Agmaset® and AgmaVet® Science Leading to Product



Agmaceuticals®:

Nutraceuticals containing the neuroprotective ingredient **G-Agmatine**[®].

G-Agmatine® The best brand of
Premium grade agmatine.

https://ForNerveHealth.com

Neuroscientists - 1977

<u>Donated to</u> – Dr. Donald J. Reis's

Laboratory of Neurobiology, Cornell

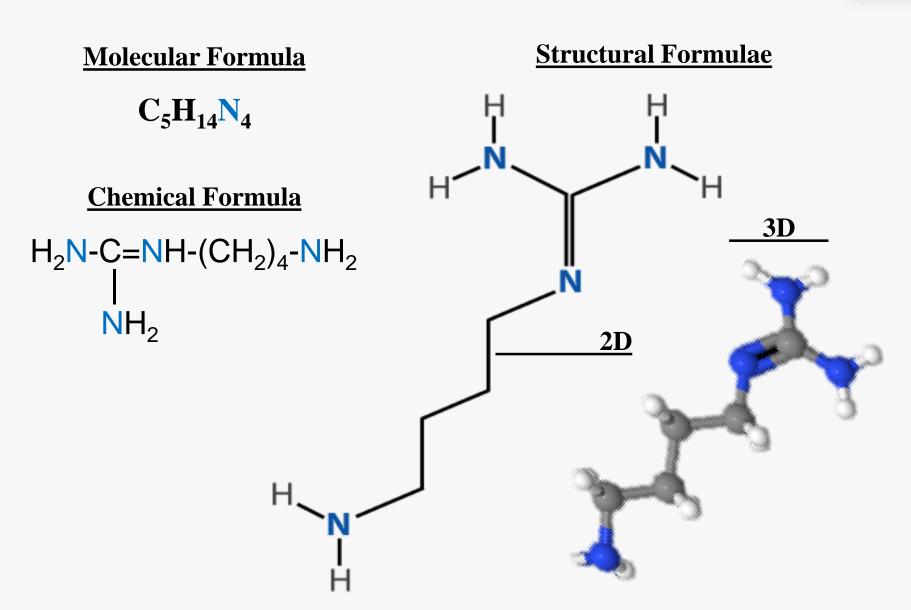
University Medical College.

(oil pastel on paper)



Agmatine

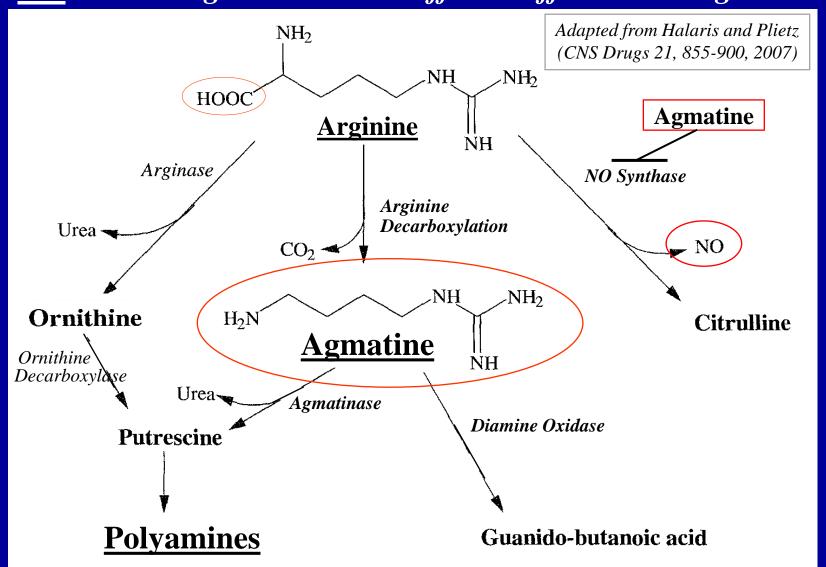




Metabolic Pathways



-Agmatine is a metabolite of the amino acid arginineBut the base agmatine exerts different effects than arginine





