

Cyclavance™

(cyclosporine oral solution)
USP MODIFIED

HAPPINESS!

is **CYCLAVANCE™** (cyclosporine oral solution) USP MODIFIED

LIQUID CYCLOSPORINE

Give atopic dogs the cyclosporine they deserve with **CYCLAVANCE™** oral solution



Virbac

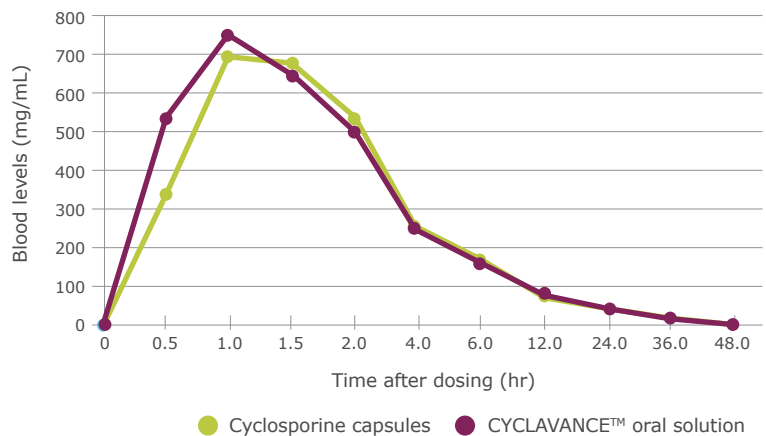
Shaping the future
of animal health

Happiness is reliability

The effective cyclosporine you know and trust—IN LIQUID FORM

- Cyclosporine is proven to significantly relieve pruritus and reduce skin lesions in just 4 weeks ($P < 0.001$)¹
- Cyclosporine maintains long-term beneficial effects with 1 dose every other day or twice weekly²
- Cyclosporine is recommended by the International Committee on Allergic Diseases of Animals (ICADA) for the treatment of atopic dermatitis in dogs³
 - Blocks the release of inflammatory cytokines
 - Exhibits strong anti-inflammatory and antipruritic effects

Mean cyclosporine plasma concentration (mg/mL) following administration at 5 mg/kg of CYCLAVANCE™ (cyclosporine oral solution) USP MODIFIED and cyclosporine capsules⁴



“A novel liquid oral formulation of ciclosporin* (CYCLAVANCE™, Virbac) was recently reported to be better accepted than ciclosporin capsules.”

– Guidelines from the ICADA³

Help your patients feel happy in their skin.



To order, contact your distributor or Virbac representative, or call 1-844-484-7222.

Visit <https://vet-us.virbac.com/cyclavance> to learn more.

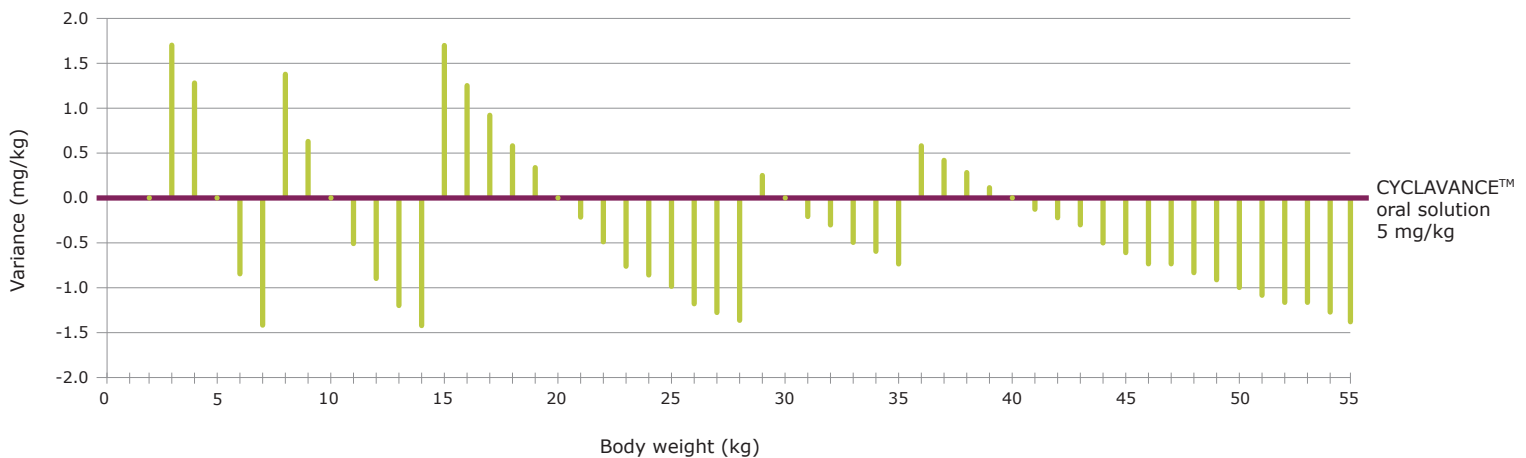
*CYCLAVANCE™ oral solution launched first in the EU where the active ingredient is spelled “ciclosporin.”

