myDNA Comprehensive Health Report





Welcome to the future of health and human potential

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TEST METHODOLOGY AND LIMITATIONS

Recommendations in this report apply to all ages, however for any patient under 18 years, a guardian must purchase the test and be present for the report recommendations. The information in this report is not intended to treat, diagnose or cure any medical condition or disease.

Gene By Gene, a wholly owned subsidiary of myDNA, Inc., is a College of American Pathologists (CAP) accredited and Clinical Laboratory Improvement Amendments (CLIA) certified clinical laboratory qualified to perform high-complexity testing. This test was developed and its performance characteristics determined by Gene by Gene. It has not been cleared or approved by the FDA. FDA does not require this test to go through premarket FDA review. This test is used for clinical purposes. It should not be regarded as investigational or for research. Only the genomic regions listed below were tested; there is a possibility that the tested individual is a carrier for additional, undetected mutations. Although molecular tests are highly accurate, rare diagnostic errors may occur that interfere with analysis. Sources of these errors include sample mix-up, trace contamination, and other technical errors. The presence of additional variants nearby may interfere with mutation detection. Genetic counseling is recommended to properly review and explain these results to the tested individual.



Speak to our Specialists

As part of our clinical service, we have a team of clinical experts available to answer any questions you may have about the report or about nutrigenomics in general. If you have any such queries, please contact us at: support@mydna.life

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Apolipoprotein E (ApoE) is a lipid-binding protein that transports triglycerides and cholesterol in multiple tissues, including the brain. The e4 allele is common in hunter-gatherer communities, while the e3 and e2 alleles are most common in agricultural communities.

- ApoE 3/3 is the most common ApoE genotype found in agricultural communities and has numerous benefits
- Extended cognitive fitness and enhanced expression of anti-aging sirtuins
- · Improved HDL and LDL profile
- Improved ability to repair synapses and neural protection
- · Higher viral protection
- · Higher response to plant bioactive compounds
- In the brain, e2 and e3 accumulate in neurons 2 to 4-fold higher than e4



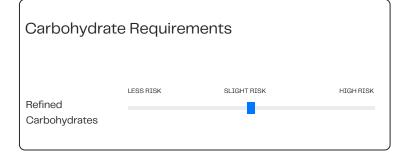
Traditional protein intake ranges based on latitude from less than 18% of total calories to approximately 35% in the far northern climates. Recommended protein intake varies based on weight and exercise intensity.

 Genetically, your requirements fall on the higher than average side of the spectrum, approximately 20% or more of your total caloric intake



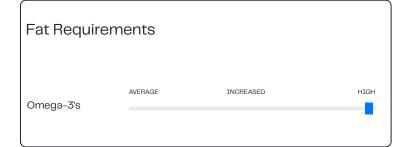
Your carbohydrate intake range is based on the latitude of your ancestors and whether a hunter-gatherer diet or modern agricultural diet made a larger imprint on your genes.

- Your genotype combination is associated with a slightly lower recommended carbohydrate intake, ranging from 30% to 40% of total calories from carbohydrates
- For a 2,000 calorie diet, this comes to 150-200 grams of carbohydrates per day



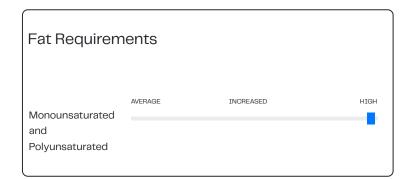
The differences between responses in individuals to refined carbohydrates have been linked to a genetic adaptation occurring during the agricultural age.

- Your genotype will benefit from limiting refined carbohydrates and choosing fiber-rich carbohydrate sources
- Refined carbohydrate sources include cookies, crackers, chips, tortillas, white bread, refined sugar, and high-sugar juices
- · Choosing low glycemic foods is recommended



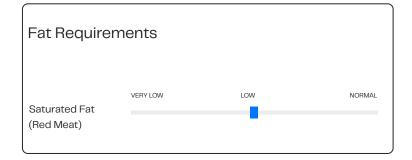
The NIH has set the recommended intake of omega-3's from 1.1 to 1.6 grams per day from a combination of ALA, EPA and DHA. Omega-3 fatty acids are essential for brain, eye, and cardiovascular health.

