







Enhancing Oral and Systemic Health Through Nutrition and Dietary Supplements

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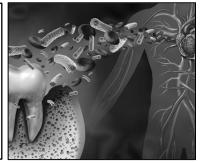
Outline of Talk

- Nutrition and micronutrients significantly impact our health
 - Oral health plays a vital role in systemic health
 - Drivers of oral and systemic inflammation
 - Role of food and supplements in inflammation
 - The microbiome's role in overall health
 - Micronutrients for gum, gut, and mucosal health

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Periodontitis

- Chronic, multi-bacterial infection that elicits low-grade systemic inflammation through the release of pro-inflammatory cytokines, as well as local invasion and long-distance translocation of periodontal pathogens.
- It can induce or exacerbate other chronic systemic inflammatory diseases, such as atherosclerosis and diabetes, and can lead to adverse pregnancy outcomes.



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Oral and Systemic Health

- 1891: Oral microbiologist Willoughby D. Miller suggests that oral infections can affect other body parts and are related to systemic diseases.
- 1912: Frank Billings speculated that tooth infection may cause rheumatoid arthritis, nephritis, endocarditis, and other diseases.
- Unchecked, this low-grade inflammation can *disrupt the body's overall health or worsen existing systemic diseases.* This underscores the crucial role of maintaining good oral health in *preserving overall health.*

Miller WD. The human mouth is a focus of infection. Lanet 1891; 138, 340–342 Billings F. Chronic focal infections and their etiologic relations to arthritis and nephritis. Arch. Intern. Med 1912; IX, 484–498 (1912).

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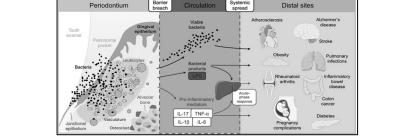
Leaky Gums

- Increased permeability of oral tissues is linked to systemic inflammation.
- Causes include poor dental hygiene, high glycemic-load diet, chronic stress, and microbiome imbalance.
- The oral microbiome:
 - · Maintains a healthy balance of bacteria.
 - Influenced by prebiotic foods (fibers) and probiotics (yogurt), etc.
 - Oral dysbiosis can lead to gum disease, cavities, and systemic issues.

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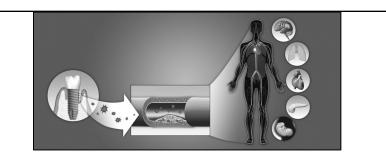
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- Severe periodontitis affects 743 million people worldwide.
- Bacteria enter the bloodstream and can translocate to extraoral tissue, such as the lung, heart, gut, placenta, brain, and joints. It can alter the gut microbiome, disrupt the intestinal barrier, and trigger a systemic inflammatory response.

From: Konkel JE, et al. Distal Consequences of Oral Inflammation Front. Immunol 2019; https://doi.org/10.3389/fimmu.2019.01403

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Clinical and experimental studies highlight the strong association between periodontitis and systemic disease, in particular cardiovascular disease, diabetes, lung diseases, and complications of pregnancy. Oral health is a crucial part of holistic wellness.

Isola G. The Impact of Diet, Nutrition, and Nutraceuticals on Oral and Periodontal Health. Nutrients. 2020 Sep 6;12(9):2724

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Inflammation and Disease

- Meta-inflammation, driven by low-grade systemic inflammation, directly contributes to *insulin resistance, metabolic syndrome, and T2DM*, which makes weight gain easier and weight loss more difficult.
- Diseases related to chronic inflammation = 50% of global deaths.
- In the US, BMI ≥ 30 is significantly associated with periodontal disease among people aged 30 to 44 (P <0.001).¹ Studies show similar results in other global populations.²

Lia L, Xia LY, Gao YJ, Dong XH, Gong RG, Xu J. Association heterescer Obesity and periodontitis in US Adults: N14NRIS 2011-2014. Oler Fasts. 2023 Nov 7. doi: 10.1119/100056751.
 Kim OL, et al. Obesity and periodontitis: A systematic review and updated meta-analysis. Finat Endonin/ (Lansance). 2022 Oct 24;13599455.

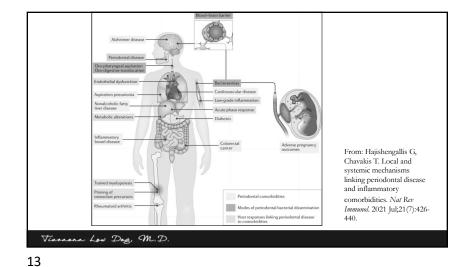
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Dental Plaque

- Organized biofilm of microorganisms attached to the tooth surface or other microorganisms, allowing survival and resistance to host defenses.
- As biofilm matures, dysbiosis occurs, shifting from Gram+ to Gramanaerobic species, biofilm forms under the gingival surface.
- Sugar metabolism by biofilm produces organic acids, lowering pH and demineralizing the tooth surface. Frequent sugar consumption induces dysbiosis of the supragingival microbiota, promoting the development of carious lesions.

Bui FQ, et al. Association between periodontal pathogens and systemic disease. Biomedical Journal 2019; 42(1):27-35

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Periodontal Disease and Alzheimer's Disease

A meta-analysis of 13 studies:

- Eight reported Alzheimer's disease (AD; 291,114 participants)
- Eight reported mild cognitive impairment (MCI, 4,805 participants)
- The risk of AD and MCI in patients with periodontal disease (PD) was significantly higher for AD (OR=1.78) and MCI (OR=1.60) and especially in those with severe PD for AD (OR=4.89, or almost 5 x more likely) and for MCI (OR=2.32).

