



# Tieraona Low Dog, MD

Chair: US Pharmacopeia Dietary Supplement Admissions, Evaluation and Labeling Expert Committee

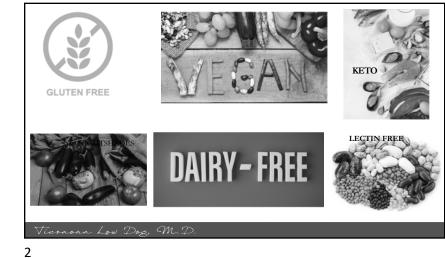
Guiding Teacher: NY Zen Center Fellowship in Contemplative Medicine

National Geographic's Life Is Your Best Medicine Healthy At Home Fortify Your Life Guide to Medicinal Herbs

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# <image>

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# Quality

- **Domestically produced products** *generally* better quality than those imported from China or India.
- 3<sup>rd</sup> party testing (e.g., **USP**, NSF, CL, NNFA) quality indicator.
- Be cautious with products in weight loss, sexual enhancing or bodybuilding categories.
- **Consumer Labs** can be a useful website for assessing quality.

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00/04/2822	Wonderful Huney	n/a	n/a	Examination of imported goods	Undeclared sildenafil	Public Notification	Oruga
02/14/2822	Rise Up Red Edition Capsules	Positive Health	n/a	amazon.com	Undeclared tadalahi	Recel	Drugs
62/94/2822	Red Mammoth capsules	Celebrate Taday	n/a	amazon.com	Undeclared tadalahi and sildenahi	Recall	Oruga
02/06/2922	The Red Pill	Your Favorite Shop	n/a	amazon.com	Undeclared tadalafi	Recall	Drugs
02/04/2922	MAC DADOY PURPLE capsules	ABC SALES 1 INC	n/a	amazon.com	Undeclared tadalahi	Recal	Druga
62/94/2822	MAC DADDY RED capsules	ABC SALES 1 INC	n/a	amazon.com	Undeclared tadalahi and sildenahi	Recall	Drugs
62/04/2822	MogNan Performance Booster capsules	Jump LLC	n/a	amazon.com	Undeclared tadalafi	Recall	Drugs
01/28/2922	Hard Dawn Rise and Shine capsules	Esupplementaalies, ULC	n/a	amazon.com	Undeclared tadalafi	Recall	Drugs
01/07/2022	Taewon Liar	Varieus Distributors	n/e	etby.com	Undeclared metosicam	Public Notification	Drugs
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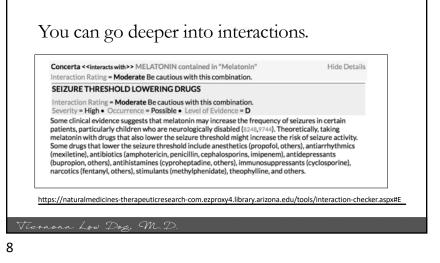
https://www.accessdata.fda.gov/scripts/sda/sdNavigation. cfm?filter=&sortColumn=1d&sd=tainted\_supplements\_cd er&page=1\_Accessed March 8, 2022

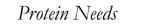


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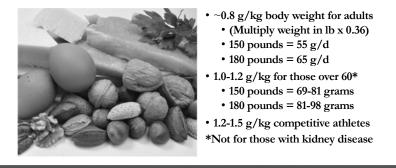




- From Greek *protos*, "first." Build **new cells**, **maintain tissues** (e.g., muscles, inner bone, hair, nails), create **enzymes**, **make hemoglobin**, **lipoproteins to transport cholesterol; present in membrane of every living cell**.
- Sources include meat, poultry, seafood, eggs, soy products, nuts, nut butters, beans, peas, and seeds.
  - Choose seafood 2 x weekly (low in mercury, high in omega 3s, sustainable)
  - Include beans and peas often in your diet, as a side or main dish
  - Chicken and poultry regularly, red meat 10 ounces per week. Avoid processed red meats.

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How Much Protein Do You Need Per Day



Per 20g servi

Calories (kcal)

17

1

190

2.3

15.3

1.2

2.4

0

4.32

Protein (g)

Fat (g)

Carbs (g)

In USD

Sodium (mg)

Cost (per 100g)

77

0

6.5 (5.2g fiber)