- Your genotype combinations are associated with a higher requirement of EPA and DHA
- ApoE e2 and e3 carriers can benefit from non-phospholipid fish oil intake, however, e4 carriers should use phospholipid-based EPA and DHA as found in fish and fish roe
- For ApoE e4 carriers, fish oil supplements do not appear as effective as phospholipid-based EPA and DHA as found in fish and fish roe
- E4 carriers may have impaired transport of free DHA and require phospholipids for successful transport



Traditional total fat intake ranges based on latitude, with as low at 25% consumed in countries closer to the equator, and up to 55% of total calories from fat being consumed in northern latitudes.

- Genetic testing can show which fats to focus on, but total fat will range based on your climate and health goals
- Your genoypes are associated with a higher emphasis on monounsaturated and polyunsaturated fats from olive oil, avocados, poultry, nuts and seeds



The 2020 Dietary Guidelines in the U.S. recommends limiting calories from saturated fats to less than 10% of the total calories you eat and drink each day. That's about 200 calories for a 2,000 calorie diet. Traditionally, saturated fat intake from animal foods ranged based on the season and the geographical location, with higher latitudes and more mountainous regions consuming more.

- Based on your genotype combinations, you should aim to get less saturated fat in your diet from red meat
- · Limit your red meat consumption to twice a week

Fat Requirements

VERY LOW NORMAL

Saturated Fat (Dairy)

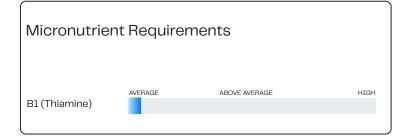
The 2020 Dietary Guidelines in the U.S. recommends limiting calories from saturated fats to less than 10% of the total calories you eat and drink each day. That's 200 calories for a 2,000 calorie diet.

 Your genotype combinations are associated with benefiting from getting less than 22 grams of saturated fat in your diet, especially from dairy



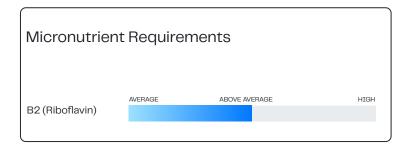
Celiac disease is an immune reaction to eating gluten, a protein found in wheat, barley and rye. Published research shows that approximately 30 percent of the general population have variants in the celiac disease risk genes HLA-DQA1 through HLA-DQB, yet only 3% of these individuals develop celiac disease.

- Your genotype combination is associated with a low genetic risk for celiac disease
- On a global level, the rates of celiac disease are not related either to the amount of wheat consumed by each country or to the prevalence of the HLA DR3-DQ2 and DR4-DQ8 genotypes worldwide
- First-degree relatives of people with celiac disease including parents, siblings and children have a 1 in 10 risk compared to 1 in 100 in the general population, which may be increased by existing autoimmune disorders



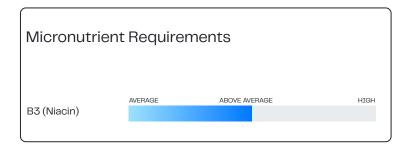
The recommended daily allowance (RDA) for thiamine is 1.2mg. Thiamine requirements are analyzed based on ethanol metabolism, however, chronic intake of alcohol depletes thiamine.

· Your genotype is associated with an average need for B1



The recommended daily allowance (RDA) for riboflavin is 1.3mg. Riboflavin is used as a co-factor for numerous reactions associated with protein, fat, and carbohydrate metabolism. Riboflavin requirements are analyzed based on MTHFR gene function.

- Your genotype is associated with a higher than average need for $\ensuremath{\mathsf{B2}}$
- B2 is high in liver (2.8mg), lamb (0.4mg), salmon (0.8mg), yogurt (0.6mg) and oyster mushrooms (0.3mg)



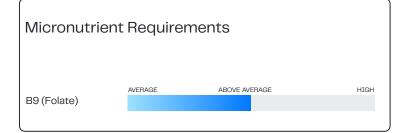
The recommended daily allowance (RDA) for niacin is 16mg. Niacin targets genes associated with cardiovascular and skin health, while also balancing methylation levels.