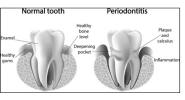
Hu X, et al. Periodontal disease and the risk of Alzheimer's disease and mild cognitive impairment: a systematic review and meta-analysis. Psychogeniatris. 2021 Sep;21(5):813-825

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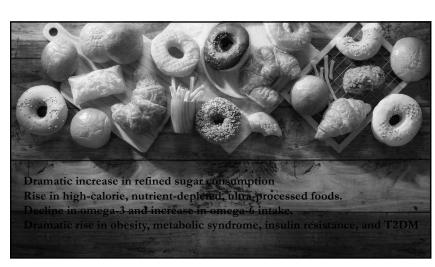
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Diet and Periodontitis

- Diet is a modifiable factor for systemic inflammation, which plays a vital role in periodontitis.
- The research underscores a robust association between **proinflammatory dietary** patterns and the development of **periodontitis.**
- Both micro- and macronutrients modulate inflammatory cascades, influencing inflammatory status.²



Lieske B, et al. *Nutrients.* 2023 Jul 21;15(14):3235 Littlemore, B., et al. *BDJ Team* 2021; 8: 55–65





By 1750 CE, those in Britain consumed ~4 pounds of sugar per year. By 1850 CE, closer to 25 pounds per person per year. By 1950 CE, average consumption in the US was ~120 pounds per year. Today, the average sugar consumption of US adults is ~152 pounds per year.

DiNicolantonio JJ, et al. Prog Cardionasc Dis 2016; 464-72

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Sugar and Dental Caries

- Excessive sugar consumption is the leading cause of dental caries. Sucrose is more cariogenic than fructose/glucose.
- American Academy of Pediatrics:
 - No added sugars to children < 2 years
 - No more than 25 grams (6 tsp) of sugar or added sugars two years and older
- American Heart Association:
- Women limit added sugar intake to 6 tsp/d
- Men limit to 9 tsp/d (37.5 g)
- We consume roughly triple this amount of sugar

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What About Sugar Substitutes?

- Despite GRAS status, sugar substitutes can negatively affect gut microbiota.
- Sucralose disrupts the diversity of gut microbiota, increasing bacterial pro-inflammatory genes.
- In animal/human studies, xylitol and erythritol increase platelet aggregation.
- NIH-funded research: those with the highest erythritol levels (top 25%) twice as likely to have CV events over three years of follow-up as the bottom 25%.

Witkowski, M, *et al. Nature Medicine* 2023 h<u>ttms://doi.org/10.1038/s41591-023-02223-9</u> Witkowski M, et al. *Eur Heart J.* 2024 Jul 12;45(27):2439-2452



Ultra-Processed Foods

- UPF are "snacks, drinks, ready meals, and other products created mostly or entirely from substances extracted from foods or derived from food constituents with *little if any intact* food."¹
- From animals or plants harms the microbiome and drives inflammation.²
- US: 57% of total calories for adults and 67% for children come from UPF. Observational studies show an association between UPF and cancer, heart disease, obesity, depression, and other chronic health problems.

Willett W, et al. Lanor 2019 February 2; 393, (10170): 447-492.
 Snour B, et al. Lanor Gastmentred Highstd 2022 Dec;7(12):1128-114
 Juul F, et al. Any J Chin Natr 2022; 115(1):211-221.
 Wang L, et al. JAMA 2021; 526(6):519-550

Anti-Inflammatory Dietary Patterns

- Mediterranean and DASH diets are associated with reduced inflammatory markers, periodontopathogenic bacteria^{1,} and better oral outcomes.²
- · Both diets have high consumption of plantbased foods (fruits, vegetables, legumes, nuts, whole grain products) and dietary fiber and *limited consumption* of sweets and highly processed foods, including processed red meats.

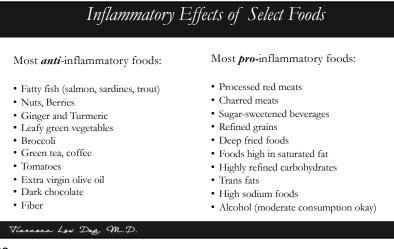
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 Laiola M et al. A Mediterranean Diet Intervention Reduces the Levels of Salivary Periodontopathogenic Bacteria in Overweight and Obese Subjects. *Appl. Environ. Microbiol.* 2020;86:e00777-20 2. Altun E, et al. Association between Dietary Patterns and Periodontitis-A Cross-Sectional Study. Nutrients. 2021;13:4167.



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IM AGINE HEALTHY!	HOME PRODUCTS	COMPANY	BLOG CONTACT US LOG IN Search Q
	DEMAN	⊮" ND™	DII on Demand \$45.00 Designed for those who want to monitor their DII [®] score on an on-going basis, DII-on-Demand TM is an in-depth, online food frequency questionnaire that generates a DII [®] score calculated by CH1 schrists. Typically, DII-on-Demand TM is repeated monthly, though sometimes, twice a month, based on your doctor's
			or nutritionist's recommendations Users receive a personalized report that includes their Dil [®] score, dietary recommendations and next steps for reducing the inflammatory potential of one's diet.
www.imaginehealthy.org			DII-on-Demand [™] Report \$45:00 USD Each New User? Create Account
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Nutrient(s)	Type of Study	Methodology	Follow- Up Period	Clinical Outcomes	Subclinical Outcomes	Author(s) and Year	
Mediterranean diet	RCT	42 patients with gingivitis, were divided into a groups. Test group had to adhere to Mediterranean diet (DM) for 6 weeks and control group did not have to change their diet. Gingival parameters were assessed at baseline, week 2 (beginning of the MD intervention), and week 8.	8 weeks	Test group showed better results in gingival inflammatory parameters (Gi, BOP) after treatment, No differences in dental bacterial plaque scores between test and control groups.	Test group achieved weight loss and waist compliance after treatment.	Bartha et al., 2021 [<u>]3</u>]	PPD= probing pocket depth
Low in carbohydrates, rich in omega-3 fatty acids, vitamins, C and D, antioxidants, and fiber	Pilot RCT	15 patients with gingivitis were divided into 2 groups. The test group was started on a dist low in carbohydrates, high in omega-5 farty acids, and rich in vitamins C and D, antioxidans, and ther for a 4 weeks, Periodonial parameters were measured after 1 and 2 weeks, followed by a 2-week transition period, and then measured weeks for 4 weeks	4 weeks	Test group showed significant improvement in Gi, BOP, PI, PPD, and CAL parameters.	N/A	J. P. Woelber et al., 2016 [14]	CAL =clinical attachment loss BoP =bleeding on probing
Low in carbohydrates, rich in omega-3 fatty acids, vitamins: C and D, antioxidants, and fiber	RCT	54 patients with gingivitis were divided into 2 groups, The test group was started on a dist low in carbohydones, high in omega-5 fatty acids, and rich in vitamins C and D, antioxidants, and fiber for 4 weeks. Periodontal parameters were measured once a week for 4 weeks, followed by a 2- week transition period, and then measured weekly for 4 weeks.	4 weeks	Test group showed significant improvement in BOP and GL	N/A	Sava Sunari Rajaram et al., 2021 [j≦]	Table From: Papathanasiou E, et al. Anti-Inflammatory Benefits of Food Ingredients in Periodontal Diseases. <i>Pathogens</i> . 2023 Mar 27;12(4):520.