15.5 9

1.1 1.9

1.5

192

3.96 2.82

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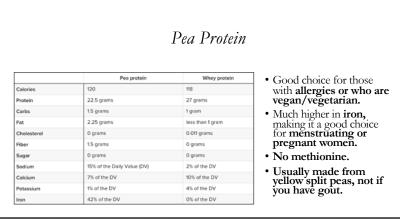
Food	Portion Size	Protein (g) (approximate)	
Meat, fish, or poultry	75g (2 ½ oz) / 125 mL (½ cup)	21	
Firm tofu	150g / 175 mL (¼ cup)	21	
Egg, chicken	2 large	13	
Cheese	50 g (1 ½ oz)	12	
Fortified soy beverage	250 mL (1 cup)	6-8.5	
Cooked dried beans, peas, or lentils	175 mL (% cup)	12	
Cow's milk	250 mL (1 cup)	9	
Yogurt	175 mL (¼ cup)	8	
Peanut butter or other nut spread	30 mL (2 Tbsp)	8	
Nuts or seeds	60 mL (¼ cup)	7	
Bread	1 slice (35g)	3	
Cereals, cold	30 g	3	
Cereals, hot	175 mL (¼ cup)	3	
Pasta or rice	125 mL (½ cup)	3	
Vegetables	125 mL (½ cup) or 250 mL (1 cup)	2	
	lettuce		
Fruit	1 fruit or 125 mL (½ cup)	1	

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# Protein Powders

- Whey protein hydrosolate: digested quickly after workout)
- Casein: "time-release protein" take before bed for muscle recovery
- Egg white: no fat, equal to whey. Allergy.
- Soy: complete protein, no carbs/fiber, highly absorbed. Allergy.
- Hemp: omega 3, fiber, less protein
- All of the above are "complete proteins" and highly absorbable.
- Brown Rice hypoallergenic
- Pea see next page

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# Collagen is Essential for Strength and Structure

 Collagen is a protein that forms a scaffold that protects tissues; it is abundant in the skin, bones, muscles, tendons. Collagen production declines with age and exposure to smoke, UV light, high sugar diet, and lack of sleep, leading to wrinkles, osteoarthritis, etc.

- 80-90% of collagen are types 1, 2 and 3.
  - Type I Skin Elasticity
  - Type II Joints and Immune System
  - Type III Skin, Blood Vessels, Vital Organs
  - Type IV Cell Health and Growth
  - Type V Skin, Tendons and Immune System
  - Type X Joints, Bones, Muscles and Hair

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# Oral Collagen: Meta-Analysis for Skin

- 19 randomized, double-blind, controlled studies were reviewed<sup>1</sup>
  - N=1125, ages 20-70 years, 95% female
- **Conclusions:** favorable results of *hydrolyzed* collagen supplementation compared with placebo in terms of **skin hydration, elasticity, and wrinkles** when 6-10 g/d taken for at least 90 days. Good safety profile, no adverse effects.<sup>1,2</sup>
- Note: studies are small and mostly subsidized by collagen companies.

1. de Miranda RB, et al. Effects of hydrolyzed collagen supplementation on skin aging: a systematic review and meta-analysis. Int J Dermatol 2021; 60(12):1449-1461.

2. Choi FD, et al. Oral Collagen Supplementation: A Systematic Review of Dermatological Applications. J Drugs Dermatol 2019; 18(1):9-16.

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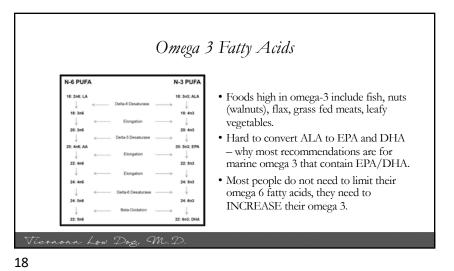
# Oral Collagen for Musculoskeletal Pain

- Studies in healthy adults and those with knee OA report that 10 g/d of *hydrolyzed* collagen reduces join pain and improves function.
- Small studies assessing consumption of **undenatured type II collagen** (most abundant protein in joints/spine) alone or with vitamin D show a reduction of **musculoskeletal pain and improvement in joint function.**
- Undenatured type II collagen *more effective* than glucosamine and chondroitin sulfate supplements in companion animals. Dose generally 40 mg per day.

Mohashari A, et al. A White Paper on Gollagen Hydrolyzates and Ultrahydrolyzates Potential Sapplements to Support Joint Health in Ostcoarthritis? *Carr Rhomani Rep* 2021 '22(11):78 Rodrigen Mondowa C, et al. Effects of Naminoal Intercentions in the Control of Manahokchell Pairs. An Integrative Review, Namina 2020, 120(10):79. Georgia H, et al. Understarted Type I Georgiange UC: In 1 ion Inter Main and Davas Rodrigen on the Carrest Annology of Company animal. Annal. 2020;102(10):70

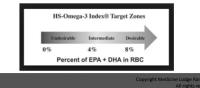
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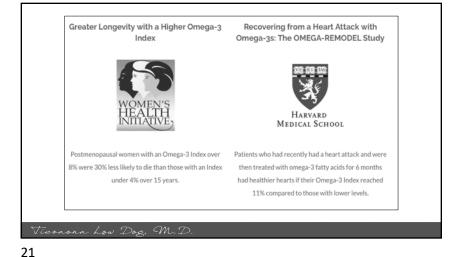


No Fish Story: The Omega 3 Index

- Omega-3 Index test is now the gold standard for omega-3 biostatus testing. It is used as a compliance marker for randomized controlled trials with fish oil supplements, and in epidemiological research.
- In 2008 Dr. Bernadine Healy, cardiologist and past President of the AHA and first woman Director of the NIH said, "Before long, your personal Omega-3 Index just could be the new cholesterol—the number you want to brag about."