Happiness is precision in dosing

Eliminates the inefficiencies of dosing with capsules

- Liquid formulation allows for precise dosing at 5 mg/kg for all dogs
- Simple dosing makes it easy for the pet owner to follow instructions and minimizes the risk of over- and underdosing
- When compared with CYCLAVANCE™ oral solution at 5 mg/kg, variance in dosing for cyclosporine capsules can range from -29% to +33%^{2,5}

Cyclosporine capsules variance from 5 mg/kg ideal dose^{2,5†}



[†]Based on dosing recommendations in cyclosporine capsules package insert²

Important Safety Information

CYCLAVANCE™ (cyclosporine oral solution)(USP Modified): For use in dogs only. Wear gloves during and wash hands after administration. Gastrointestinal problems and gingival hyperplasia may occur at the initial recommended dose of CYCLAVANCE Oral Solution for Dogs. CYCLAVANCE Oral Solution for Dogs should be used with caution: 1) in cases with diabetes mellitus as it may cause elevated levels of serum glucose; 2) in dogs with renal insufficiency since the effect of cyclosporine use on dogs with compromised renal function has not been studied; 3) in simultaneous administration with drugs that suppress the P-450 enzyme system, such as azoles (e.g. ketoconazole), that may lead to increased plasma levels of cyclosporine. Killed vaccines are recommended for dogs receiving CYCLAVANCE Oral Solution for Dogs because the impact of cyclosporine on the immune response to modified live vaccines has not been evaluated. For full prescribing information, contact Virbac at 1-800-338-3659 or visit us.virbac.com.

Happiness is

confidence

in delivery

Convenient and easy dosing to help promote compliance

- In 2 studies involving 320 individual tests comparing administration of CYCLAVANCE™ oral solution with that of cyclosporine capsules (n=70):
 - **98.3%** of dogs consumed the entire dose of CYCLAVANCE™ oral solution when mixed with a small amount of dry food (vs 2.2% consumption for cyclosporine capsules)⁶
 - **99.3%** of dogs easily accepted CYCLAVANCE™ oral solution when administered directly in the mouth (vs 27.1% acceptance for cyclosporine capsules)⁶



- 2 vial presentations:
15 mL and 50 mL
- Syringe and adaptor cap for easy dosing with no leaks or spills



