

Patient Information	Name:			
Date of Birth:	05/17/1962	Gender:	F	Lab ID:
Date Received:		Date Collected:	03/01/2022	Date Reported:
Physician:	6 D P S O H 3 K \ V L F L D Q		Clinic ID:	10804

Lab Director
Harold Alvarez, M.D.

Genomic Recommendations:

Gene	Address Need For	Nutrient Applications FOOD FIRST	RDA (amount per day according to age)	Consider Supplementation With Practitioner Guidance
COMT	Precursors for body to make SAME (cofactor for COMT enzyme): L-methionine Cofactor:magnesium	Food sources of methionine:eggs, fish, turkey, cheese,legumes, nuts/seeds. High doses of bioactive food components, such as tea catechins (green and black tea, EGCG, green coffee bean extract) and quercetin may further inhibit COMT enzyme activity. Intake may need to be altered.	0-6 months-59 (AI) 6-12 months-43 1-3 years-28 4-8 years-22 9-13 years, boys-22 9-13 years, girls-21 14-18 years, boys-21 14-18 years, girls-19 19+ years-19 Pregnancy-25 Lactation-26	IMPORTANT: The level of nutrient intake to optimize methylation status varies from individual to individual Clinical experience suggests an oral dose of 500mg methionine 1-2 times per day**
MTHFR	5-MTHF (5-methyltetrahydrofolate) Cofactors: riboflavin, niacin, magnesium, zinc	Encourage intake of green leafy vegetables, legumes, citrus fruit, beets, whole grains.	folate: 1-3 years-150ug 4-8 years-200ug 9-13 years-300ug 14+ years-400ug	A daily dose of 100-1000ug (.1- 1 mg) is typically used in research studies to achieve clinical benefit.** Additional support using vitamin B2, B6, B12, and betaine may also need to be addressed.** Avoid folic acid in supplements and fortified foods.
MTR	Vitamin B12 (hydroxycobalamin, adenosylcobalamin or methylcobalamin) Cofactors: zinc	Vitamin B12 is naturally found in animal products. Bound to protein in food, it is released by HCl and protease in the stomach. Many medications, including antacids and acid blockers, can deplete vitamin B12.	B12: 1-3 years-0.9ug 4-8 years-1.2ug 9-13 years-1.8ug 14+ years-2.4ug	1000ug typically suggested ** (Note: Homozygous positive COMT-consider hydroxycobalamin and/or adenosylcobalamin, Homozygous negative COMT-consider methylcobalamin Heterozygous positive COMT-methylcobalamin may or may not be tolerated)
MTRR	Vitamin B12 (hydroxycobalamin, adenosylcobalamin or methylcobalamin) Cofactors: riboflavin,niacin	Vitamin B12 is naturally found in animal products. Bound to protein in food, it is released by HCl and protease in the stomach. Many medications, including antacids and acid blockers, can deplete vitamin B12.	B12: 1-3 years-0.9ug 4-8 years-1.2ug 9-13 years-1.8ug 14+ years-2.4ug	1000ug typically suggested ** (Note: Homozygous positive COMT-consider hydroxycobalamin and/or adenosylcobalamin, Homozygous negative COMT-consider methylcobalamin Heterozygous positive COMT-methylcobalamin may or may not be tolerated)

***Limitations of the Recommended Dietary Allowances** The RDA is defined by The Food and Nutrition Board of the Institute of Medicine as "the average daily dietary nutrient intake level sufficient to meet the nutrient requirement of nearly all (97 to 98 percent) healthy individuals in a particular life stage and gender group." This does not mean that additional nutrients provided via supplementation would not be beneficial. The RDAs are not meant to apply to those managing inherited metabolic disorders, medical conditions, or those using nutrient depleting medications. It is generally well accepted by nutrition professionals, that higher levels of nutrient intake can help prevent chronic disease and promote optimal health.

****Consult with ordering health care practitioner to assess need for supplementation and proper dosage. Therapeutic dose to be determined by ordering health care provider. (the level of nutrient intake to optimize methylation status varies from individual to individual)**

Patient Information

Name: PATIENT II, PRETEND

Date of Birth: 05/17/1962

Gender: F

Lab ID:

68220

Date Received: 03/02/2022

Date Collected: 03/01/2022

Date Reported:

03/07/2022

Physician: Sample Physician

Clinic ID:

10804

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FOOD and LIFESTYLE FIRST

An individual's nutrient status depends on many factors. Digestion, absorption, and assimilation impacts the availability of nutrients supporting methylation, so issues potentially interfering with that availability need to be addressed. Further laboratory assessment may be indicated.

