PRN PHARMACAL

A MUST-READ, IF YOUR DOG HAS MISPLACED PEE.

PIPER THE SUPER-DOG

RIDDING THE WORLD OF INCONTINENCE ONE PUDDLE AT A TIME!

DR. PIPER HAS BEEN IMMERSED IN PEE PROBLEM SOLVING, AND HER PATIENTS ARE THRILLED TO HAVE THEIR LEAKING UNDER CONTROL! WHAT'S NEXT? DON'T MISS THIS EXCITING NEW CHAPTER OF *THE ADVENTURES OF DR. PIPER!*


THANKS!
THANK YOU, DR. PIPER – YOU'RE MY HERO!
THANK YOU!
Z Z Z z
DR. PIPER HAS BEEN BUSY HELPING DOGS FROM ALL PARTS OF THE COUNTRY WITH MISPLACED PEE.
CANINE URINARY INCONTINENCE CAN AFFECT DOGS OF ANY AGE, BREED, OR SIZE. THIS PROBLEM IS MOST COMMON IN LARGE BREED SPAYED FEMALES.[1]


SUDDENLY, DR. PIPER IS STARTLED AWAKE BY A CRY FOR HELP!
OOOHHHH NOOOOOOO! I PEEED IN THE HOWWWWSE!
Dr. Piper
PRN PHARMACAL
SOUNDS LIKE MISSY IS IN TROUBLE!
I'M GOING TO BE IN BIG TROUBLE. PLEEEASE HEELLP!
WITH ONE TOUCH OF HER MAGIC STETHOSCOPE, THE TRANSFORMATION BEGINS...
THIS IS A JOB FOR PIPER THE SUPER-DOG!


DON'T WORRY, MISSY. I'M A SUPER DOG WITH SUPER HELP FOR YOU!
I ONCE HAD A PEE PROBLEM MYSELF. I LEARNED THAT WHEN HOUSE-TRAINED DOGS HAVE URINARY ACCIDENTS, THE CAUSE CAN BE A COMMON MEDICAL CONDITION CALLED URETHRAL SPHINCTER HYPOTONUS (USH). IN FACT, ONE OUT OF FIVE OF US HAVE IT!
WHEN I WENT TO VETERINARY SCHOOL, I LEARNED ALL ABOUT IT.
PRN
PHARMACAL
URETHRAL SPHINCTER HYPOTONUS SIGNS:
DRIPPING, LEAKING URINE
WET SPOTS WHERE DOG SLEEPS
EXCESSIVE LICKING OF GENITALS
RED, IRRITATED SKIN
INAPPROPRIATE URINATION INDOORS

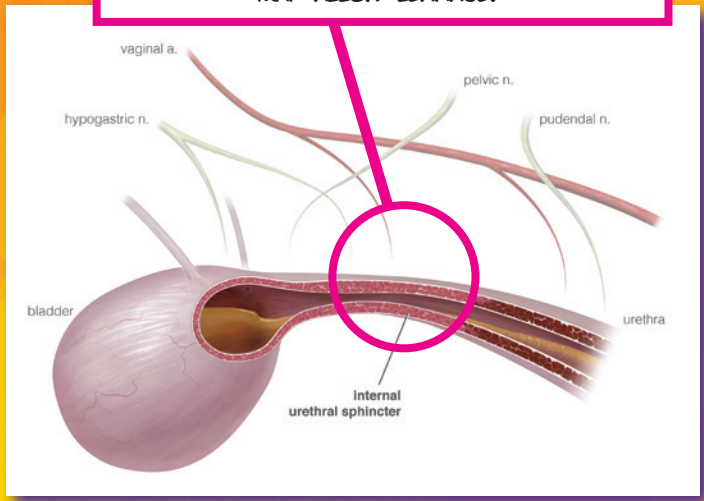
When our urethral muscles are weak from USH, urine can leak whenever we relax!