- Your genotype is associated with a higher sensitivity to low niacin intake
- Niacin in high in yellowfin tuna (37.5mg), canned tuna (21.9mg), wild salmon (17mg), ground turkey (20mg), chicken breast (16mg), liver (14.2mg), skirt steak (9.5mg), white button mushrooms (6.8mg), and brown rice (5.2mg)

Micronutrient Requirements AVERAGE ABOVE AVERAGE HIGH B6 (Pyridoxine)

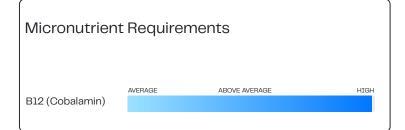
The recommended daily allowance (RDA) for B6 is 1.7mg. B6 deficiency can manifest as anorexia, irritability, anxiety, depression, muscle pain, bad PMS/low progesterone, nausea, seizures, migraines, dermatitis, age related macular degeneration (with low folate and B12) and lethargy.

- Your genotype is associated with a higher than average need for B6
- Women of reproductive age, especially current and former users of oral contraceptives, teenagers, male smokers, non-Hispanic African-American men, and men and women over age 65 are most at risk of B6 deficiency
- B6 is high in yellowfin tuna (6 oz, 1.8mg), wild salmon (6 oz, 1.2mg), liver (3oz, 0.8mg), chicken breast (6 oz, 1mg), unfiltered fermented drinks (16oz, 0.8mg), pistachios (1 oz, 0.5mg), avocado (1 whole, 0.5mg), sweet potatoes (1 whole, 0.3mg), and spinach (1/2 cup, 0.1mg)



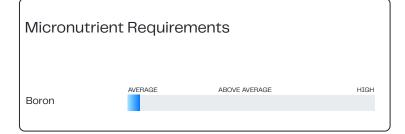
The recommended daily allowance (RDA) for folate is 400mcg. Folate is one of the – if not most – influential nutrigenomic micronutrient. It has a powerful influence on genes related to pregnancy, homocysteine, and cancer prevention.

- Your genotype is associated with a higher than average need for folate
- Folate is depleted by proton pump inhibitors, oral contraceptives, NSAIDs, anticonvulsants, antivirals, antibiotics, and antacids
- Folate is high in liver (3 oz., 215mg), collard greens (1 cup cooked, 177mcg), beets (1 cup raw, 148mcg), black-eyed peas (1/2 cup, 105mg), raw spinach (1 cup 58mg), asparagus (4 spears, 89mg), hummus (1/2 cup, 83mcg), broccoli (1/2 cup cooked, 52mg), romaine lettuce (1 cup, 64mg), strawberries (1 cup, 40mcg), orange (1 whole, 39mcg), sprouted lentils (1/2 cup, 38mcg), and parsley (1 sprig, 15.2mg)



The recommended daily allowance (RDA) for B12 is 2.4mcg. B12 influences genes related to homocysteine, brain health, pregnancy, and energy. B12 requirements are based on serum levels associated with the FUT2 gene.

- · Your genotype is associated with low serum B12 levels
- B12 is depleted by antacids, antibiotics, proton pump inhibitors, Metformin, anticonvulsants, oral contraceptives, certain psychiatric medications
- Older adults, vegans, digestive disorders, and those who take Metformin or PPI's are at risk for B12 deficiency
- B12 is highest in liver (3 oz, 70.7 mcg), clams (3oz, 17mcg), wild salmon (6 oz, 5.2mcg), ground beef (4.8mcg), yogurt (6 oz, 1.0mcg), eggs (1 whole, 0.5mcg), and cheddar cheese (1.5 oz, 0.5mcg)



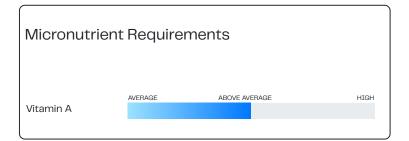
The recommended daily allowance (RDA) for boron has not been set, but 1-3mg is considered adequate. Boron is connected to bone health, hormone health and healthy SAMe levels for brain health. Men with low testosterone and women with osteoporosis or osteopenia will benefit from more boron.