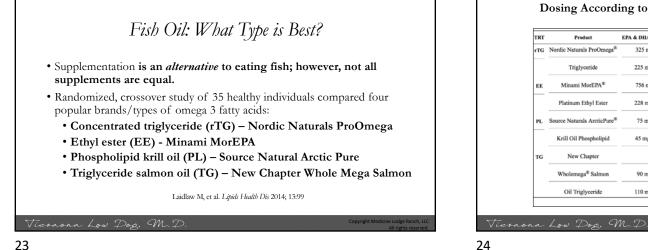






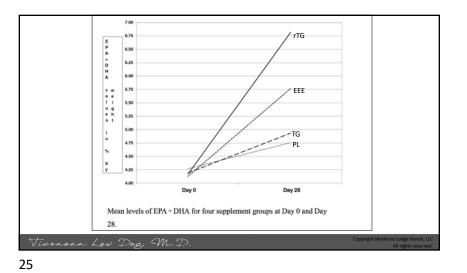


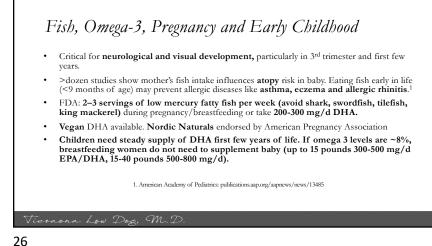


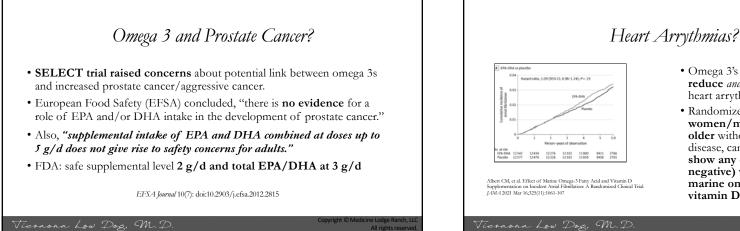


# Dosing According to Manufacturer's Recommendations

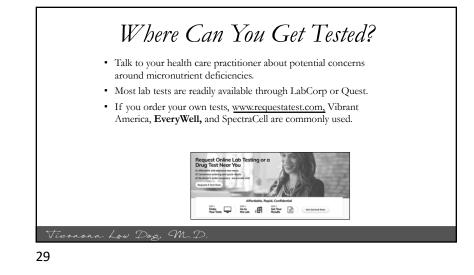
TRT	Product	EPA & DHA per capsule*	Tested values	Label use: caps/day	Daily dosage of EPA + DHA
rTG	Nordic Naturals ProOmega®	325 mg EPA	329.6 mg EPA	2	EPA: 650 mg
	Triglyceride	225 mg DHA	226.0 mg DHA		DHA: 450 mg
EE	Minami MorEPA®	756 mg EPA	774.2 mg EPA	1	EPA: 756 mg
	Platinum Ethyl Ester	228 mg DHA	233.7 mg DHA		DHA: 228 mg
PL	Source Naturals ArcticPure®	75 mg EPA	78.0 mg EPA	2	EPA: 150 mg
	Krill Oil Phospholipid	45 mg DHA.	46.7 mg DHA.		DHA: 90 mg
TG	New Chapter				
	Wholemega® Salmon	90 mg EPA	96.4 mg EPA	2	EPA: 180 mg
	Oil Triglyceride	110 mg DHA	109.5 mg DHA		DHA: 220 mg







- Omega 3's have been said to reduce *and* increase the risk of heart arrythmias.
- Randomized clinical trial 25,119 women/men aged 50 years or older without cardiovascular disease, cancer, or AF failed to show any effect (positive or negative) with 1 gram/d marine omega 3, 2000 IU vitamin D, or combo.



# Micronutrient Status of Americans

- 31% population at risk for at least one vitamin deficiency or anemia.
- Deficiency risk most *common in women* (37% overall): 19–50 years (41%), and pregnant or breastfeeding women (47%).
- Those who did NOT take dietary supplements had *highest risk* of any deficiency (40%), compared to users of full-spectrum MVI-mineral supplements (14%).
- Individuals consuming an adequate diet based on EAR had a lower risk of any deficiency (16%), compared to those with inadequate diet (57%).

Bird JK, et al. Nutrients 2017 Jun 24;9(7):655.