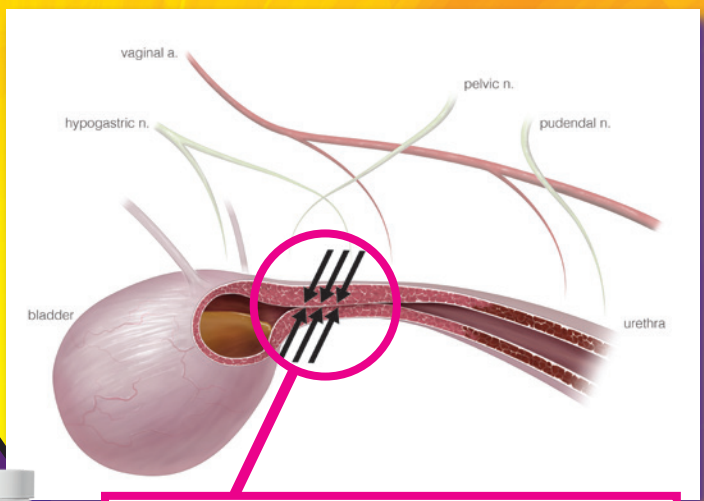
But PROIN ER™ (phenylpropanolamine hydrochloride extended-release tablets) can help. Once I started it my accidents stopped, and I began spreading the word about PROIN ER!

Up to one in five spayed dogs is affected by urinary incontinence due to urethral sphincter hypotonus (Urinary sphincter mechanism incompetence or USMI).[2]

WITHOUT PROIN ER

When the urethral muscle is weak, the tube from the bladder to the outside may allow leakage.

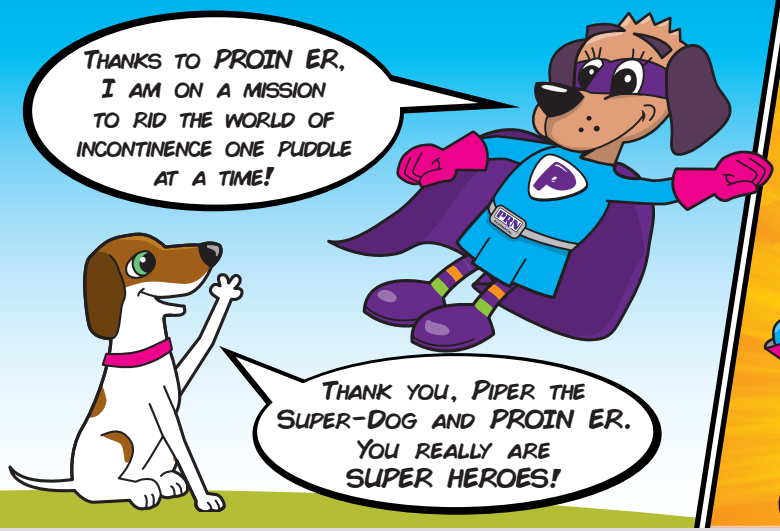


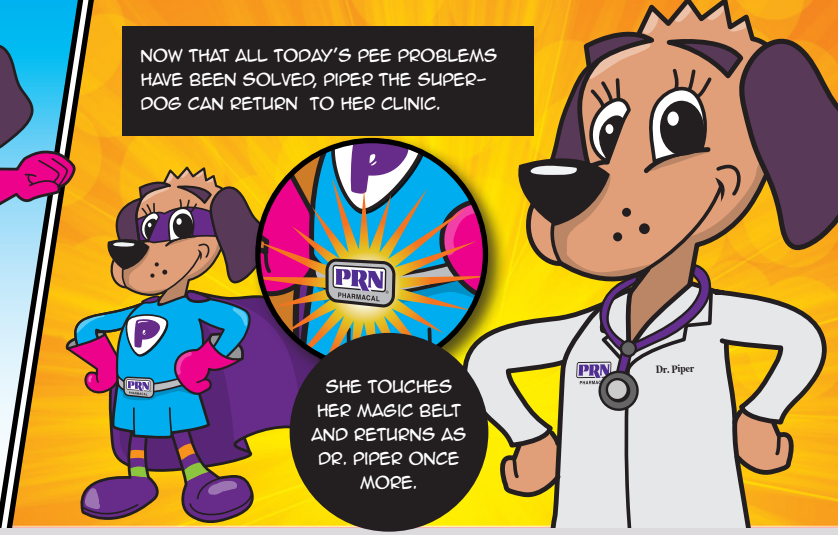


WITH PROIN ER

PROIN ER can strengthen urethral sphincter muscle tone to safely and effectively control urinary incontinence.