References: **1.** Steffan J, Parks C, Seewald W; North American Veterinary Dermatology Cyclosporine Study Group. Clinical trial evaluating the efficacy and safety of cyclosporine in dogs with atopic dermatitis. *J Am Vet Med Assoc.* 2005;226:1855–1863. **2.** CYCLAVANCE™ (cyclosporine oral solution) USP MODIFIED [product label]. **3.** Olivry T, DeBoer DJ, Favrot C, et al. Treatment of canine atopic dermatitis: 2015 updated guidelines from the International Committee on Allergic Diseases of Animals (ICADA). *BMC Vet Res.* 2015;11:210. **4.** Data on file. Virbac Corporation. **5.** ATOPICA™ (cyclosporine capsules). Greenfield, IN: Elanco US Inc.; 2020. **6.** Navarro C, Crastes N, Benizeau E, McGahie D. Voluntary acceptance and consumption of two oral cyclosporin formulations in dogs: two randomised, controlled studies. *Ir Vet J.* 2015;68:3.



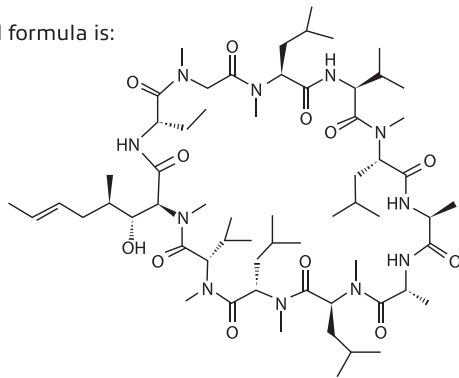
CYCLAVANCE™ (cyclosporine oral solution) USP MODIFIED 100 mg/mL

CAUTION: Federal (USA) Law restricts this drug to use by or on the order of a licensed veterinarian. Keep this and all drugs out of reach of children.

DESCRIPTION: CYCLAVANCE™ (cyclosporine oral solution) USP MODIFIED is an oral form of cyclosporine that immediately forms a microemulsion in an aqueous environment. Cyclosporine, the active ingredient in CYCLAVANCE, is a cyclic polypeptide, immune modulating agent consisting of 11 amino acids. It is produced as a metabolite by the fungal species *Beauveria nivea*.

Chemically, cyclosporine A is designated Cyclo[(E)-(2S,3R,4R)-3-hydroxy-4-methyl-2-(methylamino)-6-octenyl]-L-2-aminobutyryl-N-methylglycyl-N-methyl-L-leucyl-L-valyl-N-methyl-L-leucyl-L-alanyl-D-ananyl-N-methyl-L-leucyl-N-methyl-L-leucyl-N-methyl-L-valyl].

The structural formula is:



INDICATIONS: CYCLAVANCE is indicated for the control of atopic dermatitis in dogs weighing at least 4 lbs (1.8 kg) body weight.

DOSAGE AND ADMINISTRATION: Always Provide the Instructions for Assembling the Dispensing System and Preparing a Dose of CYCLAVANCE and the Information for Dog Owners with the prescription. The initial dose of CYCLAVANCE is 5 mg/kg/day as a single daily dose for 30 days. Following this initial daily treatment period, the dose of CYCLAVANCE may be tapered by decreasing the frequency of dosing to every other day or twice weekly, until a minimum frequency is reached which will maintain the desired therapeutic effect. CYCLAVANCE should be given at least one hour before or two hours after a meal. If a dose is missed, the next dose should be administered (without doubling) as soon as possible but dosing should be no more frequent than once daily. The dispensing system for the 5 and 15 mL vial sizes includes a 1 mL oral dosing syringe graduated in 0.05 mL increments. To dose the dog, administer 0.05 mL of CYCLAVANCE per 2.2 lbs of body weight. The dispensing system for the 30 and 50 mL vial sizes includes both a 1 mL oral dosing syringe graduated in 0.05 mL increments, and a 3 mL oral dosing syringe graduated in 0.1 mL increments. To dose the dog, administer 0.1 mL of CYCLAVANCE per 4.4 lbs of body weight. **Do not rinse or clean the oral dosing syringe between uses.** (See Instructions for Assembling the Dispensing System and Preparing a Dose of CYCLAVANCE.)

