ml
10 unit-dose pipettes of 0.4 ml

1 x 0.4 ml
3 x 0.4 ml
4 x 0.4 ml
6 x 0.4 ml
10 x 0.4 ml

5. TARGET SPECIES

Dogs

6. INDICATION(S)

EXTERNAL PARASITE CONTROL.

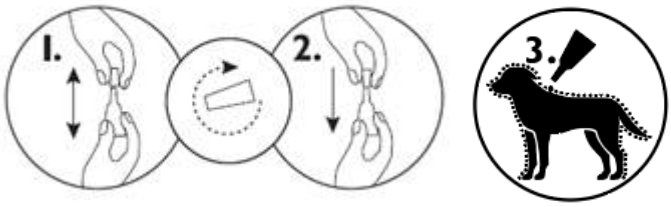
For OTC products:
- Eliminates ticks and fleas.
- Repels ticks, mosquitos and sand flies.

- Reduction of the risk of transmission of canine leishmaniosis

7. METHOD AND ROUTE(S) OF ADMINISTRATION

****Only for those countries where the product is available without prescription**.
1 pipette per dog weighing up to 4 kg

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



8. WITHDRAWAL PERIOD(S)

9. SPECIAL WARNING(S), IF NECESSARY

Read the package leaflet before use.
Do not use on cats.

10. EXPIRY DATE

EXP

11. SPECIAL STORAGE CONDITIONS

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

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

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

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

For animal treatment only.

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(S)

17. MANUFACTURER'S BATCH NUMBER

Lot

 *Ixodida*

 *Ctenocephalides felis*

 *Ctenocephalides felis larvae*

 *Phlebotomus perniciosus*

 *Aedes aegypti*

 *Leishmania infantum*

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

LABEL

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

DAMTIX 200 mg/40 mg spot-on solution for dogs up to 4 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 200 mg/40 mg spot-on solution for dogs up to 4 kg (FI)
Permetrix biocanina 200 mg/40 mg spot-on solution for dogs up to 4 kg (FR)
Permethrin (40:60)/Imidacloprid

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

Each 0.4 ml pipette contains:
Active substances:
Permethrin (40:60) 200.0 mg
Imidacloprid 40.0 mg

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

1 unit-dose pipette of 0.4 ml

1 x 0.4 ml

4. ROUTE(S) OF ADMINISTRATION

Spot-on use.

5. WITHDRAWAL PERIOD(S)

6. BATCH NUMBER

Lot

7. EXPIRY DATE

EXP

8. THE WORDS “FOR ANIMAL TREATMENT ONLY”

For animal treatment only.
Do not use on cats.

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

PIPETTE

1. NAME OF THE VETERINARY MEDICINAL PRODUCT

DAMTIX (IE, DE, EL, ES, IT, PT)
ATAXXA VET (FI)
Permetrix biocanina (FR)

Permethrin (40:60)/Imidacloprid
Permethrin (40:60)/Imidacloprid (for multilingual packaging)

2. QUANTITY OF THE ACTIVE SUBSTANCE(S)

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

0.4 ml

4. ROUTE(S) OF ADMINISTRATION

5. WITHDRAWAL PERIOD(S)

6. BATCH NUMBER

Lot

7. EXPIRY DATE

EXP

8. THE WORDS "FOR ANIMAL TREATMENT ONLY"

B. PACKAGE LEAFLET

PACKAGE LEAFLET:
DAMTIX 200 mg/40 mg spot-on solution for dogs up to 4 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 200 mg/40 mg spot-on solution for dogs up to 4 kg (FI)
Permetrix biocanina 200 mg/40 mg spot-on solution for dogs up to 4 kg (FR)

DAMTIX 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (FI)
Permetrix biocanina 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (FR)

DAMTIX 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (FI)
Permetrix biocanina 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (FR)

DAMTIX 2000 mg/400 mg spot-on solution for dogs over 25 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 2000 mg/400 mg spot-on solution for dogs over 25 kg (FI)
Permetrix biocanina 2000 mg/400 mg spot-on solution for dogs over 25 kg (FR)

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

Marketing authorisation holder and manufacturer responsible for batch release:
KRKA, d.d., Novo mesto
Šmarješka cesta 6
8501 Novo mesto
Slovenia