For Important Safety Information, see next page.





FOR MORE THAN A DECADE, VETERINARIANS HAVE TRUSTED THE PROIN BRAND TO HELP STRENGTHEN THE URETHRAL SPHINCTER MUSCLE AND CONTROL LEAKING.

PROIN ER™ (PHENYLPROPANOLAMINE HYDROCHLORIDE EXTENDED-RELEASE TABLETS) IS FDA APPROVED FOR CONTROL OF URINARY INCONTINENCE DUE TO URETHRAL SPHINCTER HYPOTONUS. PROIN ER PATENTED EXTENDED-RELEASE TABLETS PROVIDE THE CONVENIENCE OF ONCE-A-DAY DOSING.

PROIN ER IS AVAILABLE ONLY FROM VETERINARIANS.

JOIN PIPER THE SUPER-DOG IN HER MISSION TO RID THE WORLD OF INCONTINENCE ONE PUDDLE AT A TIME!

"P IS FOR PROIN, ER IS FOR EXTENDED-RELEASE"

ONCE-A-DAY PROIN ER TABLETS

IMPORTANT SAFETY INFORMATION: For oral use in dogs only. Not for human use. Keep out of reach of children. If accidentally ingested by humans, contact a physician immediately.

The most commonly reported side effects were vomiting, loss of appetite, diarrhea, excessive salivation, agitation, tiredness, vocalization, confusion, increased water consumption, weight loss, weakness, fever, panting, and reversible changes in skin color (flushing or bright pink). Abnormal gait, seizures or tremors, as well as liver enzyme elevations, kidney failure, blood in urine and urine retention have been reported. In some cases death, including euthanasia has been reported. Sudden death was sometimes preceded by vocalization or collapse.

Instances of dogs chewing through closed vials of PROIN® and eating the vial contents have been reported, in some cases resulting in overdose. Keep the product in a secured storage area out of the reach of pets in order to prevent accidental ingestion or overdose, as dogs may willingly consume more than the recommended dosage of PROIN® Chewable Tablets or PROIN ER™ tablets. Contact your veterinarian immediately if the dog ingests more tablets than prescribed or if other pets ingest PROIN Chewable tablets or PROIN ER tablets.

PROIN and PROIN ER may cause elevated blood pressure and should be used with caution in dogs with pre-existing heart disease, high blood pressure, liver disease, kidney insufficiency, diabetes, glaucoma, and other conditions associated with high blood pressure. Dogs may transition from PROIN Chewable Tablets to PROIN ER without a break in administration. However, do not alternate PROIN ER with PROIN Chewable Tablets because the effectiveness and safety of interchangeable use have not been evaluated.

The safe use of PROIN and PROIN ER in dogs used for breeding purposes, during pregnancy or in lactating bitches, has not been evaluated. Contact your veterinarian if you notice restlessness or irritability, loss of appetite, the incontinence persists or worsens, or any other unusual signs. See prescribing information for complete details regarding adverse events, warning and precautions or visit prnpharmacal.com.

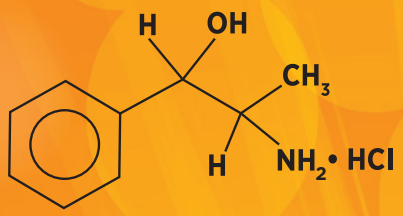
PROIN ER™

(phenylpropanolamine hydrochloride extended-release tablets)

For oral use in dogs only

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

DESCRIPTION: PROIN ER (phenylpropanolamine hydrochloride extended-release tablets) is a sympathomimetic amine closely related to ephedrine. Phenylpropanolamine hydrochloride (PPA) is the nonproprietary designation for benzenemethanol, α-(1-aminoethyl)-hydrochloride, (R*, S*)-, (±). The empirical formula is $C_9H_{13}NO \bullet HCl$ and the molecular weight is 187.67. It is a white crystalline compound having a slight aromatic odor. PPA is freely soluble in water and alcohol but is practically insoluble in ether, benzene and chloroform. The chemical structure of phenylpropanolamine hydrochloride is:



INDICATION: For the control of urinary incontinence due to urethral sphincter hypotonus in dogs.