CONTRAINDICATIONS: CYCLAVANCE is contraindicated for use in dogs with a history of neoplasia. Do not use in dogs with a hypersensitivity to cyclosporine.

WARNINGS: CYCLAVANCE (cyclosporine oral solution) is a systemic immunosuppressant that may increase the susceptibility to infection and the development of neoplasia.

HUMAN WARNINGS: Not for human use. Keep this and all drugs out of reach of children. **For use only in dogs. Special precautions to be taken when administering CYCLAVANCE in dogs:** Do not eat, drink, smoke, or use smokeless tobacco while handling CYCLAVANCE. Wear gloves during administration. **Wash hands after administration.** In case of accidental ingestion, seek medical advice immediately and provide the package insert or the label to the physician.

People with known hypersensitivity to cyclosporine should avoid contact with CYCLAVANCE.

PRECAUTIONS: The safety and effectiveness of cyclosporine has not been established in dogs less than 6 months of age or less than 4 lbs body weight. CYCLAVANCE is not for use in breeding dogs, pregnant or lactating bitches. As with any immunomodulation regimen, exacerbation of sub-clinical neoplastic and infectious conditions may occur. Gastrointestinal problems and gingival hyperplasia may occur at the initial recommended dose (See Animal Safety).

CYCLAVANCE may cause elevated levels of serum glucose, and should be used with caution in cases with diabetes mellitus. If signs of diabetes mellitus develop following the use of CYCLAVANCE, consideration should be given to tapering or discontinuing the dose.

CYCLAVANCE should be used with caution with drugs that affect the P-450 enzyme system. Simultaneous administration of CYCLAVANCE with drugs that suppress the P-450 enzyme system, such as azoles (e.g. ketoconazole), may lead to increased plasma levels of cyclosporine.

Since the effect of cyclosporine use on dogs with compromised renal function has not been studied, CYCLAVANCE should be used with caution in dogs with renal insufficiency.

There have been reports of convulsions in human adult and pediatric patients receiving cyclosporine, particularly in combination with high dose methylprednisolone (See Animal Safety).

Killed vaccines are recommended for dogs receiving CYCLAVANCE because the impact of cyclosporine on the immune response to modified live vaccines is unknown (See Animal Safety).

ADVERSE REACTIONS: A total of 265 dogs were included in the field study safety analysis. One hundred and eleven (111) dogs were treated with placebo for the first 30 days. For the remainder of the study, all dogs received cyclosporine capsules.

Fourteen dogs withdrew from the study due to adverse reactions. Four dogs withdrew from the study after vomiting. One dog each withdrew from the study after diarrhea; vomiting, diarrhea and pruritus; vomiting, depression and lethargy; lethargy, anorexia and hepatitis; gingival hyperplasia, lethargy, polyuria/polydipsia and soft stool; seizure; sebaceous cyst; pruritus; erythema; or otitis externa.

Vomiting and diarrhea were the most common adverse reactions occurring during the study. In most cases, signs spontaneously resolved with continued dosing. In other cases, temporary dose modifications (brief

interruption in dosing, divided dosing, or administration with a small amount of food) were employed to resolve signs.

Persistent otitis externa, urinary tract infections, anorexia, gingival hyperplasia, lymphadenopathy and lethargy were the next most frequent adverse events observed. Gingival hyperplasia regressed with dose tapering. Owners of four dogs reported seizures while dogs were receiving cyclosporine. In one dog, seizures were the result of a brain tumor diagnosed one month into the study. Another dog experienced seizures before and after the study.

Otitis externa, allergic otitis, or pinna erythema, with or without exudates, commonly accompanies atopy. Many dogs entered the study with otitis externa, which did not resolve without otic treatment. New cases of otitis externa, allergic otitis, or pinna erythema developed while dogs were receiving cyclosporine. However, the incidence rate was lower with cyclosporine compared to placebo. A change in the dose frequency was not necessary when new cases occurred.