2. NAME OF THE VETERINARY MEDICINAL PRODUCT

DAMTIX 200 mg/40 mg spot-on solution for dogs up to 4 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 200 mg/40 mg spot-on solution for dogs up to 4 kg (FI)
Permetrix biocanina 200 mg/40 mg spot-on solution for dogs up to 4 kg (FR)

DAMTIX 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (FI)
Permetrix biocanina 500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg (FR)

DAMTIX 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (FI)
Permetrix biocanina 1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg (FR)

DAMTIX 2000 mg/400 mg spot-on solution for dogs over 25 kg (IE, DE, EL, ES, IT, PT)
ATAXXA VET 2000 mg/400 mg spot-on solution for dogs over 25 kg (FI)
Permetrix biocanina 2000 mg/400 mg spot-on solution for dogs over 25 kg (FR)

Permethrin (40:60)/Imidacloprid

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

Each 0.4 ml pipette contains:
Active substances:
Permethrin (40:60) 200.0 mg
Imidacloprid 40.0 mg

Excipients:
Butylhydroxytoluene (E321) 0.4 mg
N-methylpyrrolidone 80.0 mg

Each 1.0 ml pipette contains:
Active substances:
Permethrin (40:60) 500.0 mg
Imidacloprid 100.0 mg

Excipients:
Butylhydroxytoluene (E321) 1.0 mg
N-methylpyrrolidone 200.0 mg

Each 2.5 ml pipette contains:
Active substances:
Permethrin (40:60) 1250.0 mg
Imidacloprid 250.0 mg

Excipients:
Butylhydroxytoluene (E321) 2.5 mg
N-methylpyrrolidone 500.0 mg

Each 4.0 ml pipette contains:
Active substances:
Permethrin (40:60) 2000.0 mg
Imidacloprid 400.0 mg

Excipients:
Butylhydroxytoluene (E321) 4.0 mg
N-methylpyrrolidone 800.0 mg

Clear yellowish to brownish solution.

4. INDICATION(S)

For the treatment and prevention of flea (*Ctenocephalides felis*) infestation. 

Fleas on dogs are killed within one day following treatment. One treatment prevents further flea infestation for four weeks. The product can be used as part of a treatment strategy for flea allergy dermatitis (FAD) where this has been previously diagnosed by a veterinary surgeon.

The product has persistent acaricidal efficacy against tick infestations (*Rhipicephalus sanguineus* and *Ixodes ricinus* for four weeks, and *Dermacentor reticulatus* for three weeks) and persistent repellent efficacy (*Ixodes ricinus*) for three weeks. 

Ticks already on the dog may not be killed within two days after treatment and may remain attached and visible. Therefore the removal of ticks already on the dog at the time of treatment is recommended, in order to prevent them from attaching and having a blood meal.

One treatment:
- provides repellent (anti-feeding) activity against the sand fly *Phlebotomus perniciosus* for three weeks, 
- provides repellent (anti-feeding) activity against the mosquito *Aedes aegypti* from 7 days up to 14 days after treatment. 

Reduction of the risk of infection with *Leishmania infantum* via transmission by sandflies (*Phlebotomus perniciosus*) for up to 3 weeks. The effect is indirect due to product's activity against the vector. 

5. CONTRAINDICATIONS

In the absence of available data do not use the product on puppies of less than 7 weeks of age, or 1.5 kg of weight (product for dogs up to 4 kg), 4 kg of weight (product for dogs over 4 kg up to 10 kg), 10 kg of weight (product for dogs over 10 kg up to 25 kg), 25 kg of weight (product for dogs over 25 kg).
Do not use in known cases of hypersensitivity to the active substances or to any of the excipients.
Do not use on cats. (Refer to section 12. – Special warnings).

6. ADVERSE REACTIONS

Pruritus, hair loss, erythema, oedema, and erosions may occur in very rare cases at the application site and are generally self-resolving.

Behaviour changes (agitation, restlessness, whining or rolling), gastro-intestinal symptoms (vomiting, diarrhoea, hypersalivation, diminished appetite) and neurological signs such as unsteady movement and twitching, or lethargy in dogs susceptible to the ingredient permethrin may be observed in very rare cases. These signs are generally transient and self-resolving.