Agmatine is a Naturally Occurring Molecule with a Broad Distribution in the Body

• Brain: 300 ng/mg

• Blood: 47 ng/ml

• Intestine: 55 ng/mg

• Fecal: 3,290 ng/g



Agmatine is a Ubiquitous Compound Present in Many Plant- and Animal-Derived Foodstuff

- Commonly found in foodstuff derived from plants, fish and animals (usual range: 0.1 10 mg/kg).
- High levels are found in certain fish products (200 and up to 650 mg/kg).
- Intestinal microbial production is considered a major source of agmatine.

Suggested Utility of Agmatine in Modulating a Spectrum of Bodily Functions -Based on Animal Model Experiments-

Cilcole Cilcole
Fight For Nerve Health

- Nerve functions.
- Mood functions.
- Reactions to stress.
- Learning and memory capabilities.
- Glucose metabolism.
- Cardiovascular functions.
- Kidney functions.
- Cellular growth.
- Mitochondria functions.

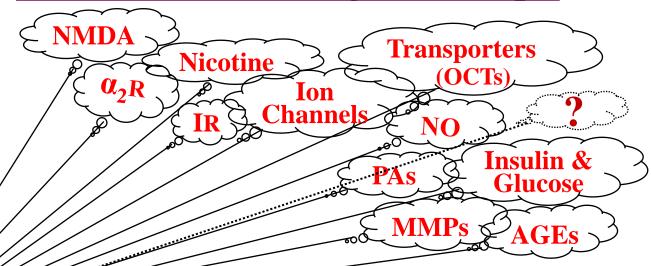
UNIQUE MECHANISM OF ACTION





Firing Squad
(Polypharmacy)

'The Molecular Shotgun Concept'