Number of Dogs Displaying Each Clinical Observation in the Field Study

Clinical sign	% out of 265
Vomiting	30.9%
Diarrhea	20.0%
Persistent Otitis Externa	6.8%
Urinary Tract Infection	3.8%
Anorexia	3.0%
Lethargy	2.3%
Gingival Hyperplasia	2.3%
Lymphadenopathy	2.3%

The following clinical signs were reported in less than 2% of dogs treated with cyclosporine in the field study: constipation, flatulence, Clostridial organisms in the feces, nausea, regurgitation, polyuria/polydipsia, strong urine odor, proteinuria, pruritus, erythema/flushed appearance, pyoderma, sebaceous adenitis, crusty dermatitis, excessive shedding, coarse coat, alopecia, papillomas, histiocytoma, granulomatous mass or lesion, cutaneous cyst, epulis, benign epithelial tumor, multiple hemangioma, raised nodule on pinna, seizure, shaking/trembling, hind limb twitch, panting, depression, irritability, hyperactivity, quieter, increased light sensitivity, reluctance to go outside, weight loss, hepatitis.

The following clinical signs were observed in 1.5-4.5% of dogs while receiving the placebo: vomiting, diarrhea and urinary tract infection. The following clinical signs were observed in less than 1% of dogs receiving the placebo: anorexia, otitis externa, cutaneous cysts, corneal opacity, lymphadenopathy, erythema/flushed appearance.

Clinical Pathology Changes: During the study, some dogs experienced changes in clinical chemistry parameters while receiving cyclosporine, as described in the following table:

Clinical Chemistry	% Affected (out of 265)
Elevated Creatinine	7.8%
Hyperglobulinemia	6.4%
Hyperphosphatemia	5.3%
Hyperproteinemia	3.4%
Hypercholesterolemia	2.6%
Hypoalbuminemia	2.3%
Hypocalcemia	2.3%
Elevated BUN	2.3%

In addition, the following changes in clinical chemistry parameters were noted in less than 2% of dogs: hypernatremia; hyperkalemia, elevated ALT, elevated ALP, hypercalcemia and hyperchloremia. These clinical pathology changes were generally not associated with clinical signs.

POST-APPROVAL EXPERIENCE: The following adverse events are based on post-approval adverse drug experience reporting. Not all adverse reactions are reported to FDA CVM. It is not always possible to reliably estimate the adverse event frequency or establish a causal relationship to product exposure using this data. The following adverse events are grouped by body system and are presented in decreasing order of reporting frequency.

Gastrointestinal: Emesis, diarrhea, gingival hyperplasia, hemorrhagic diarrhea, abdominal pain, hematemesis, digestive tract hemorrhage, hypersalivation, retching, flatulence, tenesmus, intestinal stasis, digestive tract hypermotility, melena, pancreatitis, involuntary defecation

General: Lethargy, anorexia, weight loss, polydipsia, hyperthermia, pale mucous membrane, general pain, collapse, dehydration, edema

Dermatologic: Pruritus, dermatitis and eczema, alopecia, erythema, papilloma, bacterial skin infection, skin lesion, skin and/or appendage neoplasm, pigmentation disorder, hair change, hyperkeratosis, histiocytoma, fungal skin infection, dermal cyst(s), desquamation

Behavioral: Hyperactivity, behavioral changes, anxiety, vocalization, aggression, inappropriate urination, disorientation

Neurologic: Muscle tremor, convulsion, ataxia, paresis

Respiratory: Tachypnea, dyspnea, cough

Urologic: Polyuria, urine abnormalities (hematuria, urinary tract infection, proteinuria, glucosuria, decreased urine concentration) urinary incontinence, cystitis, renal failure, renal insufficiency

Immune: Urticaria, anaphylaxis, allergic edema

Blood and lymphatic: Lymphadenopathy, anemia, hypoalbuminemia, leukopenia

Hepatic: Elevated Liver Enzymes, hepatopathy, hepatomegaly, hepatitis

Musculoskeletal: Lameness, limb weakness, myositis

Ear and labyrinth: Otitis externa

Cardio-vascular: Tachycardia

Endocrine: Diabetes mellitus, hyperglycemia

In some cases, death/euthanasia has been reported as an outcome of the adverse events listed above.