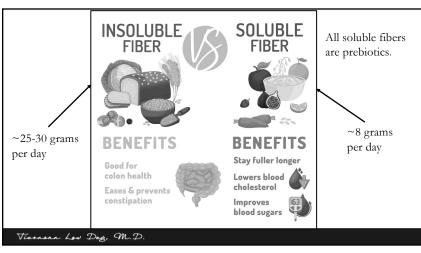
High-fiber and low-fat diet	Pilot Study	47 volunteers were included in the study. Subjects received a high-fiber, low-fat test meal 3x/day for 8 weeks, followed by a regular diet for 24 weeks. Periodontal parameters were evaluated at the beginning and end of treatment.	8 weeks	The high-fiber, low-fat diet effectively improved PPD, CAL, and BoP in patients after treatment.	After treatment, there was improvement in metabolic profiles (body weight, HbA1c, and high-sensitivity C-reactive protein levels).	Keiko Kondo et al., 2014 [16]	
Nordic diet	Cohort	2187 healthy Finns eating a Nortici diet were divided into two age groups and then into two eral hygiene groups (good and poor oral hygiene). Periodonial parameters were used as outcome variables. Dietary data were collected using a validated food frequency questionnaire.	6 weeks	Nordic diet provides evidence that it is associated with less gingival bleeding and reduced PPD in patients with poor oral hygiene.	N/A	Jauhiainen L et al., 2016 [<u>17</u>]	
Nitrate	RCT	44 patients with gingivitis were divided into 2 groups. Test group received 100 m.L.O daily for 14 days, and the control group received placebo. Periodontal parameters of salivary mirate were evaluated before and after treatment.	2 weeks	Test group showed better results in GI on day 14.	Test group showed higher levels of salivary nitrate.	Jockel- Schneider Y et al., 2016 [<u>18]</u>	
Nitrate	RCT	37 patients with gingivitis and reduced periodontitum were divided into 2 groups. Test group received lettuce juice (200 mg of nitrate) daily for 14 days and test group received placebo. Microbial samples, salt collection, and assessment of gingival inflammation were analyzed before and after treatment.	2 weeks	Test group showed reduction in gingival inflammation after treatment.	Test group showed compositional changes within the subgingival microbiome after treatment.	Yvonne Jockel- Schneider et al., 2020 [12]	Table From: Papathanasiou E, et al. Anti-Inflammatory Benefits of Food Ingredients in Periodonial Diseases. Pathogens: 2023 Mar 27;12(4):520.

Fiber and Oral Health: Systematic Review

- 5 RCT, one sequential feeding trial; fiber-rich diet for 4-8 weeks
- Fiber-rich dietary intervention significantly reduced:
 - Clinical Attachment Loss/Level by 0.48 mm/tooth (p < 0.001)
 - Bleeding On Probing by 27.57% sites/tooth (p = 0.02)
 - Periodontal Inflamed Surface Area by 173.88 mm^2 (p = 0.003),
 - **Plaque Index** by 0.02 (*p* = 0.04)
 - Gingival Index by 0.41 (p= 0.002).
 - Probing Depth -0.17 mm/tooth (p = 0.09 non-significant)

Swarnamali H, et al. Role of Dietary Fibre in Managing Periodontal Diseases-A Systematic Review and Meta-Analysis of Human Intervention Studies. Natrinut. 2023 Sep 18:15(18):4034.





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Fresh & Drie	d Fruits	Serving	Soluble Fiber (g)	Insoluble Fiber (g)	Total Fiber (g)
Apple with s	kin	1 medium	4.2	1.5	5.7
Apricots, dri	ed	4 medium	1.8	1.7	3.5
Banana		1 medium	2.1	0.7	2.8
Blackberries		½ cup	3.1	0.7	3.8
Figs, dried		3 medium	3.0	2.3	5.3
Grapefruit		½ of large	2.4	0.7	3.1
Kiwi		1 large	2.4	0.8	3.2
Orange		1 medium	2.1	1.3	3.4
Pear		1 medium	0.8	3.2	4.0
Plums		2 medium	1.2	1.0	2.2
Prunes		4 medium	1.3	1.8	3.1
Raspberries		½ cup	0.9	2.3	3.2
Strawberries		1 cup	1.8	2.6	4.4
Nuts, Seeds	& Beans				
Almonds, ray	N	1 ounce	0.7	3.5	4.2
Black beans,	cooked	½ cup	3.8	3.1	6.9
Flaxseeds		2 tbsp.	2.7	2.1	4.8
Garbanzo be	ans, cooked	½ cup	1.2	2.8	4.0
Kidney bean	s, cooked	½ cup	2.9	2.9	5.8
Lentils, cook	ed	½ cup	2.8	3.8	6.6
Peanuts, dry	roasted	1 ounce	1.1	1.2	2.3
Pinto beans,	cooked	½ cup	5.5	1.9	7.4
Psyllium see	ds	2 tbsp.	7.1	0.9	8.0
Sesame seed	ls	¼ cup	0.7	2.6	3.3
Split peas, co	ooked	½ cup	1.1	2.4	3.5
Sunflower se	eds	¼ cup	1.1	1.9	3.0
Walnuts		1 ounce	0.6	2.5	3.1