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Real State of Our Nutrition

- 90 million Americans are vitamin D deficient (using the Endocrine Society guidelines < 20ng/mL)</li>
- 30 million are deficient in vitamin B6
- 18 million people have B12 deficiency
- 16 million have scurvy (by serum levels)
- 13% of Latinas and 16% of African American women (ages 12–49) are **iron deficient**
- Women 25–39 overall have borderline iodine insufficiency

CC: 2<sup>M</sup> Attional Report on the Biochemical

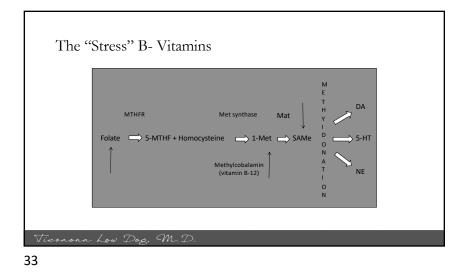
CDC: 2<sup>na</sup> National Report on the Biochemical Indicators of Diet and Nutrition in the U.S. population

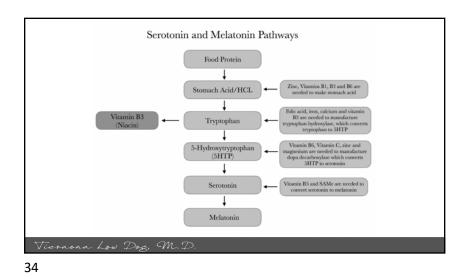
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# **B-Vitamins Fast Facts**

- 8 B-vitamins partner together, which is why you almost always want to take them together.
- Vitally important for energy, and maintenance of healthy brain, nervous and cardiovascular system.
- Low levels of vitamin B6 and B12, increase risk for *depression and impair* cognition, attention, and memory.
- Subset of women taking birth control pills are deficient in vitamin B6.
- $2/3^{rd}$  of those with B12 deficiency are over age 50.
- Women may not get sufficient folate, significant if pregnancy occurs.

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# Oral Contraceptives & Vitamin B6

- Majority of women on OCPs have low serum B6 levels, even when meeting dietary RDA.1
- Oral contraceptive use in US 2017–2019:2
- 19.5% of women aged 15–19
- 21.6% aged 20-29
- 10.9% aged 30-39
- 6.5% of women aged 40-49
- Low B6: microcytic anemia, depression, poor concentration, fatigue, etc.
- Women who discontinue OCs and become pregnant may be at increased risk for preterm birth, early pregnancy loss, and difficulty conceiving.3,4



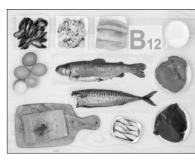
Wilson SMC. Nutr Rev 2011 Oct; 69(10):572-83 Ho CL, et al. Nutrients 2016; Sep 1;8(9)

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# Vitamin B12

- 18 million Americans deficient
- Geriatric population prevalence 21%
- Risk for deficiency:
  - · Inadequate intake
  - Veganism
  - Malabsorption
  - Medications (PPI, metformin)
  - · Obesity
  - Aging

• 25-250 mcg per day if at risk



Singh NN, et al. Vitamin B-12 Associated Neurological Disease. 2018; Oct 22 https://emedicine.medscape.com/article/1152670-overview#a6

# Screening for B12 Deficiency

**Even marginal B12 status** reduces cognitive function, mood, and energy.

Screening *should be considered* in those with one or more risk factors for low B12:

- Inflammatory bowel disease
- Use of **metformin** >4 months
- Use of PPIs/H2 blockers >12 months
- Vegans, possibly vegetarians
- Bariatric surgery
- · Those over 70 years old

Langan RC, Goodbred AJ. Am Fam Physician 2017 Sep 15;96(6):384-389

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# Metformin & PPI Increase Risk for Low B12

- 2015 meta-analysis: 80% increased risk B12 deficiency after 10 months of regular proton pump inhibitor use.
- Meta analysis 29 studies: 245% increased risk B12 deficiency with metformin use. Low B12 increases progression of diabetic neuropathy.
- B12 deficiency can lead to **difficulty walking**, **tingling/numbness** in hands and feet, **fatigue**, shortness of breath, **loss of appetite**, **joint pain**, **depression**, **loss of taste and smell**, **cognitive impairment**, and **dementia**.
- B12 should be monitored every 1-2 years if taking these medications.

### Jung BJ, et al. Association hereeves viamin B12 deficiency and long-term use of acid-lowering agents: a systematic review and next-analysis. *Intern Mal J* 2017; 55(3):409-36. Out M, et al. Long-term treatment with metformin in type 2 diabetes and methylmatonic acid. Post hoc analysis of a randomized controlled 4.3year trial. J Diabatic Complications 2017; 52(2):171-178. Nature M, et al. Intern Euroge Md 2015; 10(3):973-106.