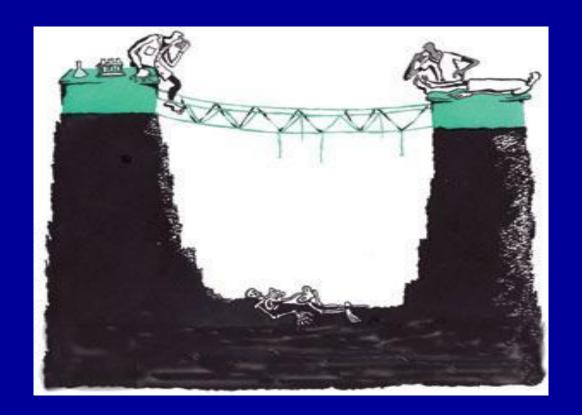


Agmatine Modulates Multiple Molecular Targets



Translating Research Into Practice

"The Valley of Death"
Where some 99% of potential treatments fail





Human Studies

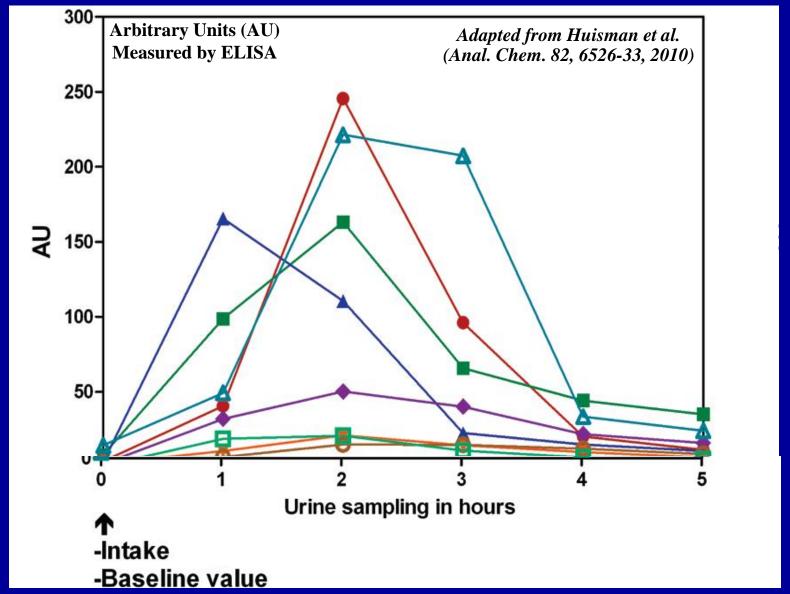


Absorption, Distribution, Metabolism, and Elimination

- Oral agmatine is absorbed by the gastrointestinal tract.
- Agmatine is distributed throughout the body. It crosses the blood-brain-barrier (animal studies).
- Agmatine is mostly converted to putrescine, precursor for polyamine synthesis, and urea. Some may be oxidized.
- Agmatine is excreted by the kidneys.

Agmatine in Urine Following Oral Intake (2 hr apparent blood half-life)







Addressed Conditions

Approximately 30 million people in the United States alone cope everyday with circumstances that challenge healthy nerve functions...

Safety Clinical Trial



-Open Label, Dose-Escalating Clinical Study Design-

Keynan et al., Pain Med., 2010;11(3):356–368

Cohort	Participants [Male] [Female]		Treatment Regimen
1 st Cohort	[3]	[2]	1 capsule 3 times daily (1.335 g G-Agmatine/day) - 10 day treatment
2 nd Cohort	[1]	[4]	2 capsules 3 times daily (2.670 g/day) - 10 day treatment
3 rd Cohort	[7]	[5]	8 daily capsules (2 in the morning, 3 at noontime and 3 in the evening) (3.560 g/day) - 10 day treatment
4 th Cohort	[8]	[4]	8 daily capsules (2 in the morning, 3 at noontime and 3 in the evening) (3.560 g/day) - 21 day treatment

Adverse Effects During 7-month Follow-up	1st Cohort	2 nd Cohort	3 rd Cohort	4th Cohort
Mild Diarrhea			1	2
Discontinuations – Due to the above Adverse Effects				1
Free Will Discontinuations				1

Long-Term Safety

-A Case Report-



Gilad and Gilad, J. Medicinal Food, 2014;17 (11):1–4

Regimen: The recommended daily dose of 2.67 g oral G-Agmatine® was taken over a period of 4–5 years. The regimen consisted of six gelatin capsules, each containing 445 mg, taken twice daily three in the morning and three in the evening after meals.

Measures: Clinical follow-up consisted of periodic physical examinations and laboratory blood and urine analyses.

Results: All measurements remained within normal values and good general health status sustained throughout the study period, up to 5 years.

The studies provide evidence for the long-term safety of a high daily dosage of G-Agmatine® and indicate that it is non-addictive.

Translation of Research into Treatment Fight For Nerve Health



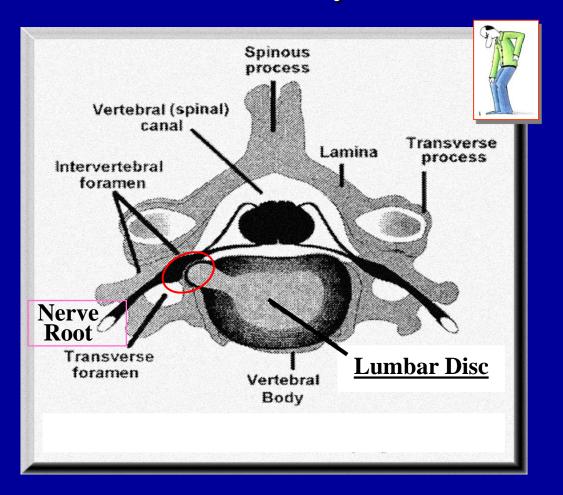
-Clinical Study Focused on Sciatic Nerve-



Clinical Study



- ➤ About 1.5% of the population cope everyday with circumstances that challenge healthy sciatic nerve functions.
- > Sciatic nerve functions can be clearly defined and measured.