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Vegetables	Serving	Soluble fiber (g)	Insoluble fiber (g)	lotal fiber (g)
Artichoke, cooked	1 medium	4.7	1.8	6.5
Asparagus, cooked	½ cup	1.7	1.1	2.8
Broccoli, raw	½ cup	1.3	1.4	2.7
Brussel Sprouts, cooked	1 cup	1.7	1.9	3.6
Carrot, raw	1 medium	1.1	1.5	2.6
Green peas, cooked	½ cup	3.2	1.2	4.4
Green beans, cooked	½ cup	0.8	1.2	2.0
Kale, cooked	1 cup	2.1	5.1	7.2
Lima beans, cooked	½ cup	2.1	2.2	4.3
Potato with skin	1 medium	2.4	2.4	4.8
Soybeans (edamame)	½ cup	2.7	2.2	4.9
Squash, summer, cooked	½ cup	1.3	1.2	2.5
Sweet potato, peeled	1 medium	2.7	2.2	4.9
Tomato with skin	1 medium	0.3	1.0	1.3
Zucchini, cooked	½ cup	1.4	1.2	2.6
Whole Grains				
Barley, cooked	½ cup	3.3	0.9	4.2
Brown rice, cooked	½ cup	1.3	0.1	1.4
Oat bran, cooked	¾ cup	2.2	1.8	4.0
Oatmeal, cooked	1 cup	2.4	1.6	4.0
Popcorn, air-popped	3 cups	3.2	0.4	3.6
Dumpernickel bread	1 elica	15	1 2	17
Rye bread	1 slice	1.9	0.8	2.7
Wheat bran	½ cup	11.3	1.0	12.3
Wheat germ	3 tbsp.	3.2	0.7	3.9
Wholegrain bread	1 slice	2.8	0.1	2.9
Whole wheat bread	1 slice	1.6	0.3	1.9
Wholegrain pasta	1 cup	4.1	2.2	6.3

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Dietary Protein

- Essential for collagen production, a key structural component of gingival tissue, periodontal ligament, and alveolar bone.
- Supports the synthesis of antibodies, cytokines, and other immune molecules that protect against oral infections.
- Enough, but NOT too much!



Dietary Protein

- Saliva contains proteins that contribute to antimicrobial activity, buffer acid, and protect teeth from demineralization.
- Protein-rich diets ensure the production of high-quality saliva, enhancing its protective functions.
- Arginine-containing proteins increase oral pH, reducing the risk of dental caries (e.g., nuts, eggs, chicken, pumpkin seeds, soybeans, dairy, chickpeas)



Dietary Protein and Oral Health

- Elders are particularly at risk for low protein diet, which increases the risk of sarcopenia and poor bone health.
- Low protein intake in elders is associated with poor oral health, dental caries, enamel hypoplasia, and salivary gland atrophy.

Jayasinghe TN, et al. Nutrients. 2022 Oct 25;14(21):4478.

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Protein Fast Facts ity to digest, • Meat cuts with round, chuck, or

loin in the name are usually lean.

• Beware of buying ground chicken

or turkey unless the package says

chicken breast. The meat is likely

100% ground turkey breast or

• Eat low-mercury fatty fish 2

ground with skin/fat.

times per week.

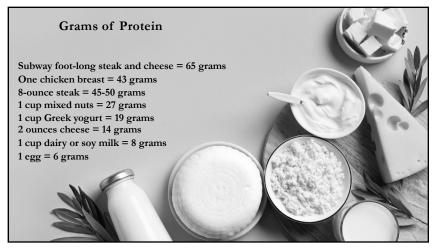
- Maximize the ability to **digest**, **absorb**, **and use protein** by distributing it **evenly throughout meals and snacks**.
- A high-protein snack within **one hour of exercise is best** - muscles are sensitive to nutrients they use to repair and grow.
- Beans, peas, quinoa, and lentils are rich in protein, fiber, vitamins, and *minerals*.

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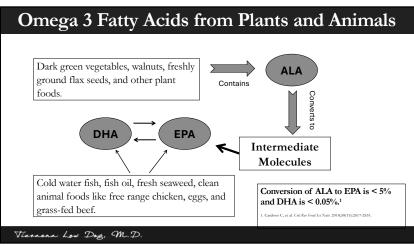
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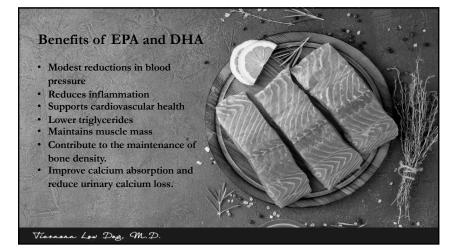
How Much Protein Do You Need? ~0.8 g/kg ideal body weight adults (Multiply weight in lb. x 0.36) 150 pounds = 55 g/d 180 pounds = 65 g/d 10–1.2 g/kg for those over age 60* 150 pounds = 81–98 grams 180 pounds = 81–98 grams 12–1.6 g/kg competitive athletes *Caution with advanced kidney disease.