• Your genotype is associated with an average need for boron

Micronutrient Requirements Choline & Betaine AVERAGE ABOVE AVERAGE HIGH

The recommended daily allowance (RDA) for choline is 550mg, while betaine hasn't been set. The more betaine you consume, the less choline you require. Choline is crucial for pregnancy, lowers anxiety, prevents fatty liver, assists detoxification, and improves memory.

- Your genotype is associated with a higher than average need for choline and betaine
- Choline is depleted by nighttime pain relievers, antihistamines, sleep aids, antidepressants, incontinence drugs and narcotic pain relievers
- Intense endurance exercise depletes choline levels, and increasing phosphatidylcholine has been found to improve exercise capacity during high-intensity cycling and running, as well as reduce muscle soreness
- Choline is highest in liver (3 oz, 356mg), pastured eggs (2 eggs, 294mg), beef round (6 oz, 234mg), heart (3 oz, 194mg), chicken (6 oz, 144mg), wild cod (6 oz, 142mg), bacon (3.5 oz, 125mg), and edamame (1/2 cup, 107mg)
- Betaine is highest in spinach (3.5 oz., 645mg), shrimp (3.5 oz., 218mg), beets (3.5 oz., 200mg) and whole grain sourdough wheat bread (2 slices, 201mg)



The recommended daily allowance (RDA) for vitamin A is 900 mcg for men and 700 mcg per day for women. Vitamin A assists digestive lining repair, oral health, eye health, iron mobilization, mitochondria health, skin health, healthy lung function, and increased immunity.

- Your genotype is associated with a 32% lower conversion rate of betacarotene to vitamin A, making it important to include more animal-based vitamin A to hit your daily target
- Vitamin A is high in liver (3 oz., 6,600mcg), pastured eggs (1 egg, 75mcg), cod liver oil (378mcg), wild salmon oil (206mcg), pickled herring (219mcg) and sockeye salmon (118mcg)

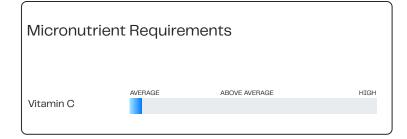
Micronutrient Requirements

AVERAGE ABOVE AVERAGE HIGH

Vitamin D

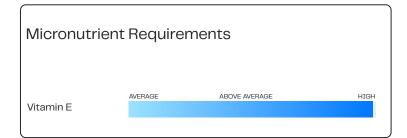
The recommended daily allowance (RDA) for vitamin D is 20mcg. Vitamin D has a wide role in immune function, bone health, cardiovasular health and cancer prevention.

- Your genotype is associated with below average circulating levels of vitamin D
- Vitamin D is depleted by obesity, pesticides, a high fructose intake, anticonvulsants, barbiturates, benzodiazepines, calcium channel blockers, corticosteroids, antidepressants, and bronchodilators
- Vitamin D is highest in sockeye salmon (6 oz., 28.4mcg), cod liver oil (1 tsp., 11mcg), canned tuna (1 can, 6.7mcg), wild herring (3 oz., 5.4mcg), sardines (1 can, 4mcg)



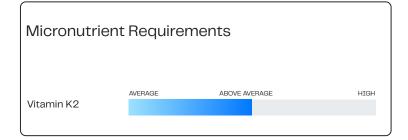
The recommended daily allowance (RDA) for vitamin C is 90mg, however, the amount consumed in the Paleolithic era was 400mg per day. Studies show the best results occur with over 500mg per day. Extensive research shows that adequate vitamin C reduces the risk of cancer, heart disease, colds, flu, cataracts, hypertension and even depression.

• Your genotype is associated with average serum vitamin C levels



The recommended daily allowance (RDA) for vitamin E is 15mg. Vitamin E is important for antioxidant protection, skin health, fertility, brain health, and cardiovascular health.