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# Downside of PPI Acid Suppression

- · Dysbiosis and SIBO (probiotics)
- Rebound hypersecretion of acid (taper)
- · Gastrointestinal infections: FDA warning (probiotics)
- Pneumonia (vitamin D, immune support)
- Fracture: FDA warning (vitamins D3, K2, calcium, mag)
- Food allergies (protease, betaine HCl, probiotics)
- Deficiencies: FDA warning magnesium. In addition, vitamins B12 and C, and iron
- Kidney injury (monitor, limit salt)
- Cognitive decline\*
- Gastric cancer\*
- \* Correlation but not strong evidence at this time

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R S K K

# Vitamin B12 in Elders



- 10–30% of elders have **atrophic gastritis**, which can interfere with B12 absorption. Deficiency can cause weakness, cognitive/behavioral changes.
- National Academy of Medicine recommends *all* adults over 50 get majority of vitamin B12 from supplement or fortified foods.
- Intakes *well above RDA* may be required in older adults to maintain vitamin B12 status.

Langan RC, Goodbred AJ. Am Fam Physician 2017 Sep 15;96(6):384-389

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Proton Pump Inhibitors & Fracture

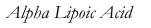
- FDA: patients taking high doses of PPIs and/or taking one year or more at highest risk. Warning label mandated.
- Recent study of adverse event reporting at FDA showed that PPI also increases fracture of ribs and other sites.
- American Geriatrics Society recommends *against* PPIs for longer than eight weeks in older adults, except in high-risk patients, due to the potential risk of bone loss, fractures and *C. difficile* infection

http://www.fda.gov/Drugs/DrugSafety/PostmarketDrugSafetyInformationforPatientsandProviders/ucm213206.htm

Wang J., et al. Proton Pump Inhibitors and the Risk for Fracture at Specific Sites: Data Mining of the FDA Adverse Event Reporting System. Sci Rep 2017 Jul 17,7(1):5527. American Genatrics Society 2015 Updated Beers Criteria for Potentially Inappropriate Medication Use in Older Adults. J. Am Genatr Soc, 63(11):2227-46.

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- α-lipoic acid (ALA, thioctic acid) naturally occurring compound produced by humans (and others), resides in mitochrondria.
- Johns Hopkins review: multiple studies show ALA can improve pain of diabetic peripheral neuropathy (moderate strength of evidence).<sup>1</sup>
- 200 people diabetic peripheral neuropathy: significant improvement in vibration perception threshold, neurological symptom score and disability score, and visual analog scale in group receiving add on 600 mg ALA (BID) for 6 mo., compared to placebo.<sup>2</sup>
- May be beneficial for **burning mouth syndrome**.
- 1. Nesbit SA, et al. Non-pharmacologic treatments for symptoms of diabetic peripheral neuropathy: a systematic review. Curr Med Res Opin 2019
- El-Nahas MR, et al. Oral Alpha Lipoic Acid Treatment for Symptomatic Diabetic Peripheral Neuropathy: A Randomized Double-Blinded Placebo-Controlled Study Endor Metab Immune Diand Drug Targets. 2020

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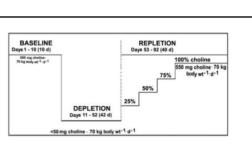




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- Choline deficiency causes abnormal deposition of fat in the liver, which results in a condition called *nonalcoholic fatty liver disease*.
- Necessary for healthy cell membranes and cognition as we age.
- Particularly crucial during pregnancy and first three years of a child's life, improves cognitive function in adulthood, prevents age-related memory decline, and protects brain from neuropathological changes associated with Alzheimer's disease, and neurological damage associated with epilepsy, fetal alcohol syndrome, and Down's.
- New daily value set in 2016: 550 mg per day Jiang X, et al. Trends Endocrinol Metab 2014; 25(5):263-73. Wozniak JR, et al. Nutr Res 2013; 33(11):897-904

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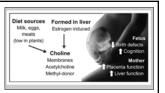
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- 57 healthy adults were fed choline-deficient diets under controlled conditions.
- Results showed that 77% of men, 80% of postmenopausal women, and 44% of premenopausal women developed fatty liver, liver damage, and/or muscle damage.
- Dysfunction corrected when choline was reintroduced into diet. Fischer 1M, et al. Am J Clin Nutr. 2007;85(5):1275-1285.

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# Choline in Pregnancy



- ACOG, AMA, AAP, EFSA, WHO: all conclude that choline is *crucially important* during pregnancy.<sup>1</sup>
- ~90-95% pregnant women consume less than recommended amounts of choline.
- Review found supplementing mother/child's diet with choline over first 1,000 days of life:
  - Supports normal brain development.
  - Protects against neural/metabolic insults, particularly when fetus exposed to alcohol.<sup>2</sup>
  - Improves neural and cognitive functioning.3

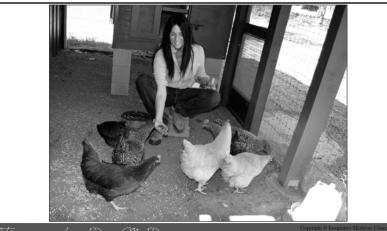
1. Schwarzenberg SJ. Paliatrix. 2018:141. doi: 10.1542/peds.2017-3716. 2. Derbyshine E, et al. Natrients 2020 Jun 10;12(6):1731. 3. Brunst KJ, et al. RMJ 2010 May 20;340:c2181.