For food and lifestyle based support of methylation:

- Address GI function, intestinal permeability, dysbiosis, and food sensitivities. Avoid offending foods and ingredients.
- Consume a variety of organic, whole, colorful plant foods providing fiber, anti-inflammatory and anti-oxidant benefit. Include omega 3 fatty acids.
- Consume enough protein from lean- antibiotic/hormone free animal sources and/or plant sources- legumes, nuts/seeds.
- Manage weight and regulate blood glucose.
- Hydrate well with filtered water. Overall fluid need = 1 ounce/kg body weight (~½ body weight in fluid ounces, unless fluid restriction prescribed by physician)
- Avoid sugar, refined/fortified grains, conventionally raised animal products, trans fats, charbroiled foods (avoid grilling and deep frying)
- Avoid air pollutants, pesticides, bisphenol A, phthalates, automobile fumes, jet fuel, benzene, heavy metals, plastic food/beverage containers. Avoid high mercury fish- tuna, shark, swordfish, King mackerel.
- Avoid excessive alcohol consumption.
- Don't smoke.
- Work with a nutrition expert who can tailor the eating pattern to meet individual requirements.
- Lead an active lifestyle. Adopt a moderate exercise routine. Consult an exercise specialist to individualize routine. Avoid over-training.
- Reduce and manage stress. Consider meditation, yoga, prayer, positive thinking, acupuncture, social interaction, journaling. Get adequate sleep.

Dietary sources of key methylation nutrients

Folate - leafy greens- spinach, turnip greens, mustard greens, collard greens, legumes- mung beans, chickpeas, pinto beans, great northern beans, lentils, black beans, fava beans, kidney beans, soybeans, navy beans, pinto beans, black eye peas, split peas, peanuts, leeks, asparagus, broccoli, Brussels sprouts, avocado, citrus fruit, beets, spearmint, rosemary, daikon radishes, basil, cilantro (coriander leaf), marjoram, oregano, sage, tarragon, thyme, peanuts, sunflower seeds, wakame seaweed, quinoa, kelp seaweed, bay leaf, parsley, shitake mushrooms, dill, okra, egg, artichokes

Riboflavin - spirulina, egg, paprika, chives, cilantro, spearmint, tarragon, shiitake mushrooms, parsley, almonds, fish roe, cayenne pepper, chili powder, soybeans, game meat, daikon radish, chervil, goat cheese, mackerel, brie cheese, sesame, liver-lamb, beef, chicken, duck, goose

Niacin - peanuts, sunflower seeds, chicken, shiitake mushrooms, sesame seeds, salmon, spirulina, pork cilantro, mackerel, parsley, beef, game meats, sun-dried tomatoes, tarragon, trout, lamb, chili powder, mustard seed, duck, cod, anchovy, liver- beef, lamb, chicken

Magnesium - agar seaweed, herbs, spices, bran, pumpkin seeds, cocoa, flaxseed, Brazil nuts, sunflower seeds, sesame seeds, poppy seeds, almonds, cashews, buckwheat, amaranth, rye, molasses, walnuts, quinoa, great northern beans, mung beans, teff, tofu, chickpeas, oats, daikon radish, bulgur, lamb's quarters, hazelnuts, leeks, black beans, kidney beans, horseradish

Vitamin B12 - meat- beef, chicken, goose, pork, lamb, game meat, fish- mackerel, whitefish, salmon, cod, herring, snapper, trout, crab, clams, lobster, oysters, mussels, eggs, liver (lamb, beef, turkey, duck, goose, chicken) milk and milk products

Zinc - oysters, pumpkin seeds, sesame seeds, chervil, beef, game meats, lamb, poppy seed, shiitake mushroom, cardamom, celery seed, crab, bison, turkey, pork, peanuts, pine nuts, cocoa, thyme, parsley, rice bran, basil, agar seaweed, cashews, lobster, mustard seed, dark rye

Methionine - egg, cod, whitefish, sesame seeds, spirulina, Parmesan cheese, sunflower seeds, Brazil nuts, chicken, beef, lamb, salmon, buffalo, turkey, halibut, anchovy, Romano cheese, game meats, gruyere cheese, goat cheese, goose, duck, snapper, tilapia, mackerel, haddock, lobster, pumpkin seeds, sardine, herring, bison