- Your genotype is associated with a higher sensitivity to low vitamin E intake
- Vitamin E is highest in sunflower seeds (1 oz., 7.4mg), almonds (1 oz. 7.3mg), avocado (1 whole, 4.2mg), spinach (1 cup cooked, 3.7mg), butternut squash soup (1 cup, 2.6mg) and olive oil (1 tablespoon, 1.9mg)



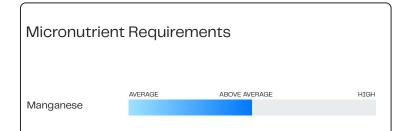
The recommended daily allowance (RDA) for vitamin K2 has not been established, but based on amounts found in food and research, 60–70mcg of MK–4 and MK–7 is a good target. MK–4 targets sex hormones, the brain, possesses anti–cancer and anti–inflammatory activity, and also supports bone health. MK–7 is considered better for reducing arterial calcification, increasing bone density, anti–cancer, improving salivary buffering (minimizes the demineralization of enamel and enhances its re–mineralization), and increasing cardiac output (12% increase) in athletes.

- Your genotype is associated with a higher sensitivity to low vitamin K2 intake
- Vitamin K2 is highest in natto (3.5oz., 108mcg MK-7), Münster cheese (1.7 oz., 50mcg of MK-4 and MK-7), Camembert cheese (1.7 oz., 34mcg of MK-4 and MK-7), dark chicken meat (6 oz., 90mcg MK-4), and pork chops (6 oz. 112mcg MK-4)

Micronutrient Requirements AVERAGE ABOVE AVERAGE HIGH Magnesium

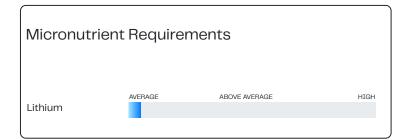
The recommended daily allowance (RDA) for magnesium is 400mg, however, higher amounts may be required for certain individuals, stress levels and athletes. Magnesium levels vary drastically based on the soil, and therefore in the food. Magnesium is involved in 300 biochemical reactions, and deficiency has widespread effects on every aspect of health. The most common symptoms of low magnesium includes calf cramps at night, headaches, arrhythmia, calcification, and muscle fatigue.

- Your genotype is associated with a higher than average need for magnesium
- Magnesium is depleted by smoking, sugar, chronic stress, high alcohol
 intake, coffee, tea, fluoridated water, phosphoric acid, non-fermented
 grains, intense exercise, high protein diets, high calcium supplementation,
 high arsenic levels, antacids, proton pump inhibitors, ACE inhibitors, birth
 control, hormone replacement therapy, Estradiol, Premarin, antibiotics,
 antivirals, immunosuppressants, methylphenidate, Tamoxifen and
 corticosteroids
- Magnesium is highest in sprouted pumpkin seeds (2 tablespoons, 156mg), hemp seeds (2 tablespoons, 116mg) chia seeds (2 tablespoons, 111mg), Gerolsteiner mineral water (1 liter, 100mg), spinach (1/2 cup cooked, 78mg), wild salmon (6oz, 52mg), and peanut butter (2 tablespoons, 49mg)



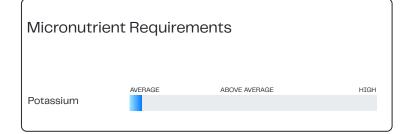
The recommended daily allowance (RDA) for manganese has not been set, however, 1.8 to 2.3mg per day is considered adequate. Manganese has a special role in protecting the mitocondria of the cells against toxicity through superoxide dismutase. Manganese is crucial for heart health, blood sugar, male fertility, bone health and protecting the brain against glutamate toxicity.

- Your genotype is associated with a higher sensitivity to low manganese intake
- Manganese is highest in mussels (3 oz., 5.8mg), wild blueberries (1/2 cup, 2.87mg), hazelnuts (2 tablespoons, 1.6mg), pecans (2 tablespoons, 1.1mg), oysters (3 oz., 1mg), clams (3 oz., 0.9mg), hummus (1/2 cup, 0.9mg), spinach (1/2 cup cooked, 0.8mg), and cultivated blueberries (1/2 cup, .33mg)



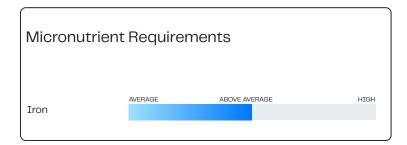
The recommended daily allowance (RDA) for lithium has not been set, with normal intake ranging from 250mcg to 3mg. Studies found an association between higher levels of lithium in local water and "beneficial clinical, behavioral, legal and medical outcomes." In the context of your genetic analysis, we are looking at lithium requirements in regards to B12 transportation.