Protein Choices• Protein intake should be
of high biological
value and from healthy
dietary choices:
legumes, nuts, eggs,
fish, lean meats, dairy,
etc.Image: Constraint of the state of











Preg 1 serv Eat 2 from	ing? As a guide, uso nancy and breastfee ing is 4 ounces to 3 servings a weel the "Best Choices" I serving from the "Go	ding: k ist	r hand.	1 ounce at 2 ounces a 3 ounces a 4 ounces a	e, a serving is about: age 1 to 3 at age 4 to 7 at age 8 to 10 at age 11	e "Best Choices" list.
Best Choice	5		Good C	hoices		
Anchovy Atlantic croaker Atlantic mackerel Black sea bass Butterfish Catfish Clam Cod	Herring Lobster, American and spiny Mullet Oyster Pacific chub mackerel Perch, freshwater and ocean	Scallop Shad Shrimp Skate Smelt Sole Squid Tilapia	Bluefish Buffalofish Carp Chilean sea Patagoniar Grouper Halibut Mahi mahi/	ı bass/	Monkfish Rockfish Sablefish Sheepshead Snapper Spanish mackerel Striped bass (ocean)	Tilefish (Atlantic Ocean) Tuna, albacore/ white tuna, canned and fresh/frozen Tuna, yellowfin Weakfish/seatrout White croaker/ Pacific croaker
Crab Crawfish	Pickerel Plaice	Trout, freshwater Tuna, canned light	Choice	s to Avoi		EVELS
Flounder Haddock Hake	Pollock Salmon Sardine	(includes skipjack) Whitefish Whiting	King macke Marlin Orange rou		Shark Swordfish	Tilefish (Gulf of Mexico) Tuna, bigeye
fish. If there is no a		erving and no other fis	sh that week.	Some fish ca		ou can safely eat those nds, such as larger carp,
				gov/fishadvic .gov/fishadvic		

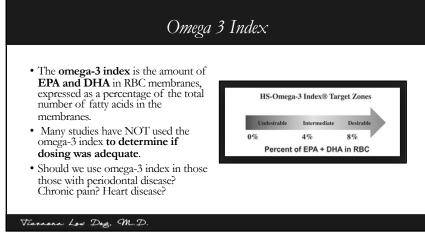
Omega 3 Fatty Acids and Periodontitis

- Meta-analysis 8 RCTs: Significant overall reduction in probing pocket depth of 0.42 mm and clinical attachment loss gain of 0.58 mm associated with the use of omega-3 fatty acids.
- "In individuals with periodontitis, omega-3 fatty acid supplementation as an *adjunct* to non-surgical periodontal treatment can provide additional benefits in CAL gain and PPD reduction, compared with nonsurgical periodontal treatment alone."

Castro Dos Santos NC, et al. Does the use of omega-3 fatty acids as an adjunct to non-surgical periodontal therapy provide additional benefits in the treatment of periodontitis? A systematic review and meta-analysis. J Periodontal Res. 2022 Jun;57(3):435-447."

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Global Omega-3 Status Map shows low levels for most of the world



Update on Omega 3 Index Status

- · The US (excluding Alaska), Canada, Germany, and Italy have seen improvement and are now in the "low category."
- Spain has moved from the low to the "moderate" category.
- South Korea, Japan, and Alaska's average Ω -3i were >8%.
- Total fish and shellfish intake:
 - US 4.38 kg/capita/y
 - Germany 14.1 kg/capita/y
 - Japan (45.5 kg/capita/y)
 - South Korea 55.0 kg/capita/y

Schuchardt JP, et al. Red blood cell fatty acid patterns from 7 countries: Focus on the Omega-3 index. Prostaglandins Laukot Essent Fatty Acids: 2022 Apr;179:102418.

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Looking at Labels

upplement facts

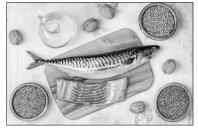
ring Size: 2 Veggie Softgels

vings Per Container: 30

Omega 3 Supplement Forms

- The lowest effective dose to raise Ω -3i: >1,000 mg/d DHA + EPA for 12 weeks.1
- Triglyceride and ethyl-ester forms more bioavailable and effective.¹
- Microalgal oil supplementation increased Ω -3i levels in all studies.²
- High-dose flaxseed or echium seed oil: no change in Ω -3i.²

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Supplement Facts

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Serving Size 2 Soft Gels



400 mg Vegan form EPA/DHA 2:1

%Dail

Value

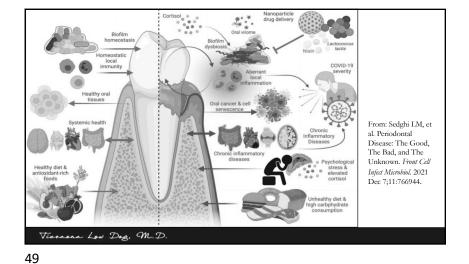
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2 g 3%**

2000 mg †

1300 mg †

800 mg †



Christell Guide to Products Available in USA
 Activities
 Superior Constraints and available in USA
 Activities
 Superior Constraints
 Superior Constan

- Probiotics utilized in periodontal therapy have included tablets containing live cultures of *Lactobacillus reuteri, Streptococcus salivarius,* probiotic drinks containing *Lactobacillus casei,* and chewing gum containing *Lactobacillus reuteri.*
- These can be used to help restore the oral microbiome.

Nguyen T, et al. Probiotics, including nisin-based probiotics, improve clinical and microbial outcomes relevant to oral and systemic diseases. Periodontol 2020 Feb;82(1):173-185.

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Nisin

- Nisin is an antimicrobial peptide produced by Lactococcus lactis.
- It is active against both Gram(+) and Gram(-) bacteria, including Streptococcus aureus, Listeria monocytogenes, Fusobacterium nucleatum, Porphyromonas gingivalis, and Treponema denticola.
- *L. lactis* can prevent oral biofilm formation and disrupt 24-h and 48-h pre-formed biofilms.

Rachaic A, et al. Modulation of pathogenic oral biofilms towards health with rissin probiotic. J Out Microbiol. 2020 Aug 24;12(1):1809302.

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The Enterosalivary Circuit

- Dietary nitrate (e.g., spinach, arugula, celery, beets) is absorbed into the bloodstream from the stomach and small intestine.
- ~25% is concentrated in the salivary glands and secreted into the saliva.
- Bacteria in the oral cavity reduce nitrate to nitrite. The saliva containing nitrite is swallowed. In the stomach's acidic environment, nitrite is reduced to nitric oxide.
- Nitrite and NO are absorbed systemically, contributing to vasodilation, improved blood flow, lowered blood pressure, and immune defense.
- Studies show PPIs and antiseptic mouthwash blunt these effects. Monengro MF; et al. Blood Pressure Lowering Effect of Orally Ingested Nitrite Is Abdeled by a Proton Pump Inhibitor. *Hypermains*. 2017 Jarc/9(1):23-31.