DOSAGE AND ADMINISTRATION: The recommended dosage is 2 to 4 mg/kg (0.9 to 1.8 mg/lb) of body weight once daily according to Table 1 below. Administer PROIN ER with food (see **Clinical Pharmacology**). **Do not split or crush tablets**.

Dogs weighing less than 10 pounds cannot be safely dosed because tablet administration would result in a dose over 4 mg/kg.

Table 1. Dose Administration[a]

Body weight in pounds	PROIN ER
10-20	18 mg
21-40	38 mg
41-80	74 mg
81-125[b]	145 mg

[a]Body weight should be rounded to the nearest pound

[b]Dogs exceeding 125 lbs should receive the appropriate combination of tablets to achieve the recommended dosage.

Dogs may transition from PROIN® Chewable Tablets to PROIN ER without a break in administration. However, do not alternate PROIN ER with PROIN Chewable Tablets because the effectiveness and safety of interchangeable use have not been evaluated.

WARNINGS: Not for human use. Keep out of reach of children. Consult a physician in case of accidental ingestion by humans.

Keep PROIN ER in a secured location out of reach of dogs, cats, and other animals to prevent accidental ingestion or overdose.

PRECAUTIONS: Proin ER may mask signs of incontinence due to urinary tract infection. PROIN ER is not effective in dogs with incontinence due to neurologic disease or malformations.

PROIN ER may cause hypertension; therefore, use with caution in dogs with pre-existing heart disease, hypertension, liver disease, kidney insufficiency, diabetes, glaucoma, and conditions with a predilection for hypertension.

Use with caution in dogs receiving sympathomimetic drugs, tricyclic antidepressants, or monoamine oxidase inhibitors as increased toxicity may result. Use with caution in dogs administered halogenated gaseous anesthetics as this may increase the risk of cardiac arrhythmias.

A laboratory study on human blood revealed that phenylpropanolamine (PPA) used in conjunction with aspirin may potentiate decreased platelet aggregation.[1]

PROIN ER may cause increased thirst; therefore, provide dogs with ample fresh water.

The safe use of PROIN ER has not been evaluated in dogs that are intended for breeding, or that are pregnant or lactating.

ADVERSE REACTIONS: Adverse Reactions are listed below for both PROIN ER (NADA Number 141-517) and PROIN Chewable Tablets (NADA 141-324).

PROIN ER (NADA 141-517)

In the open-label clinical study involving 119 dogs administered PROIN ER once a day for 180 days, the following adverse reactions were observed.

Table 2. Number and percentage of dogs with adverse reactions in the 180-day open-label clinical study for PROIN ER

Adverse Reactions	Total N=119
Emesis	39 (32.8%)
Body weight loss (≥5%)	34 (28.6%)
Hypertension (≥160 mmHg) developed during study[a]	15 (12.6%)
Diarrhea	20 (16.8%)
Proteinuria	16 (13.4%)
Tachycardia (≥160 bpm)	11 (9.2%)
Lethargy	11 (9.2%)
Decreased appetite	10 (8.4%)
Urinary Tract Infection	10 (8.4%)
Elevated Alkaline phosphatase and/or Alanine Aminotransferase	7 (6.0%)
Hypoglycemia	4 (3.3%)
Hypercalcemia	3 (2.5%)
Increased BUN	2 (1.7%)
Bradycardia (<60 bpm)	2 (1.7%)
Seizures/twitching	2 (1.7%)

[a]There were an additional 21 dogs enrolled with hypertension who remained hypertensive throughout the study.