Efficacy Study — Sciatic Nerve Functions

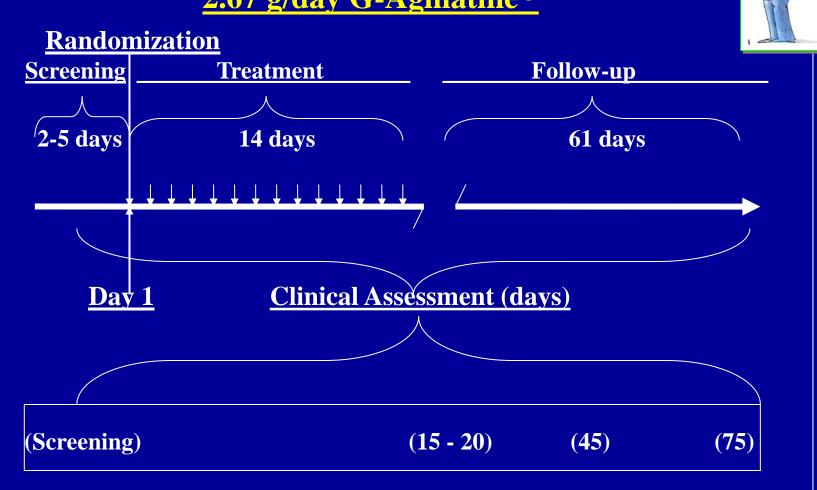


Randomized, Double-Blind, Placebo-Controlled Trial (RCT)

Keynan et al., *Pain Med.*, 2010;11(3):356–368

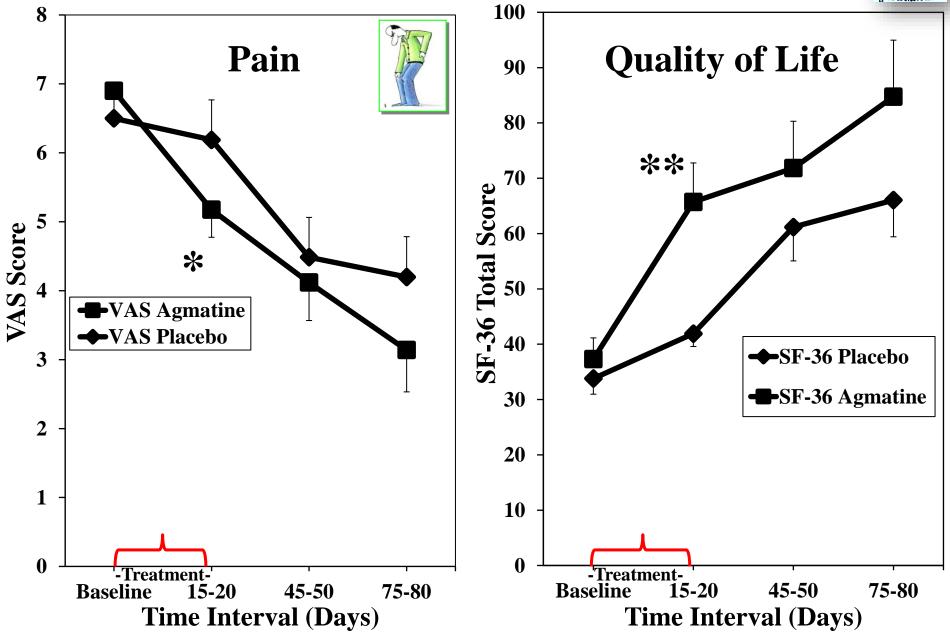
-Study Design-





Efficacy of G-Agmatine®







Effectiveness in Small Nerve Fiber- Associated Pain

Open-Label Consecutive Case Series Study

Rosenberg et al., Nutrients, 2020;12(2):576

> Small fiber nerve malfunctions underlie symptoms associated with various types of bodily systems disorders