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Choline Content Per Serving	Milligrams (mg) per	Percent
Food	serving	DV*
Beef liver, pan fried, 3 ounces	356	65
Egg, hard boiled, 1 large egg	147	27
Beef top round, separable lean only, braised, 3 ounces	117	21
Soybeans, roasted, ½ cup	107	19
Chicken breast, roasted, 3 ounces	72	13
Beef, ground, 93% lean meat, broiled, 3 ounces	72	13
Fish, cod, Atlantic, cooked, dry heat, 3 ounces	71	13
Potatoes, red, baked, flesh and skin, 1 large potato	57	10
Wheat germ, toasted, 1 ounce	51	9
Beans, kidney, canned, ½ cup	45	8
Quinoa, cooked, 1 cup	43	8
Milk, 1% fat, 1 cup	43	8
Yogurt, vanilla, nonfat, 1 cup	38	7
Brussels sprouts, boiled, ½ cup	32	6
Broccoli, chopped, boiled, drained, ½ cup	31	6
Mushrooms, shiitake, cooked, ½ cup pieces	27	5
Cottage cheese, nonfat, 1 cup	26	5

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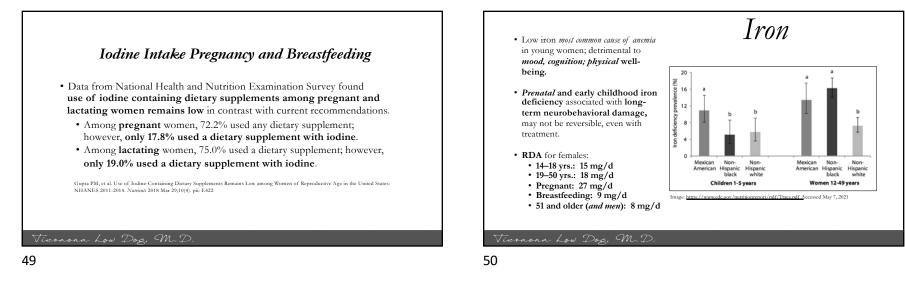
# Iodine Crucial Brain Health

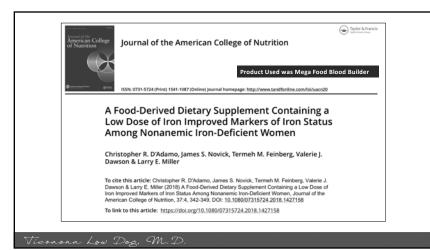
- Crucial nutrient, especially during pregnancy, infancy, and childhood when thyroid hormones regulate growth in developing brain.
- Mild deficiency associated with autistic spectrum disorder, ADHD, learning disabilities, and dyslexia.<sup>1</sup>
- American Thyroid Association: women planning on becoming pregnant, are pregnant or lactating take supplement containing 150 mcg/d potassium iodide.<sup>2</sup>

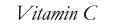


Hay I, et al. Nutrients 2019 Aug 22;11(9):1974
 Stagnaro-Green A, et al. Thyroid. 2011;21(10):1081-1125

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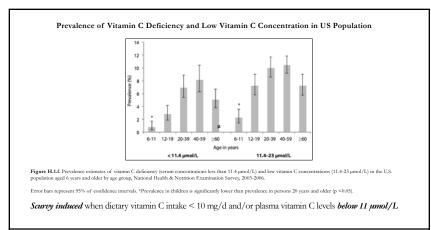


- Plays crucial role in innate and adaptive immune responses.
- Maintains *epithelial barrier* function, increases alveolar fluid clearance, and attenuates proinflammatory response.
- Deficiency leads to impaired immunity and greater risk of infection.<sup>1</sup>
- Only 1 in 10 Americans get recommended daily intake for fruits and vegetables.<sup>2</sup>
- ~15.7 million Americans have serious vitamin C deficiency; ~60 million have marginal status.

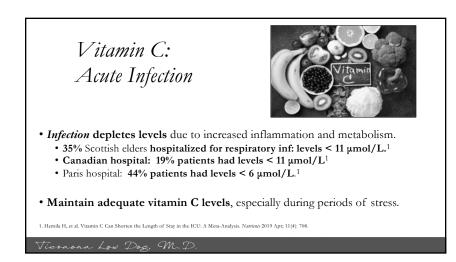


 Vorilhon P, et al. Eur J Clin Pharmaol 2019; 75(3):303-311.
 <u>2https://www.ckc.gov/netwis/wolumes/06/wir/mm6645a1.htm?s\_cid=nem6645a</u> <u>1\_w Accessed November 24, 200</u>
 <u>3https://www.ckc.gov/netmioneport/pdf/Nutrition\_Book\_complete508\_final.p</u>

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*'Picky'' Eater?* 

- Healthy 4-year-old boy seen by ortho/rheumatology with right-leg pain and progressively worse limping, became unable to weight bear.
- Intermittent non-blanching rash arms and legs past 2 years, topical emollients not effective. Bleeding when brushing teeth, gingivitis.
- Diet primarily waffles, yogurt, pasta with butter, goldfish crackers, peanut butter, chicken nuggets, and water.
- Workup negative except for iron, vitamins C and D deficiencies.
- 100 mg ascorbic acid q 8 hours x 7 days, then 1 x daily with iron and vitamin D.
- Limp and rash completely disappeared within weeks.