During the first week of administration of PROIN ER, 15% of dogs had reported emesis, diarrhea, or decreased appetite which improved or resolved prior to the Day 21 visit.

Four deaths occurred during the study. One dog was euthanized for pulmonary metastasis and one dog for poor quality of life due to hindlimb weakness. One dog had emesis and died at home; upon necropsy a foreign body was present in the small intestine. The fourth dog had been treated for a urinary tract infection three weeks prior to sudden death of undetermined cause.

PROIN Chewable Tablets (NADA 141-324):

Table 3 below includes the most common adverse reactions observed in the masked, placebo-controlled 28-day clinical study involving 123 PROIN Chewable Tablet-treated dogs and 61 placebo-treated dogs. In addition, one dog exhibited disorientation, nervousness, a 7.7% loss of body weight, and hypertension with proteinuria. A second dog exhibited restless behavior, lethargy, a 2.8% body weight loss, and proteinuria.

Table 3. Number and percentage of dogs with adverse reactions in the 28-day placebo-controlled clinical study for PROIN Chewable Tablets

Adverse Reactions	PROIN-treated (N=123)	Placebo-treated (N=61)
Emesis	20.3%	8.2%
Hypertension (≥160 mmHg)[a]	19.5%	14.7%
Anorexia	16.3%	3.3%
Body weight loss (>5%)[b]	16.1%	6.8%
Proteinuria	13.0%	8.2%
Anxiety/aggression/behavior change	9.7%	3.2%
Diarrhea	7.3%	9.8%
Polydipsia	6.5%	9.8%
Lethargy	5.7%	1.6%
Musculoskeletal Disorder	3.2%	1.6%
Insomnia/sleep disorder	2.5%	0.0%

[a]One or more systolic blood pressure readings of ≥160 mmHg.

[b]The "N" for weight loss is PROIN-treated N=118 and placebo N=59 because seven dogs did not have a final weight at the time of withdrawal from the study.

One-hundred fifty-seven dogs continued into the 6-month open-label clinical study for PROIN Chewable Tablets. The most common adverse reactions are listed in Table 4 below. In addition, one dog exhibited progressively worsening hypertension with proteinuria. Five dogs enrolled in the study with pre-existing heart disease. Of these, one dog developed systolic failure with an unknown relation to treatment.

Table 4. Number and percentage of dogs with adverse reactions in the 6-month open-label clinical study for PROIN Chewable Tablets

Adverse Reactions	Total N=125
Hypertension (≥160 mmHg)[a]	34.6%
Body weight loss (>5%)	24.8%
Emesis	19.7%
Proteinuria	15.3%
Anorexia	10.2%
Diarrhea	6.4%
Lethargy	5.7%
Anxiety/aggression/behavior change	5.7%

[a] Percent of dogs with systolic blood pressures of ≥160 mmHg on day -7 were 30.2% and on day 0 were 33.3%.

Post Approval Experience for PROIN Chewable Tablets (2015):

The following adverse reactions are based on voluntary, post approval reporting for PROIN Chewable Tablets (2015). Not all adverse events 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 these data. The signs reported are listed in decreasing order of reporting frequency by body system:

Gastrointestinal: Emesis, anorexia, diarrhea, hypersalivation

Behavioral: Agitation, lethargy, vocalization, confusion

General body system: Polydipsia, weight loss, weakness, fever

Respiratory: Panting

Dermatological: Erythema, piloerection

Hepatic: Elevated serum alanine aminotransferase (ALT), elevated serum alkaline phosphatase (ALP)

Neurologic: Ataxia, seizures, tremors

Renal/Urinary: Renal failure, hematuria, urinary retention

Cardiovascular: Tachycardia, hypertension, bradycardia, arrhythmias

Sensory: Ophthalmic disorders, mydriasis and eye redness

In some cases, death, including euthanasia, has been reported. Sudden death was sometimes preceded by neurologic signs, vocalization, or collapse. A necropsy of one dog revealed subarachnoidal and intraventricular hemorrhage in the brain.