Neoplasms have been reported in dogs taking cyclosporine, including reports of lymphoma/lymphosarcoma and mast cell tumor. It is unknown if these were preexisting or developed de novo while on cyclosporine.

Diabetes mellitus has been reported; West Highland White Terriers are the most frequently reported breed.

To report suspected adverse drug events, for technical assistance or to obtain a copy of the Safety Data Sheet, contact Virbac AH, Inc. at 1-888-338-3659. For additional information about adverse drug experience reporting for animal drugs, contact FDA at 1-888-FDA-VETS or <http://www.fda.gov/reportanimalae>.

CLINICAL PHARMACOLOGY: Cyclosporine is an immunosuppressive agent that has been shown to work via suppression of T-helper and T-suppressor cells and inhibition of interleukin-2. It does not depress hematopoiesis or the function of phagocytic cells. A decrease in CD4 and CD8 cells was not seen in dogs receiving 20 mg/kg/day of cyclosporine for 56 days. Cyclosporine is not a corticosteroid or an antihistamine.

METABOLISM: Cyclosporine is extensively metabolized by the cytochrome P-450 enzyme system in the liver, and to a lesser degree in the gastrointestinal tract and the kidney. The metabolism of cyclosporine can be altered by the co-administration of a variety of agents (See *Precautions*).

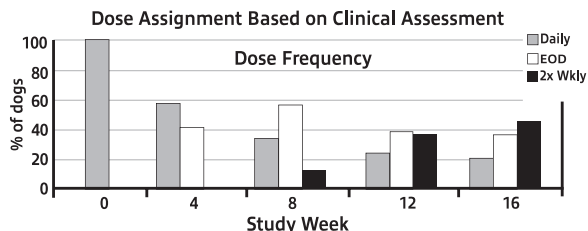
EFFECTIVENESS FIELD STUDY: A multisite, placebo controlled, double masked, field study was conducted in the United States and Canada using 16 investigators. Two hundred sixty five (265) dogs aged 1-10 years, weighing 4-121 lbs received either cyclosporine capsules at 5 mg/kg/day or placebo capsules. After 30 days, placebo dogs were switched to cyclosporine capsules.

Dogs were treated with cyclosporine capsules for a total of 4 months. No additional therapy with antihistamines, corticosteroids or medicated shampoos was permitted. Evaluations for pruritus and for skin lesions to derive a Canine Atopic Dermatitis Extent and Severity Index (CADESI) score occurred at enrollment and at monthly intervals. One hundred ninety-two (192) dogs were included in the statistical analysis of effectiveness.

At the end of the 30 day placebo controlled period, CADESI scores of dogs treated with cyclosporine capsules improved by 45% from enrollment, while CADESI scores of dogs treated with placebo worsened by 9%. Seventy-four percent (74%) of cyclosporine capsule treated dogs showed improvement in their pruritus scores over the first 30 day period, while only 24% of the placebo treated dogs showed an improvement. Owner and Veterinary Global Assessment in response to treatment also demonstrated statistically significant ($p < 0.0001$) improvement. After 4 weeks of therapy, Owner and Veterinary Global Assessments showed approximately twice as much improvement in the cyclosporine capsule treated dogs as compared to placebo treated dogs.

Improvements in pruritus accompanied by 50% or 75% improvements in CADESI scores resulted in dose reductions to every other day or twice weekly respectively. Not all dogs were able to decrease to twice weekly dosing. Some animals required upward or downward dosage adjustments during the study. Such adjustments should be expected during therapy of this disease. Dogs unable to decrease from once daily dosing after 60 days were considered dose reduction failures for the purposes of the study.