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Nitrate and Oral Health

- The nitrate-reducing bacteria *Rothia* and *Neisseria* are consistently found at higher levels in individuals free of oral disease (vs. individuals with caries, periodontitis, and/or halitosis) and increase when nitrate is consumed in clinical studies.
- Preliminary in vitro and clinical evidence show that **bacteria normally associated with disease**, such as *Veillonella* (caries) and *Prevotella* (periodontal diseases and halitosis), **decrease in the presence of nitrate**.

Rosier BT, Takahashi N, Zaura F, et al. The Importance of Nitrate Reduction for Oral Health. Journal of Dental Research. 2022;101(8):887-89

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Saliva

- Facilitates clearance of dietary carbohydrates and microorganisms from the oral cavity.
- Supplies bacteria with nutrients through enzymatic breakdown of dietary starch, proteins, and salivary glycoproteins.
- Antimicrobial activity through numerous proteins and peptides, including mucins, lactoferrin, lysozyme, lactoperoxidase, and SIgA.
- Up to 10⁸ microorganisms/ml of saliva have been found.
- Salivary gland hypofunction causes dysbiosis and increases the risk of oral disease.

Lynge Pedersen AM, Belstrom D. The role of natural salivary defences in maintaining a healthy oral microbiota. J Dent. 2019 Jan;80 Suppl 1:S3-S12

Ginger Mouthwash

- Several studies have found ginger spray or mouthwash can be helpful in patients with xerostomia.
- · A study of 105 patients with diabetes and xerostomia randomized to one of the following groups: ginger (25%) mouthwash, aloe vera mouthwash, or control with normal saline.
- All the mouthwashes were used at 20 cc three times per day.

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			intervent [n (%)]	ion		after intervention [n (%)]				Com- parison	
Symptom	Response	Ginger	Aloe	control	P*	Ginger	Aloe vera	control	P*	P**	
Experience of Mouth dryness during meals	yes	23(65.7)	18(51.4)	23(65.7)	0.368	2 (5.7)	4(11.4)	5(14.3)	0.491	< 0.001	
Difficulty in swallowing	yes	27(77.1)	20(57.1)	21 (60)	0.166	4(11.4)	3(8.6)	7(20)	0.343	< 0.001	
The need of Drinking fluids while swallowing dry food	yes	31(88.6)	35(100)	32(91.4)	0.120	12(34.3)	17(47.6)	27(77.1)	<0.001	< 0.001	
The feeling of decreased saliva in mouth	yes	33(94.3)	29(82.9)	33(94.3)	0.171	6(17.1)	12(34.3)	26(74.3)	< 0.001	< 0.001	
Experience of Mouth dryness after waking up	yes	34(97.1)	30(85.7)	33(94.3)	0.233	8(22.9)	10(28.6)	25(71.4)	< 0.001	< 0.001	
Waking up during night due to thirst	yes	28(80)	29(82.9)	23(65.7)	0.196	5(14.3)	14(40)	9(25.7)	< 0.051	< 0.001	
Chewing gum or using chocolate to diminish dryness in mouth	yes	12(34.3)	10(28.6)	10(28.6)	0.955	3(8.6)	3(8.6)	9(25.7)	<0.061	< 0.001	
Difficulty in tasting foods	yes	12(34.3)	12(34.3)	16(45.7)	0.524	6(17.1)	5(14.3)	11(31.4)	0.168	< 0.001	
Burning sensation in mouth	yes	14(40)	10(28.6)	17(48.6)	0.228	6(17.1)	2(5.7)	11(31.4)	0.020	< 0.001	
Feeling dry mouth during travel	yes	34(97.1)	35(100)	35(100)	0.364	13(37.1)	25(71.4)	31(88.6)	< 0.001	< 0.001	

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Reference

Chandrahas et al.

Ajmera et al.

Rezaei et al.

Vangipuram et al.

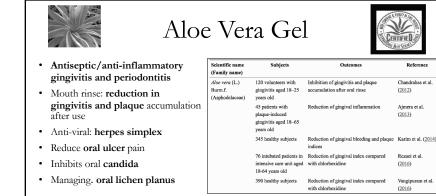
(2016)

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Outcomes



Sujatha G, et al. Aloc vera in dentistry. J Clin Diagn Res 2014; 8(10): Z101–Z102 Ali S. Wahbi W. The officient of the second statement of the second

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Glutamine as an Immunomodulatory Agent

- Immune cells depend on glutamine to survive, proliferate, and function. During catabolic/hypercatabolic circumstances, demand for glutamine increases dramatically; low glutamine may severely impair immune function. (Preop, post-op, elite athletes, trauma)
- A systematic review of eight studies (six RCTs) recommended glutamine oral care to mitigate oral mucositis.





ory agent

Glutamine C5H10N2O3

Mahendran VJ, et al. Advances in the use of anti-inflammatory to manage chemotherapy-induced oral and gastrointestinal mu *Curr Pharm Des.* 2018;24(14):1518–32.



▶ Med Sci Monit. 2024 Feb 22;30:e942585-1-e942585-10. doi: 10.12659/MSM.942585

Effectiveness of Glutamine Oral Care in Reducing Oral Mucositis and Improving Oral Health After Neurosurgery: A Randomized Controlled Trial with Microbiome Analysis

Yan Gao^{1,D,E}, Hong Yang^{1,B}, Xiaohong Zhang^{1,C}, Ying Ma^{1,D,F}, Ling Wang^{1,A,E,®}

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 PMCID: PMC10898191 PMID: <u>38384124</u>

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Glutamine and Oral Care

- In a double-blind study, 300 patients undergoing neurosurgery without pneumonia or oral infection were randomized into three groups:
 - The **control group** (n=100) received oral care with routine oral nursing methods with saline
 - The **experimental group** (n=100) received oral care with 5% glutamine.
 - The saline chlorhexidine combo (n=100) was positive control.
- Oral health complications, such as local debris, oral mucositis, halitosis rates, oral flora, and pneumonia incidence, were included in the analysis.