The following signs have been reported more often with a dose higher than the recommended dosage: agitation, arrhythmia, bradycardia, erythema, fever, hypersalivation, hypertension, lethargy, mydriasis, panting, piloerection, tachycardia, tremor, and urinary retention.

Contact Information For a copy of the Safety Data Sheet (SDS) or to report suspected adverse drug events, contact Pegasus Laboratories at 1-800-874-9764. For additional information about adverse drug experience reporting for animal drugs, contact FDA at 1-888-FDA-VETS or http://www.fda.gov/AnimalVeterinarySafetyHealth.

INFORMATION FOR DOG OWNERS:

Always follow the dosage instructions for PROIN ER provided by your veterinarian. Give PROIN ER with food and do not split or crush the tablet. Monitor your dog after giving PROIN ER to be sure all of it was consumed. If you have difficulty giving PROIN ER, contact your veterinarian.

PROIN ER may cause increased thirst; therefore, provide dogs with ample fresh water.

If you forget to give your dog a dose, then resume dosing at the next scheduled dose. Keep PROIN ER in a secured location out of reach of dogs, cats, and other animals to prevent accidental ingestion or overdose.

Contact your veterinarian immediately if the dog ingests more tablets than prescribed or if other pets ingest PROIN ER. In the case of accidental ingestion by humans, contact a physician immediately.

Contact your veterinarian if you notice restlessness, irritability, loss of appetite, the incontinence persists or worsens, or any other unusual signs.

Consult your veterinarian before administering PROIN ER with any other medications.

CLINICAL PHARMACOLOGY:

Phenylpropanolamine is a chemical analogue of the endogenous sympathomimetic amines. It is an alpha-adrenergic agent which has been reported to increase urethral tone in dogs.[2] Its mechanism of action is not well determined, but it is believed to cause the release of norepinephrine by indirectly stimulating both the alpha and beta-adrenergic receptors of the smooth muscle to increase smooth muscle tone of the urethra, bladder neck, and the internal urethral sphincter.[3,4]

In a crossover pharmacokinetic study of PROIN ER in fed and fasted dogs, post-prandial drug administration was associated with approximately a 23% increase in the maximum plasma concentration (C_{max}), but the area under the concentration vs time curve to the last quantifiable concentration (AUC_{last}) was similar in both fed and fasted states. The small decrease in the post-prandial AUC_{last} appeared to be attributable to the corresponding increase in the terminal elimination rate constant under the fed conditions. The time to C_{max} (T_{max}) was more variable in the fasted state, ranging from 1.5 to 8 hours compared to 2 to 6 hours for the fed state. The elimination half-life ($t_{½}$) was also more variable in the fasted state, ranging from 3.89 to 10.35 hours compared to 2.98 to 7.81 hours for the fed state.

EFFECTIVENESS:

Effectiveness of PROIN ER was demonstrated in a multi-center, prospective, open-label, 6-month study in client-owned dogs of various breeds. In this study, 119 dogs (113 spayed females and 6 neutered males, aged 1-16 years and weighing 4.9-81.8 kg) who were considered well controlled for signs of urinary incontinence (UI) while receiving PROIN Chewable Tablets for at least 30 days prior to study start were enrolled in the study. Of these dogs, 104 were evaluated for effectiveness. The owners continued to administer PROIN Chewable Tablets twice a day and recorded episodes of UI during a baseline period (Day -7 through Day -1). After the baseline period, the owners transitioned to administration of PROIN ER once a day, at the labeled dose (see **Dosage and Administration**), and recorded urinary accidents for 28 days.

The primary variable was the ratio of average daily incidence of UI during the 7 days preceding the Day 28 clinic visit compared to the baseline period. It was concluded that PROIN ER was effective for the control of urinary incontinence due to urethral sphincter hypotonus in dogs.