Nastro A, et al. Scurvy Due to Selective Diet in a Seemingly Healthy 4-Year-Old Boy. Pediatrics September 2019; 144 (3) e20182824.

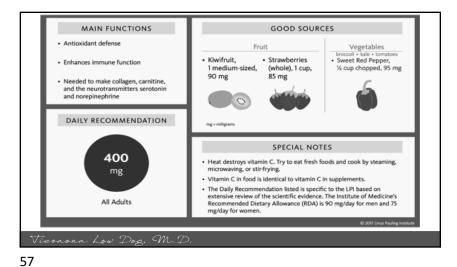
Vitamin C



- Skin changes, easy bruising, slow healing wounds, dry mouth, dry eyes. Emotionally labile. Weakened capillaries. Hemorrhage is hallmark of scurvy and hair follicles are common site of cutaneous bleeding.
- Inflammation of gingiva followed by bleeding, ulceration, and bad breath.
- Swelling of periodontal membranes occur, followed by loss of bone, and loosening of the teeth.
- Low ascorbic acid levels found in **healthy subjects with gingivitis** and in **diabetics with periodontitis**. 250 mg BID improved symptoms.<sup>1</sup>

1. Gokhale NH et al. J Diet Suppl. 2013;10:93–104.

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Vitamin D & Respiratory Infection

people/year.

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# Vitamin D

- Deficiency common globally, more severe in elders due to environmental/biological factors.
- 75% elders in nursing homes severely vitamin D deficient (25(OH)D < 10 ng/mL).
- Obesity, dark skin, living northern latitudes, use of sunscreen, all increase deficiency risk.
- Vitamin D increases innate immunity via secretion of antiviral peptides, strengthening mucosal defenses and reducing risk of respiratory infections.

Fig Pt et al. Fir J Factoriad. 2019;18:25–34. Ab. N. J Hajer Paula: Haudi 2020; Oce; 13(10); 157–1380.

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### Martineau AR, et al. Vitamin D supplementation to prevent acute respiratory tract infections: systematic review and metaanalysis of individual participant data. *BMJ* 2017; 356: i6583.

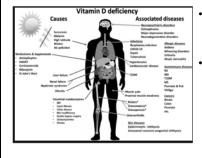
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- Vitamin D releases antimicrobial peptides in the lungs, helps to mount immune response.
  - 25 eligible randomized controlled trials (n=10,933, aged 0–95 years).

• Acute respiratory infection kills ~2.65 million

 Supplementation reduced risk of acute respiratory infection among all participants (NNT=33) and those who were vitamin D deficient experienced the most benefit (NNT=4).

# Vitamin D & Microbiome



### Vitamin D ensures appropriate level of antimicrobial peptides in mucus; maintains intestinal barrier function.

If **bacteria penetrate** epithelial layer and enter interstitium, immune cells trigger adaptive immune response by activating Th1/Th17 cells. *Vitamin D/VDR signaling in these cells ensures clearance* of the bacteria.

Fakhoury HMA, et al. J Steroid Biochem Mol Biol 2020

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# Vitamin D & COVID

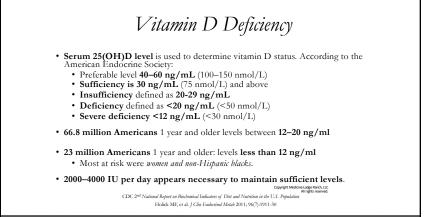
- 212 cases COVID-19: vitamin D deficient patients had 19.6-fold higher risk of critical outcome compared to those with sufficient levels (p < 0.001).<sup>1</sup>
- Retrospective study: **780 confirmed cases SARS-CoV-2** infection found those vitamin D deficient ~13 times more likely to die.<sup>2</sup>
- Israeli data: 26% of COVID patients died if vitamin D deficient soon before hospitalization, compared to 3% who had normal levels of vitamin D.
- Hospitalized patients who were vitamin D deficient 14 times more likely to end up in severe or critical condition than others.<sup>3</sup>

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 Alipia, A. Vianna D. Stephenmanian Gold Prosbil Improve Clinical Outcomes of Patterns Infectod with Generativa-2019 (CVUID-2019). SIRV Fedame J. 2020. doi:10.2193/smrs.3571484.
 Zaharuson P., et al. Patterns of COVID-19 Mortality and Vitamin D. An Indonesian Study. 31704. 2020 doi: 10.2199/smrs.358564
 Sanda S. Silve Cardio and Study Study Control Study. 2019.