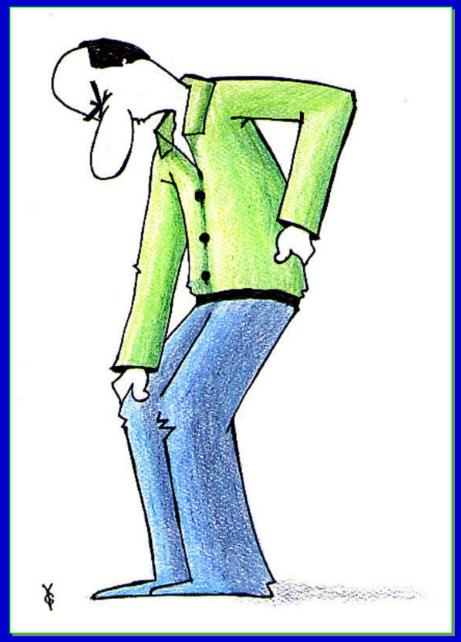
Study Regimen- 2.67 g/day G-agmatine® for 60 days

Pain questionnaire ratings before and after 60-day treatment with G-Agmatine®

Patients (11)	Average Pain Before Treatment	After	Absolute Decrease (Rating Points)	Percent Decrease
Average ± SD	56.1 ± 18.2	30.1 ± 16.9	26.0 *	46.4% *

^{*}Highly significant (p < 0.00001, paired t-test). SD, standard deviation.





'Proof-of-concept'

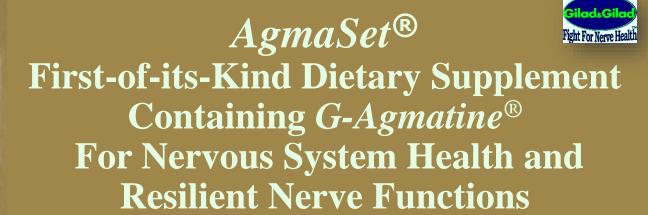
The human clinical studies
serve a

Proof-of-Concept indicating the
safety and effectiveness of
G-Agmatine® treatment for
healthy nerve functions under
stressful conditions



Recommended Effective Regimen

- > 2 to 3 capsules twice a day with or after meals for as-long-as needed (each capsule contains 450mg of *G-Agmatine*®).
- Time to effect -
 - In acute circumstances: 2 4 days in most but may be as-long-as 2 to 4 weeks.
 - Chronic conditions: 3 6 weeks.
- ➤ After the effect is established the dosage may be gradually reduced for maintenance regimen.



For Healthy Nerve Functions*

DIRECTIONS FOR USE: Adults, 2 capsules twice or three-times a day with or after meals, or as recommended by your physician, pharmacist or healthcare professional.

<u>NOTE</u> - Pregnant or nursing women, children and ill or medication-taking people should consult a physician befor use. May cause diarrhea.

- o Keep Out of the Reach of Children.
- Do Not Use If Printed Seal is Broken or Missing
- Store in a Closed Bottle at Temperature Not Exceeding 30°C (86°F) and Humidity Not Above 70% RH.

*These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure, or prevent any disease.



Gilad&Gilad LLC Las Vegas, NV 89129. Tel: 1-888-484-4523 info@ForNerveHealth.com https://ForNerveHealth.com



For Healthy Nerve Functions

Dietary Supplement 120 Capsules

Supplement Facts

Serving Size: 2 Capsules Servings per Container: 60

Amount per Servin	% Daily Value	
Calories:	less than 1	
Agmatine Sulfate (as G-Agmatine®)	900 mg	†

†Daily Value Not Established

Other Ingredients: Gelatin, titanium dioxide (as capsule color), water. No allergens, preservatives or additives. Gluten-Free. Each bottle contains one (1) nontoxic desiccant packet. G-Agmatine® is a trademark brand of Gilad&Gilad LLC. Patents: US 8,916,612; US 9,585,852; US 11,622,949 B1.