The results of dose assignments, based on the study criteria, for each 4-week dosing period, are shown in the graph below.



Analysis of blood levels of cyclosporine drawn during the study demonstrated no correlation between blood cyclosporine levels and CADESI scores or pruritus; therefore monitoring blood cyclosporine levels is not an appropriate predictor of effectiveness.

ANIMAL SAFETY: In a 52-week oral study with dose levels of 0, 1, 3, and 9 times the target initial daily dose, emesis, diarrhea and weight loss were seen in all cyclosporine treated groups with increasing frequency as the dose increased.

Multilocal papilloma-like lesions of the skin were observed in 5 out of 8 high dose animals between weeks 20 and 40. These changes regressed spontaneously after drug was withdrawn.

Other findings in the mid and high dose animals included swollen gums due to chronic gingivitis and periodontitis, lower serum albumin and higher cholesterol, triglyceride, IgA and IgG. Hematological findings consisted of anemia and decreased leukocyte counts in a few high dose animals. Erythrocyte sedimentation rates were increased at all dose levels in a dose dependent fashion. Notable histopathological findings were limited to lymphoid atrophy, hypertrophic gums (from gingivitis) and slight regenerative changes of the renal tubular epithelium in high dose animals. The findings were shown to be reversible during a 12-week recovery phase of the study.

In a 90-day study with cyclosporine, dogs were dosed in one of two patterns: either 1, 3, or 5X the maximum recommended target initial daily dose for 90 days, or 1, 3, or 5X the maximum recommended target initial daily dose for 30 days followed by tapering to mimic the recommended clinical dosing pattern. The maximum recommended dose, when administered for 90 days causes callus-like lesions on the footpads, red/swollen pinnae, mild to moderate gingival proliferation, hyperkeratotic areas on the integument, hair loss, salivation, vomiting, and diarrhea/abnormal stools. These clinical signs lessened in severity or resolved as the drug was tapered to a lower dose. Increased erythrocyte sedimentation rate, hyperproteinemia, hyperglobulinemia, hypoalbuminemia, hypocalcemia, hypophosphatemia, and hypomagnesemia were observed at three and five times the maximum recommended dose. These resolved as the dose was tapered.

When administered at higher than the maximum recommended dose, raised skin lesions, papilloma-like areas on the integument, popliteal lymph node enlargement, and weight loss were also seen. There were no cyclosporine related changes in urinalysis, ECG, blood pressure, or ophthalmologic exams.

Gross necropsy revealed epithelial changes consistent with those seen on physical examination. Proliferation of gingiva and toe pad epithelium was seen in all cyclosporine dosed groups, and was seen in a dose dependent fashion. The degree of the proliferation was greater in dogs in the non-tapered groups as compared to the tapered groups. Similar changes were noted on histopathologic examination of the cutaneous changes seen on physical examination. These lesions were characterized by epidermal hyperplasia, chronic dermatitis and hyperkeratosis.

Methylprednisolone combination: Twenty-four dogs were administered 1 mg/kg/day methylprednisolone alone for 14 days followed by 20 mg/kg/day cyclosporine either alone or in combination with methylprednisolone, or placebo for 14 days. There was no evidence of seizures/convulsions or neurological signs.

Vaccination effect: The effect of cyclosporine administration on the immunological response to vaccination was evaluated in a study in which 16 dogs were dosed with either cyclosporine at 20 mg/kg/day (4X the initial daily dose) or placebo for 56 days. All dogs were vaccinated on Day 27 with a killed commercial rabies virus and a multivalent vaccine (DHLPP) which included a modified live virus. Antibody titers for rabies, canine distemper, canine adenovirus type 2, parainfluenza, parvovirus, *Leptospira canicola*, and *Leptospira icterohaemorrhagiae* were examined on Days 0, 27 (prior to vaccination), 42 and 56. Quantification of CD4, CD8, and CD3 T-lymphocytes was analyzed.