Accidental oral uptake may result in transient vomiting and neurological signs such as tremor and incoordination. Treatment should be symptomatic. There is no known specific antidote.

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 serious 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.

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

Route of administration and dosage:
Spot-on use only. Apply only to undamaged skin.

The recommended minimum dose is:
10 mg/kg body weight (bw) imidacloprid and 50 mg/kg body weight (bw) permethrin.

Administer by topical application to the skin according to the bodyweight as follows:

Dogs (kg body weight)	**Strength**	**Volume (ml)**	**Imidacloprid (mg/kg body weight)**	**Permethrin (mg/kg body weight)**
≤ 4 kg	200 mg/40 mg spot-on solution for dogs up to 4 kg	0.4 ml	minimum of 10	minimum of 50
>4 kg ≤ 10 kg	500 mg/100 mg spot-on solution for dogs over 4 kg up to 10 kg	1.0 ml	10 - 25	50 - 125
>10 kg ≤ 25 kg	1250 mg/250 mg spot-on solution for dogs over 10 kg up to 25 kg	2.5 ml	10 - 25	50 - 125
>25 kg ≤ 40 kg	2000 mg/400 mg spot-on solution for dogs over 25 kg	4.0 ml	10 - 16	50 - 80

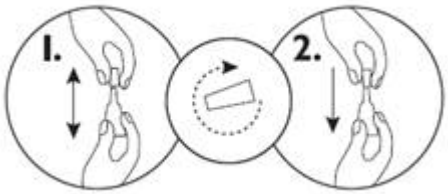
For dogs > 40 kg the appropriate combination of pipettes should be used.

To ensure correct dosage, body weight should be determined as accurately as possible.

Transient cosmetic changes (e.g. skin scaling, white deposits and spiking of the hair) may be observed at application sites.

Method of administration:

Remove one pipette from the package. Hold applicator pipette in an upright position. Tap the narrow part of pipette to ensure the contents are within the main body of the pipette, twist and pull cap off. Turn the cap around and place the other end of cap back on pipette. Push and twist the cap to break seal, and then remove the cap from the pipette.



For dogs 10 kg body weight or less:

With the dog standing still, part the coat between the shoulder blades until the skin is visible. Place the tip of the pipette on the skin and squeeze firmly several times to empty the contents directly onto the skin.

For dogs of more than 10 kg body weight:

With the dog standing still, the entire contents of the pipette should be applied evenly to four spots on the top of the back from the shoulder to the base of the tail. At each spot, part the hair until the skin is visible. Place the tip of the pipette on the skin and gently squeeze to expel a portion of the solution on the skin. Do not apply an excessive amount of solution at any one spot that could cause some of the solution to run off the side of the dog.

9. ADVICE ON CORRECT ADMINISTRATION

To reduce re-infestation from emergence of new fleas, it is recommended to treat all dogs in a household. Other animals living in the same household should also be treated with a suitable product. To aid further in reducing environmental challenge, the additional use of a suitable environmental treatment against adult fleas and their developmental stages is recommended.

Depending on the ectoparasite challenge, it may be necessary to repeat the treatment. The interval between two treatments should be 4 weeks. However, in cases of frequent and/or prolonged water exposure the persistent efficacy may be reduced. In these cases do not retreat more frequently than once weekly.

To protect a dog over the whole sand fly season, treatment should be compliantly continued throughout.

10. WITHDRAWAL PERIOD(S)

Not applicable.

11. SPECIAL STORAGE PRECAUTIONS

Keep out of the sight and reach of children.

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

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

12. SPECIAL WARNING(S)

Special warnings for each target species:
There may be an attachment of single ticks or bites by single sand flies or mosquitoes. For this reason, a transmission of infectious diseases cannot be excluded if conditions are unfavourable.
As the product exerts a repellent (anti-feeding) activity against *Aedes aegypti* mosquitoes 7 days after treatment, the product should preferably be applied 1 week before animals are likely to become exposed to these mosquitoes.
The product remains effective against fleas if the animal becomes wet. After weekly immersions in water for one minute the period of persistent insecticidal efficacy against fleas was not reduced. However, prolonged, intense exposure to water should be avoided. In cases of frequent and/or prolonged water exposure the persistent efficacy may be reduced. In these cases do not retreat more frequently than once weekly. If a dog requires a shampoo, it should be administered before applying the product or at least 2 weeks after application, to optimise efficacy of the product.
The effectiveness of the product against ticks, and the repellent effectiveness of the product following swimming or shampooing has not been investigated.
Immediate protection against sandflies bites is not documented. Treated dogs for the reduction of the risk of infection with *Leishmania infantum* via transmission by sandflies *P. perniciosus* should be kept in a protected environment during the first 24 hours after the initial treatment application.