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Checklis	of oral care procedures.	
Number	Procedure	Status (√
1	Patients were asked to sit fully upright or in an elevated side-lying position to prevent aspiration during mouth cleaning	
2	Take off the denture if there is one, spread the treatment towel under the jaw, and wet the mouth	
3	Rinse the teeth and mouth with saline 0.9% solution for 150–180s, and check the condition of the oral cavity	
4	Brush the teeth with a suction toothbrush (or soft toothbrush) with toothpaste for 180s, ensuring all four quadrants were cleaned	
5	Dip swab into a standard 0.9% saline oral rinse	
6	Clean the mouth with swab for 150-180s	
7	Clean the hard palate, tongue and the bottom of the tongue with swab for 150-180s	
8	Wipe the mouth and lips for 20-30s	
9	Inquire about the patient's feelings about the whole care process, and check the oral condition with the flashlight (whether it is clean, whether there are any residues left, whether there is bleeding, ulcer, etc.)	

Results											
Comparison of	oral disorders.										
Group	Case number	Local debris cases	Oral mucositis cases	Halitosis cases	Dryness cases						
Control	100	38 (38.00)	52 (52.00)	46 (46.00)	24 (24.00)						
Glutamine	100	14 (14.00)	23 (23.00)	23 (23.00)	18 (18.00)						
Chlorhexidine	100	12 (12.00)	22 (22.00)	21 (21.00)	19 (19.00)						
χ²		24.95	26.54	18.38	1.28						
Р		< 0.001	< 0.001	< 0.001	0.528						

Nutrient	Dietary Source(s)	etary Source(s) Importance in Periodontal Reported References Heath Improvement in PD and CAL (Mean mm, SD)							
Vitamin A	Cod liver oil, carrots, capsicum, liver, sweet potato, broccoli, leafy vegetables	Not clear. Research indicates insignificant improvement in periodontal health upon supplementation.	PD: 0.52 ± 0.03 CAL: n.d.	[33,65]	Vitamin C	Citrus fruits, vegetables, liver	Gingival bleeding and inflammation are hallmarks of scurvy. Supplementation may improve outcomes of periodontal therapy.	PD: 0.58 ± 0.14 CAL: n.d.	[66]
B- vitamins	B1-Liver, oats, pork, potatoes, eggs B2-Bananas, dairy, green beans B3-Eggs, fish, meat, mushrooms, nuts B2-Avocados, meat,	Supplementation may accelerate post-surgical healing.	PD: 1.57 ± 0.34 CAL: 0.41 ± 0.12	[56]	Vitamin D	Fish eggs, mushrooms, liver, milk	Deficiency may lead to delayed post-surgical healing. Local application may accelerate post-surgical healing/osseointegration	PD: 1.35 (SD n.d.) CAL: 1.4 (SD n.d.)	[34,56,57,67]
	B5-Avocados, meat, broccoli B6-Meat, vegetables, nuts, banana				Vitamin E	poultry, meat, fish, nuts, seeds and cereals	Impaired gingival wound healing	PD: 0.39 ± 0.18 CAL: n.d.	[33,68,69,70]
	B ₇ —Raw egg, liver, leafy vegetables, peanuts B ₉ —Cereals, leafy vegetables				Vitamin K	Green vegetables, egg yolk	Deficiency may lead to gingival bleeding. No known effects on periodontal therapy if supplementation used as an	n.d.	[71,72,73]

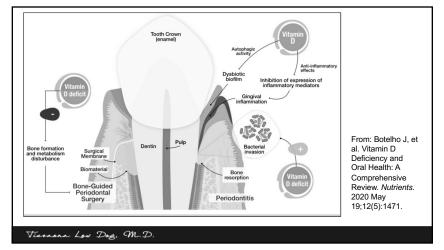
From: Najeeb S, The Role of Nutrition in Periodontal Health: An Update. Nutrients. 2016 Aug 30;8(9):530.

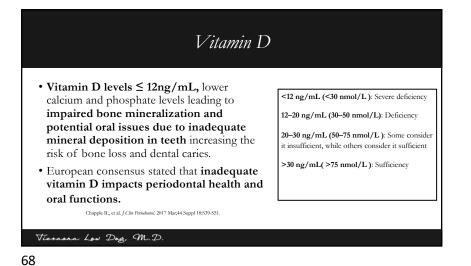
adjunct.

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B12-Animal product

	VITAMIN D		
Function	Promotes calcium absorption and maintains adequate serum calcium and phosphate concentrations. Involved in modulation of cell growth, neuromuscular, hormone, and immune function; and glucose metabolism. Involved in regulation of hundreds of genes.		
Clinical Use	Bone pain, muscle weakness, osteomalacia, high risk for falls/fractures Prediabetes		
Deficiency Signs	Musculoskeletal pain, muscle twitches, spasms, poor gait Poor immune function		
Status Indicator	25(OH)D: IOM: sufficiency 20 ng/mL, severe deficiency <12 ng/mL Endocrine Society: sufficiency 30 ng/mL higher, deficiency <20 ng/mL		
Typical Dosing	400 IU daily for infants less than one year, exclusively or partially breastfed 600 IU daily for those ages 1 to 70 years of age 800 IU daily for all adults up >70 years Tolerable upper limit: 4-8 years 3000 IU/d those 9 years and older: 4000 IU/d Deficiency: 4000-5000 IU/d (100-125 μg) or 50,000 IU/wk for 2-3 months, recheek		
Forms	D2 (ergocalciferol) – from mushrooms D3 (cholecalciferol) – from lanolin or lichen, superior form		
auhan K, Shahrokhi M, Huecker	MR. StatPearls Publishine: 2024 Jan. and Berger MM, et al. ESPEN micronutrient guideline. Clin Nutr. 2022 Jun;41(6):1357-1424.		