Table 5: Clinical Effectiveness Results for PROIN ER

Ratio	Number of Dogs N=104
Ratio >1, indicating response measurement period was better than baseline period	19 (18.3%)
Ratio of 1, indicating no difference between response measurement period and baseline period	75 (72.1%)
Ratio <1, indicating response measurement period was worse than baseline period	10 (9.6%)

The secondary outcome variable was owner assessment of the control of UI at the end of the 28 day study period. The owner assessment was "improved" for 13 (12.5%) dogs, "stayed the same" for 90 (86.5%) dogs and "worsened" for 1 dog (1%).

ANIMAL SAFETY:

The safety of PROIN ER was established based on the safety data from PROIN Chewable Tablets (see below) and a comparative analysis of pharmacokinetic (PK) data for PROIN ER and PROIN Chewable Tablets. The statistical analysis of observed and simulated post-prandial pharmacokinetic data resulted in confidence limits consistent with equal or lower oral bioavailability for PROIN ER when administered once daily versus PROIN Chewable Tablets when administered twice daily. Therefore, the safety data from PROIN Chewable Tablets could be applied to PROIN ER. Emesis and hyperemia of the ventral abdomen were observed during the PK studies.

Target Animal Safety Study (PROIN Chewable Tablets, NADA 141-324)

In a target animal safety study, PROIN Chewable Tablets were administered to 32 healthy male and female Beagle dogs at 0, 2, 6 and 10 mg/kg of body weight (0, 1, 3 and 5 times the recommended dose; 8 dogs per group) twice daily for 26 consecutive weeks. The most pronounced finding was a dose-dependent increase in blood pressure. Mean systolic blood pressure was increased in all PPA-treated groups compared to the control, but mean values for all 4 groups were within the normal range. Mean diastolic and mean MAP (mean arterial pressure) were higher in the 3X and 5X groups, and in the 1X males. Dogs in the 3X and 5X groups had more individual systolic, diastolic, and MAP values above the normal range than the control group dogs. A dose-dependent decrease in heart rate was observed in the 3X and 5X dogs. In the 0, 1, 3, and 5X groups, 5%, 34%, 44%, and 40% of the total number of heart rates obtained from electrocardiograms for each group over the course of the study were below the normal range (70-120 beats per minute), with the lowest value being 51 bpm in 4 of the 1X group dogs. One dog in each of the 1X and 5X groups had an elevated heart rate between 150-180 beats per minute on at least 2 of the 13 physical exams. One dog in each of the 1X and 3X groups developed gallop heart sounds after treatment began that were noted in 12 of 13 and 6 of 13 physical exams respectively. Dogs in the PPA-treated groups exhibited anxious/restless behavior more frequently than the control group. One dog each in the 1X and 3X groups were responsible for the majority of the observations. A decline in mean body weight and body condition was observed in females in all 4 groups, including the control. One female in the 1X group lost 33% body weight. Emesis and loose stool occurred in a dose-related fashion, and most of the emesis episodes took place within 1 hour of dosing. Mean platelet counts were higher in at least one of the PPA-treated groups, with individual values up to 1.4X the upper limit of normal (ULN) in the 3X and 5X groups. The 3X and 5X groups had higher mean serum ALT values compared to the control. Mean ALT was within the normal range for all 4 groups. There were more dogs with ALT levels above the normal range in the 3 PPA-treated groups compared to the control, but increased values were transient and less than 1.8X ULN. All dogs had ALT values in the normal range at the conclusion of the study.

Tolerance study (PROIN Chewable Tablets, NADA 141-324)