Special precautions for use in animals:
Care should be taken to avoid the content of the pipette coming into contact with the eyes or mouth of the recipient dogs.

Care should be taken to administer the product correctly as described under section 8. In particular oral uptake due to the licking of the application site by treated or in-contact animals should be avoided.

Do not use on cats.

This product is extremely poisonous to cats and could be fatal due to the unique physiology of cats which is unable to metabolise certain compounds including permethrin. To prevent cats

from being accidentally exposed to the product, keep treated dogs away from cats after treatment until the application site is dry. It is important to ensure that cats do not groom the site of application on a dog, which has been treated with this product. Seek veterinary advice immediately if this occurs.

Consult your veterinary surgeon before using the product on sick and debilitated dogs.

The product is toxic for aquatic organisms. Treated dogs must not under any circumstances be allowed into any type of surface water for at least 48 hours after treatment.

Special precautions to be taken by the person administering the veterinary medicinal product to animals:
Avoid contact between the product and skin, eyes or mouth.
Do not eat, drink or smoke during application.
Wash hands thoroughly after use.
In case of accidental spillage onto skin, wash off immediately with soap and water.
People with known skin sensitivity may be particularly sensitive to this product.
The predominant clinical symptoms that in extremely rare case may be shown are transient sensory irritations of the skin like tingling, burning sensation or numbness.
If the product gets accidentally into the eyes, they should be thoroughly flushed with water. If skin or if eye irritation persists, seek medical advice immediately and show the package leaflet or the label to the physician.
Do not ingest. In case of accidental ingestion seek medical advice immediately and show the package leaflet or the label to the physician.
Treated dogs should not be handled especially by children until the application site is dry. This may be ensured by treating the dogs e.g. in the evening. Recently treated dogs should not be allowed to sleep together with their owner, especially children.
Keep the stored pipettes in the original blister. In order to prevent children from getting access to used pipettes, used pipettes should be disposed of immediately.
The solvent in the product may stain certain materials including leather, fabrics, plastics and finished surfaces. Allow the application site to dry before permitting contact with such materials.
Strength 2000 mg/400 mg spot-on solution for dogs over 25 kg: Laboratory studies in rabbits and rats with the excipient N-methyl pyrrolidone have shown evidence of foetotoxic effects. The veterinary medicinal product should not be administered by pregnant women and women suspected of being pregnant. Personal protective equipment consisting of gloves should be worn when handling the veterinary medicinal product by women of childbearing age.

Pregnancy and lactation:
The safety of the veterinary medicinal product has not been established in dogs during pregnancy, lactation or in animals intended for breeding. Laboratory studies in rabbits and rats with the excipient N-methyl pyrrolidone have shown evidence of foetotoxic effects. Use only according to the benefit-risk assessment by the responsible veterinarian.

Interaction with other medicinal products and other forms of interaction:
None known.

Overdose (symptoms, emergency procedures, antidotes):
No adverse clinical signs were noted in healthy puppies or adult dogs exposed to 5x overdosage or for puppies whose mothers were treated with 3x overdosage of the combination of imidacloprid and permethrin. The severity of skin erythema, which sometimes occurs at the application site, increases with overdose.

Incompatibilities:

None known.

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

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

14. DATE ON WHICH THE PACKAGE LEAFLET WAS LAST APPROVED

15. OTHER INFORMATION

White polypropylene pipette closed with a polyethylene (HDPE) cap. Each pipette is packed in a polyethylene terephthalate/aluminium/low density polyethylene triplex bag.
1 ml pipette containing 0.4 ml of solution
3 ml pipette containing 1.0 ml of solution
6 ml pipette containing 2.5 ml and 4.0 ml of solution
Box containing 1, 3, 4, 6, 10 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.