Vitamin D Supplementation

Endocrine Society's new guidelines highlight the following groups:

- **1-18 years:** prevent nutritional **rickets** and potentially lower risk of RTI* (estimated weighted average ~ **1200 IU/d***)
- Pregnancy: potential to lower the risk of preeclampsia, intrauterine mortality, preterm birth, small for gestational age birth, and neonatal mortality (~2500 IU/d).
- Adults: high-risk prediabetes to reduce risk of progression to diabetes.
- 75 years and older: potential to lower the risk of mortality.

Endocrine Society Clinical Practice Guidelines. Vitamin D for the Prevention of Disease Guideline Resources. https://www.endocrine.org/clinical-practice-puidelines/vitamin-d-for-prevention-of-disease_Accessed November 4, 2024

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Uwitonze AM, et al. Zine Adequacy Is Essential for the Maintenance of Optimal Oral Health. Nationa. 2020 Mar 30;12(4):049. Monaffer B, et al. The Effectivenesss of Zine Sapplementation in Taste Disorder Treatment: A Systematic Review and Meta-Analysis of Bandomized Controlled Trials. J Natr Metak. 2023 Mar 8:2021-27:102.

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Zinc

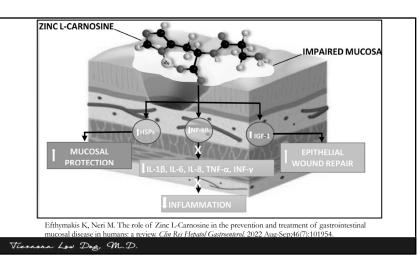
- Found in saliva, dental plaque, and dental hard tissues.
- Supplementation is beneficial in gingivitis, periodontitis, and halitosis, while deficiency is associated with poor oral and periodontal health.¹
- Maintains periodontal health due to immunological effect on oral soft tissues.
- Systematic review: Supplementation is an effective treatment for taste disorders in patients with zinc deficiency or idiopathic taste disorders.²

Zinc L-Carnosine

- Zinc provides anti-inflammatory and wound-healing benefits, while carnosine has cytoprotective effects. Together, they exert more substantial mucosal healing properties than zinc alone.
- It adheres to ulcerated and inflamed oral or gastric mucosa, protecting against further damage while promoting healing.¹
- Evidence supports the safety and efficacy of maintaining, preventing, and treating the mucosal lining, particularly for oral mucositis (approved in Japan for treatment of gastric ulcers).²

 Efthymakis K, Neri M. The role of Zinc L-Carnosine in preventing and treating gastrointestinal mucosal disease in humans: a review. Chr Ru Hupatol Gastroentoril. 2022 Aug. Sep46(7):10195-2. Hewings S, Kalman D. A Review of Zinc-L-Carnosine and Its Positive Effects on Oral Macositis, Taste Disorders, and Gastrointestinal Disorders. National. 2020 Feb 29;12(3):665.





Dose and Safety Profile

- Recommend Dose: 75-150 mg/d zinc L-carnosine (~16-30 mg elemental zinc).
- Duration: 8–12 weeks, depending on the condition being treated.
- · Safety: Generally well-tolerated with minimal adverse effects.
- Toxicity: There is a low risk at the recommended dose. Excessive elemental zinc intake (>40 mg/d) may cause copper deficiency.
- Side Effects:
- Mild gastrointestinal discomfort (rare).
- Zinc-related nausea or metallic taste (occasionally reported).

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Labels

- Recommend Dose: 75-150 mg/d zinc L-carnosine (~16-30 mg elemental zinc).
- Duration: usually 8-12 weeks, depending on the condition being treated.
- Safety: Generally well-tolerated with minimal adverse effects.
- Toxicity: Low risk at recommended dose. Excessive elemental zinc intake (>40 mg/day) may cause copper deficiency or immune dysregulation.
- Side Effects:
- Mild gastrointestinal discomfort (rare).
- Zinc-related nausea or metallic taste (occasionally reported).

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Supplement facts			Supplement	
Serving Size: 1 Capsule				
Servings Per Container: 90			Serving Size 1 Tablet Servings Per Container 60	
	Amount Per Serving	%Daily Value	Amount Per Serving	
Zinc (from Zinc-L-Carnosine)	12 mg	109%	Zinc 17 mg (from 75 mg zinc-carnosine)	
Zinc-L-Carnosine	75 mg	t		
L-Carnosine (from Zinc-L-Carnosine)	43 mg	+		
†Daily Value not established.			Other Ingredients: Microcrystalline of acid (vegetable), and silica.	

Amount Per Serving % Daily Value Zinc 17 mg 155% (from 75 mg zinc-carnosine) 155% Other Ingredients: Microcrystalline cellulose, stearic	Serving Size 1 Tablet Servings Per Container 60	
(from 75 mg zinc-carnosine)	Amount Per Serving	% Daily Value
Other Ingredients: Microcrystalline cellulose stearic		
acid (vegetable), and silica.	Other Ingredients: Microcrystall acid (vegetable), and silica.	ine cellulose, stearic

Resources · Micronutrient Information https://lpi.oregonstate.edu/mic https://ods.od.nih.gov https://www.who.int/health-topics/micronutrients#tab=tab_1 · Micronutrient Calculator https://www.nal.usda.gov/human-nutrition-and-food-safety https://www.osteoporosis.foundation/educational-hub/topic/calciun · Identifying Drug-Nutrient Interactions: https://lpi.oregonstate.edu/mic/drug-nutrient-interactions //www.drugs.com/drug-interactions/multiv niludoaton one (day https://medlineplus.gov/druginformation.html https://naturalmedicines.therapeuticresearch.com (paid subscription) Viernona Low Dog, M.D. 76