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References:

- Daijun Zhou et al.: The Polymorphisms in Methylene tetrahydrofolate Reductase, Methionine Synthase, Methionine Synthase Reductase, and the Risk of Colorectal Cancer. *Int. J. Biol. Sci.* 8(6):819-830, 2012
- David Watkins et al.: Hyperhomocysteinemia due to Methionine Synthase Deficiency, cbIG: Structure of the MTR Gene, Genotype Diversity, and Recognition of a Common Mutation, P1173L. *Am. J. Hum. Genet.* 71:143-153, 2002.
- David Watkins & David S. Rosenblatt. Update and new concepts in vitamin responsive disorders of folate transport and metabolism. *J Inherit Metab Dis*, DOI 10.1007/s10545-011-9418-1, 2011
- Erocole L. Cavalieri et al.: Catechol ortho-quinones: the electrophilic compounds that form depurinating DNA adducts and could initiate cancer and other diseases. *Carcinogenesis* vol.23 no 6 pp.1071-1077, 2012.
- Gaughan DJ et al.: The methionine synthase reductase (MTRR) A66G polymorphism is a novel genetic determinant of plasma homocysteine concentrations. *Atherosclerosis*. 2001 Aug;157(2):451-6.
- Fitzgerald ND, Kara. Hodges, Romily. 2016 Methylation Diet and Lifestyle, Whole Being Support for Healthy Methylation and Epigenetic Expression.
- Jamerson B.D. et al.: Folate Metabolism Genes, Dietary Folate and Response to Antidepressant Medications in Late-Life Depression. *Int J Geriatr Psychiatry*, 2013 September; 28(9).
- Jane C. Figueiredo et al.: Global DNA Hypomethylation (LINE-1) in the Normal Colon and Lifestyle Characteristics, Dietary and Genetic Factors. *Cancer Epidemiol Biomarkers Prev.*, 18(4): 1041-1049, April 2009.
- Mojgan Hosseini: Role of Polymorphism of Methyltetrahydrofolate-Homocysteine Methyltransferase (MTR) A2756G and Breast Cancer Risk. *POL. J. PATHOL* 2013; 64 (3): 191-195
- Qiping Feng et al.: Human S-adenosylhomocysteine hydrolase: common gene sequence variation and functional genomic characterization. *J. Neurochem.* (2009) 110, 1806-1817.
- Sheila Dawling et al.: Catechol-O-Methyltransferase (COMT)-mediated Metabolism of catechol Estrogens: Comparison of Wild-Type and variant COMT Isoforms. *CANCER RESEARCH* 61, 6716-6722, September 15, 2001.
- Seong-Gene Lee et al.: Association of Ala72Ser polymorphism with COMT enzyme activity and the risk of schizophrenia in Koreans. *Hum Genet*, 116: 319-328, 2005.
- Xiang Tan and Minhwa Chen. Association between Catechol-O-methyltransferase rs4680 (G>A) polymorphism and lung cancer risk. *Diagnostic Pathology*, 9:192, 2014
- Ziva Cerne, et al: Combined effect of CYP1B1, COMT, GSTP1, and MnSOD genotypes and the risk of postmenopausal breast cancer, *Gynecol Oncol.* Jun 30;22 (2):110-9 2011 <https://pubmed.ncbi.nlm.nih.gov/21860737/>
- Pedro Vizán, Luciano Di Croce, Sergi Aranda Functional and Pathological Roles of AHCY, *Front Cell Dev Biol* 2021; 9 <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC8044520/>
- Sak, Katrin: The Val158Met polymorphism in COMT gene and cancer risk: role of endogenous and exogenous catechols, *Drug Metab Rev.* 2017 Feb;49 (1):56-83 <https://pubmed.ncbi.nlm.nih.gov/27826992/>
- Nagai, et al: Strong inhibitory effects of common tea catechins and bioflavonoids on the O-methylation of catechol estrogens catalyzed by human liver cytosolic catechol-O-methyltransferase, *Drug Metab Dispos.* 2004 May;32 (5): 497-504 <https://pubmed.ncbi.nlm.nih.gov/15100171/>
- Qiping Feng, et al: Human S-Adenosylhomocysteine Hydrolase: Common Gene Sequence Variation and Functional Genomic Characterization, *J Neurochem*, 2009 Sep; 110 (6): 1806-1817 <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC2838417/>