MADE IN THE

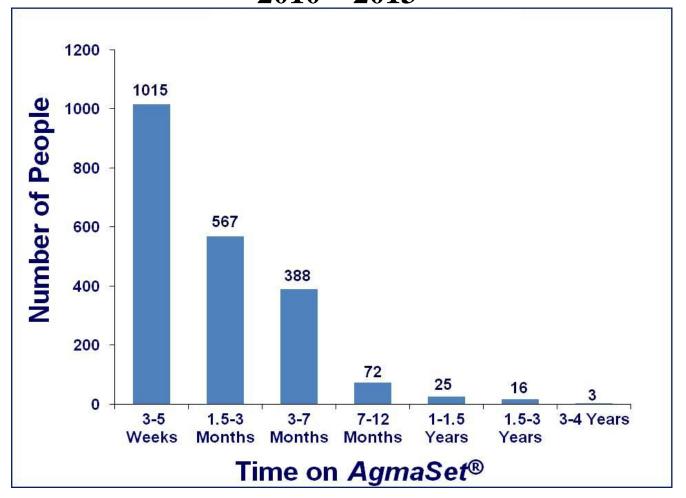
USA

with domestic and imported ingredients



Safety and Effectiveness of *AgmaSet*® 2010 – 2013*





*Surveillance of people who took AgmaSet® during the years 2010 and 2013.

-No adverse effects were reported. -Three people reported soft stool or mild diarrhea.



Animal Studies

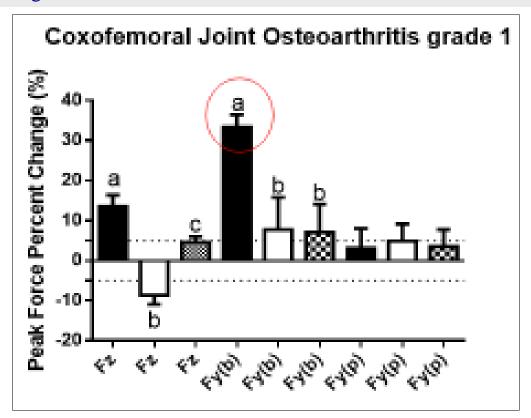


Effectiveness in Dog Hip Joint-Associated Symptoms



-A Randomized, placebo-controlled, crossover design-

Taguchi et al., Frontiers Vet. Med., 2018; 5, Article 311



Peak force percent change (mean +/- SEM) in dogs with grade 1 coxofemoral (hip) joint osteoarthritis following treatment with agmatine (black bars), placebo (white bars), or carprofen (gray bars). Fz, vertical force; Fy(b), braking force; Fy(p), propulsion force. Letters indicate significant differences among treatment groups (p < 0.05). Dashed horizontal lines demarcate 5% positive (upper) or negative (lower) changes.

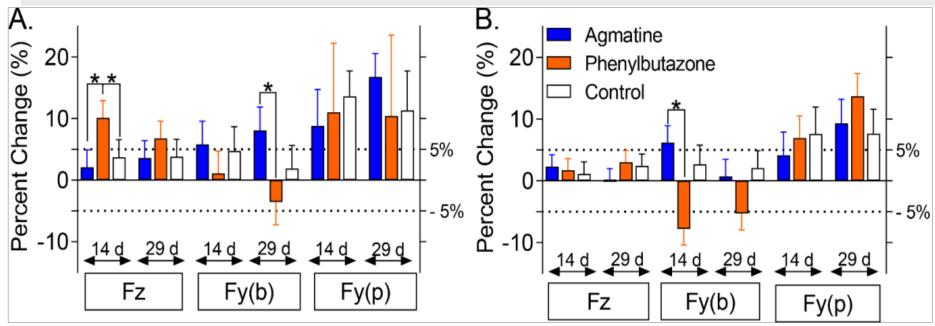




Effectiveness in Horse Lameness and Gastric Ulceration

-A Randomized, placebo-controlled, crossover design-

Taguchi et al., J. Clin. Med., 2022; 11, 7283



Percent change in lameness scale (LS mean \pm SEM) in thoracic limb ground reaction forces over the baseline. Peak force (A) and impulse (B) in vertical (Fz) and craniocaudal braking (Fy(b)) and propulsion (Fy(p)) after 14 and 29 days of oral agmatine (blue), phenylbutazone (orange), or control (white) administration. Asterisks indicate differences among treatments within time points (p < 0.05). A 5% change from the baseline (dotted lines) is indicated.