Clinical changes included soft stool and dermatologic changes consistent with those seen in previous studies. Antibody titers did not rise in dogs treated with cyclosporine or the placebo for any component of the multivalent vaccine which included a modified live virus while all animals demonstrated a significant increase in antibody rabies titer by Day 42 or 15 days post-revaccination. No effect was seen on T-lymphocytes.

STORAGE INFORMATION: CYCLAVANCE (cyclosporine oral solution) USP MODIFIED should be stored and dispensed in the original container at temperatures between 68-86°F (20-30°C). Do not refrigerate because a precipitate may be observed below 68°F (20°C).

Once opened, use contents within 12 weeks.

HOW SUPPLIED: CYCLAVANCE is supplied in glass amber vials of 5, 15, 30 and 50 mL at 100 mg/mL.

- 5 and 15 mL vials are supplied with a 1 mL Luer Lok® oral dosing syringe.

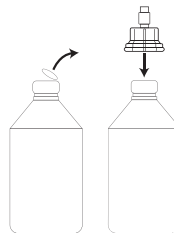
- 30 and 50 mL vials are supplied with a 1 mL and 3 mL Luer Lok® oral dosing syringes.

Instructions for Assembling the Dispensing System and Preparing a Dose of CYCLAVANCE™ (cyclosporine oral solution) USP MODIFIED.

Assembling the Dispensing System

The dispensing system consists of three parts:

1. A vial containing the medicine sealed with a rubber stopper
2. A plastic adapter (dispensing system) that you will push onto the top of the vial. The adapter must always remain on the vial after first use.
3. An oral dosing syringe that fits into the top of the plastic adapter to withdraw the prescribed dose of medicine from the vial. (1 mL syringe with the 5 and 15 mL vial sizes; 1 and 3 mL syringes with the 30 and 50 mL vial sizes)



Fitting the Plastic Adapter into the New Bottle of Medicine

1. Remove the plastic lid from the top of the vial.
2. Hold the vial upright on a table and push the plastic adapter firmly onto the top of the vial until it is firmly and evenly seated.

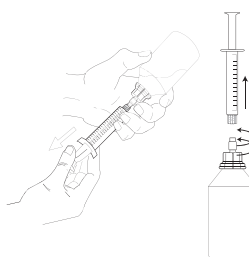
Note: To prepare a dose, carefully follow the instructions for **Preparing a Dose of Medicine.**

Preparing a Dose of Medicine

1. Check that the plunger of the oral dosing syringe is pushed all the way down.
2. Keep the vial upright and push the oral dosing syringe firmly into the plastic adapter while turning the syringe clockwise to secure the dispensing system.
3. Turn the vial with the attached dosing syringe upside down and slowly pull the plunger up so that the oral dosing syringe fills with the medicine.
4. **Expel any large bubbles by pushing and pulling the plunger a few times. The presence of a few tiny bubbles is not important for dosing accuracy.**
5. Withdraw the dose of medicine prescribed by your veterinarian.

Note: If the prescribed dose is more than the maximum volume marked on the oral dosing syringe, you will need to reload the syringe to withdraw the full dose.

6. Return the vial to its upright position and remove the oral dosing syringe by twisting it counterclockwise out of the plastic dispenser.



You can now introduce the syringe into the mouth of the dog according to your veterinarian's instructions, and push the medicine out of the syringe.

See **Information for Dog Owners** for complete administration instructions.

Do not rinse or clean the oral dosing syringe between uses.

Store the medication and the dosing syringe securely. CYCLAVANCE should be stored and dispensed in the original container at temperatures between 68-86°F (20-30°C). Do not refrigerate because a precipitate may be observed below 68°F (20°C). **Once opened, use contents within 12 weeks.**

Keep out of reach of children

Approved by FDA under ANADA # 200-692

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