In the separate tolerance study, 6 healthy female Beagle dogs were administered PROIN Chewable Tablets at 20 mg/kg body weight (10 times the recommended dose) twice daily for 21 consecutive days. Mean systolic blood pressure was increased in the 10X group compared to the control, but mean values were within the normal range for both groups. Mean diastolic pressures were above the normal range on days 7 and 21 for the 10X group, and day 14 for the control. The 10X dogs had hypertensive mean MAP values on days 7 and 21, whereas the control dog mean MAP values were in the normal range. There was a trend in 10X dogs for lower heart rates following initiation of PPA treatment. Four of 6 dogs in the 10X group had heart rates below the normal range on day 7, whereas none of the control dogs did. The 10X group dogs had increased hematocrit, hemoglobin, RBC counts, urine specific gravity, and water intake consistent with transient, sub-clinical dehydration that occurred shortly after PPA treatment was started. All 6 dogs in the 10X group developed emesis during the treatment period, whereas only 1 of the control dogs did. Most of the emesis episodes took place within 1 hour of dosing. Mean platelet counts were also higher in 10X dogs on all 3 exam days; mean values were above the normal range on day 7, with individual values up to 1.5X ULN. The 10X group had a higher mean serum ALT value on day 7 than the control. Mean ALT values for both groups were in the normal range on all 3 exam days, but 2 dogs in the 10X group had ALT values up to 1.4X ULN on day 7; these elevated values were transient, and all dogs had normal ALT values on days 14 and 21.

For either study, there was no evidence of chronic hypertension-induced target organ damage; there were no clinical findings attributable to PPA on the ophthalmic exams, electrocardiogram evaluation, or gross necropsy and histopathology.

STORAGE: Store at controlled room temperature 20-25°C (68-77°F), excursions permitted between 15-40°C (59-104°F).

HOW SUPPLIED: PROIN ER tablets contain 18, 38, 74 or 145 mg phenylpropanolamine hydrochloride per tablet. PROIN ER is packaged in bottles containing 30 or 90 tablets.

REFERENCES:

[1] Watson R, et al. Ephedra alkaloids inhibit platelet aggregation. Blood coagulation and Fibrinolysis, 2010, 21:266-271.

[2] Richter K.P., Ling G.V. Clinical response and urethral pressure profile changes after phenylpropanolamine in dogs with primary sphincter incompetence. JAVMA, Vol. 187, No 6, September 15, 1985. 605-611.

[3] Scott, L., Leddy M. and Bernay, F. Evaluation of phenylpropanolamine in the treatment of urethral sphincter mechanism incompetence in the bitch. J. Small Anim. Pract. 2002;43(11): 493-6.

[4] Noel, S., et al. Combined pharmacokinetic and urodynamic study of the effects of oral administration of phenylpropanolamine in female Beagle dogs. Vet. Journal, 2010; 184(2): 201-207.

Approved by FDA under NADA #141-517

PROIN ER is a trademark of Pegasus Laboratories, Inc.

01/2019

Manufactured By:
Pegasus Laboratories, Inc.
Employee-Owned
Pensacola, Florida 32514, USA
Manufactured in the USA

LIVER-FLAVORED TABLETS ARE AVAILABLE IN MULTIPLE CONVENIENT SIZES FOR ACCURATE DOSING.

PROIN ER™ *(phenylpropanolamine hydrochloride extended-release tablets)* TABLETS

- ☐ PROIN ER 18 MG TABLETS
- ☐ PROIN ER 38 MG TABLETS
- ☐ PROIN ER 74 MG TABLETS
- ☐ PROIN ER 145 MG TABLETS

GIVE YOUR DOG **ONE** TABLET PER DAY

REMEMBER:

"P IS FOR PROIN, ER IS FOR EXTENDED-RELEASE"

Clinic Contact Information

For more information, visit
prnpharmacal.com/proin, **ProinForCanines.com**,
or **Cuhi.org**

INFORMATION FOR DOG OWNERS Always follow the dosage instructions for PROIN ER™ provided by your veterinarian. Give PROIN ER with food and do not split or crush the tablet. Monitor your dog after giving PROIN ER to be sure all of it was consumed. If you have difficulty giving PROIN ER, contact your veterinarian.

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Consult your veterinarian before administering PROIN ER with any other medications.

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1. Cahn CM, Line @. *The Merck Veterinary Manual. 9th ed.* Whitehouse Station, NJ 2005.

2. Michael DeTora, DVM, and Robert J. McCarthy, DVM, MS, DACVS. "Ovariohysterectomy versus ovariectomy for elective sterilization of female dogs and cats: is removal of the uterus necessary?" *JAVMA*, Vol. 239, No 11. December 1, 2011.

PROINER-0219-38