2

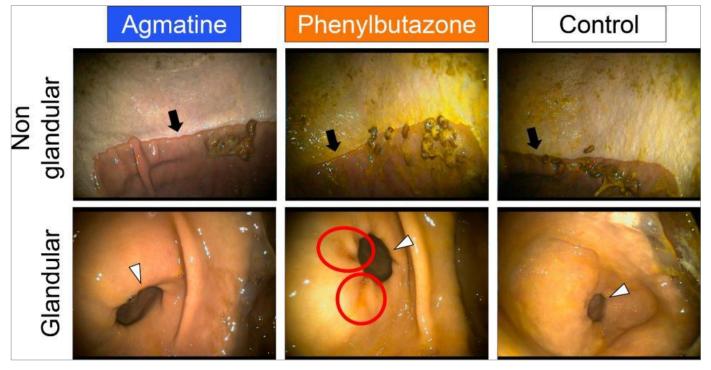




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Representative gastroscopic images of equine non-glandular and glandular gastric mucosa after 30 days of oral agmatine (**left**), phenylbutazone (**middle**) or control (**right**) administration. Ulcers in the glandular region are indicated with red circles. The margo plicatus (black arrows) and pylorus (white arrowheads) are also visible.





AgmaVet® For Your Animal's Daily Diet Containing G-Agmatine® For Resilient Nerve Functions

For Healthy Nerve Functions

Directions for use:

Adult animals - Recommended administration is 1 to 3 capsules a day (depending on body weight) with or after meals. Initial administration period is 4 to 6 weeks; however, some animals may respond in a shorter time period. The number of capsules may then be gradually reduced to an effective maintenance level.



Note - For pregnant or nursing females, young ones or ill and medication-taking animals, consult a veterinarian or health professional before use. May cause diarrhea.

Visit http://ForNerveHealth.com



Gilad&Gilad LLC Georgetown, TX 78633. Tel: 1-888-484-4523 info@ForNerveHealth.com http://ForNerveHealth.com



120 Capsules

Each Capsule Contains:

Agmatine Sulfate (as G-Agmatine_®)......450 mg*

Other Ingredients: Gelatin, titanium dioxide (as capsule color), water.
No allergens, preservatives or additives. Gluten-Free.

Each bottle contains one (1) nontoxic desiccant packet.

*Not recognized by the AAFCO as an essential animal nutrient.

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- Keep out of the reach of children.
- Do Not Use If Printed Seal is Broken or Missing.
- Store in a Closed Bottle at Temperature Not Exceeding 30°C (86°F) and Humidity Not Above 70% RH.

MADE IN THE

USA

with domestic and imported ingredients





"Alternative" Agmatine Sulfate-Containing Products

- ➤ All alternative products containing agmatine sulfate are based on borrowed science. All clinical studies are "borrowed" from Gilad&Gilad.
- ➤ Products targeting the bodybuilding field are all touted based on scientifically unsubstantiated claims and on downright erroneous scientific information.
 - No evidence for enhanced muscle protein synthesis.
 - ❖ No evidence for enhanced Growth Hormone functions.
 - Agmatine may modulate Nitric Oxide (NO) synthesis but does not enhance NO content.

THANK YOU! PLEASE SPREAD THE GOOD NEWS





Any Questions?

Call us: (toll-free) 1-888-484-4523, or 1-702-233-5553

Contact us via email: info@ForNerveHealth.com

https://ForNerveHealth.com