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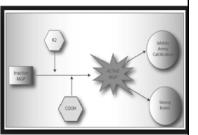
Life-stage Group <sup>a</sup>	IOM Recommendations, IU/d		Endocrine So	ciety Recommendations, IU/db
Life-stage Group*	Intake	Upper Limit <sup>c</sup>	Intake	Upper Limit <sup>c</sup>
0-6 months	400 <sup>d</sup>	1,000	400-1,000	2,000
6-12 months	400 <sup>d</sup>	1,500	400-1,000	2,000
1-3 years	600	2,500	600-1,000	4,000
4-8 years	600	3,000	600-1,000	4,000
9-18 years	600	4,000	600-1,000	4,000
19-30 years	600	4,000	1,500-2,000	10,000
31-50 years	600	4,000	1,500-2,000	10,000
51-70 years	600	4,000	1,500-2,000	10,000
71+ years	800	4,000	1,500-2,000	10,000
Pregnant or lactating women (14-18 years)	600	4,000	600-1,000	4,000
Pregnant or lactating women (19-50 years)	600	4,000	1,500-2,000	10,000
<sup>a</sup> Includes normal healthy individ <sup>b</sup> Estimated intake needed to ma <sup>◦</sup> Maximum level that is expected equivalent to 400 IU. <sup>d</sup> Refers to adequate intake (intak with limited sun exposure and v because of insufficient evidence	intain blood 25 I to have no ris ke estimated to ritamin D store	(OH)D levels above 30 i k of adverse effects to h o maintain protective 25(	ng/mL. ealthy individuals. 1 μg OH)D levels in a group	o of healthy individuals

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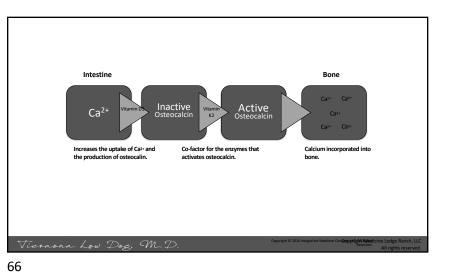
# Don't Forget the Vitamin K

- Calcium, vitamins D and K, and magnesium contribute independently and collectively to bones.
- Beneficial role of vitamin K, particularly vitamin K2, in bone and cardiovascular health reasonably well supported scientifically, with several preclinical, epidemiological, and clinical studies published over the last decade.
- Vitamin K2 (MK-7) 100-200 mcg per day.

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Karpinski M, et al. Roles of Vitamins D and K, Nutrition, and Lifestyle in Low-Energy Bone Fractures in Children and Young Adults. J Am Coll Nutr 2017 Jul;36(5):399-412.



# Magnesium

- Low magnesium intakes and serum levels associated with type 2 diabetes, metabolic syndrome, inflammation, high blood pressure, atherosclerotic vascular disease, sudden cardiac death, chronic pain, osteoporosis, migraine, asthma, and colon cancer.
- 50% of U.S. population consumes less than the required amount of daily magnesium.
- Deficiency associated with negative effects on calcium and vitamin D homeostasis.

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# Magnesium and Inflammation

- Adults consuming < RDA of magnesium **1.48–1.75 times more likely to have elevated hs-CRP** than those with adequate intake.
- Oral magnesium supplementation decreases CRP levels in seniors, those who are obese and/or with prediabetes.
- Hypomagnesemia may accentuate pain by unblocking the NMDA receptor (involved in central sensitization). Magnesium creates a blockade of the NMDA receptor in the spinal cord.
- Meta-analysis 20 studies: magnesium alleviates acute postoperative pain and *enhances effect of opioids* without increase in side effects.

Shmagel A, et al. Low magnesium intake is associated with increased knee pain in subjects with radiographic knee osteoarthritis data from the Osteoarthritis Initiative. Initiative. In

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# Magnesium Deficiency

- Magnesium deficiency: *menstrual cramps, leg cramps, increased pain, migraines, fatigue, anxiety, insulin resistance, heart arrythmia, etc.*
- Severe cases of deficiency, seizures, tingling and numbness in arms and legs, bizarre muscle movements (especially of eyes and face), personality changes, and coronary spasms can occur.
- Magnesium citrate, malate, glycinate are much better tolerated than magnesium oxide.
- **Supplementing 300–600 mg/d** very safe. Larger doses should not be used in those with impaired kidney function.

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# Resources

- · Fortify Your Life and Guide to Medicinal Herbs with National Geographic
- Dietary Supplement Label Database: dsld.nlm.nih.gov
- NIH National Center for Complementary and Integrative Health (NCCIH): nccih.nih.gov
- Office of Dietary Supplements: ods.od.nih.gov
- Linus Pauling Institute: lpi.oregonstate.edu
- Consumer Labs: www.ConsumerLabs.com
- Natural Medicines Research Collaboration
  https://naturalmedicines.therapeuticresearch.com

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