Your genotype is associated with an average need for lithium



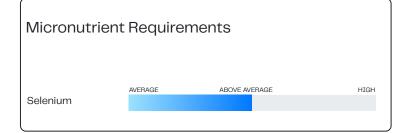
The recommended daily allowance (RDA) for potassium is 3,400mg for males and 2,600mg for females. Potassium is an electrolyte that helps maintain normal levels of fluid inside our cells, muscle contraction, and regulates blood pressure. The Paleolithic hunter–gatherers took in about 11,000 milligrams of potassium a day from fruits, vegetables, leaves, flowers, roots, and other plant sources, and under 700 mg of sodium.

· Your genotype is associated with an average requirement for potassium



The Recommended Dietary Allowance (RDA) for all age groups of men and postmenopausal women is 8 mg/day and the RDA for premenopausal women is 18 mg/day.

- Your genotype combinations are associated with with lower serum iron levels and a moderate need for dietary iron intake
- Animal-based foods and seafood contains heme iron, while plant foods contain non-heme iron
- Heme iron has a higher absorption rate compared to non-heme iron
- Iron is highest in oysters (3oz, 8mg), beef liver (3 oz, 5mg) beef 6 oz, 4mg), sardines (3 oz, 2mg) white beans (1 cup, 8mg) dark chocolate (3 oz, 7mg), spinach (1/2 cup cooked, 3mg)



The recommended daily allowance (RDA) for selenium is 55mcg. Selenium levels in plant and animal foods vary drastically based on the soil. Selenium is a crucial mineral linked to numerous genes involved in glutathione (the master antioxidant), detoxification, immunity, thyroid health, skin health, and cancer prevention.

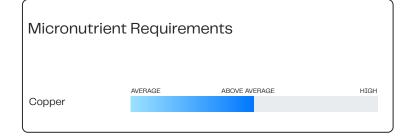
- Your genotype is associated with a higher than average need for selenium
- Selenium is depleted by oral contraceptives, Statin drugs, and hormone replacement therapy
- Selenium is hightest in oysters (3.5 oz., 130mcg), pork chops (6 oz., 80.6 mcg), beef (6 oz., 61.2 mcg), chicken breast (6 oz., 54.2 mcg), shrimp (3 oz., 42.1 mcg), eggs (2 whole, 40mcg), shiitake mushrooms (1 cup, 36 mcg), and sourdough wheat bread (2 slices, 24.1mcg)

Micronutrient Requirements

AVERAGE ABOVE AVERAGE HIGH

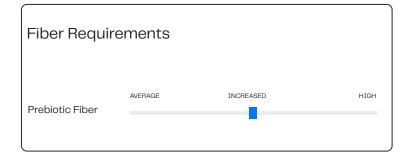
The recommended daily allowance (RDA) for zinc is 11mg. Zinc is poorly absorbed from plant foods and is highest in animal foods. Zinc plays a special role with numerous genes connected to immunity, cancer prevention, detoxification, skin health, eye health and more.

- Your genotype is associated with an average serum zinc levels $% \left(1\right) =\left(1\right) \left(1\right) +\left(1\right) \left(1\right) \left(1\right) +\left(1\right) \left(1\right) \left$



Typical diets meet or exceed the copper RDA and copper deficiency is rare. The RDA for copper is 900mcg. Copper is involved in the regulation of gene expression, brain development, neurotransmitters, cardiovascular health, and immune system functioning.

- Your genotype is associated with slightly lower serum copper levels
- Copper is highest in liver (3 oz. 12,400mcg), oysters (4,850mcg), potatoes (675mcg), shiitake mushrooms (1/2 cup, 650mcg), cashew (1 oz., 629mg), sunflower seeds (1/4 cup, 615mg) and dark chocolate (1 oz., 615mcg)



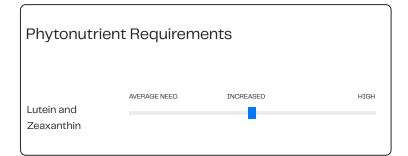
The recommended amount of fiber is up to 25 grams per day for women and up to 38 grams per day for men. $\,$

 Your genotypes are associated with a slighter higher than average requirement for prebiotic fiber

Phytonutrient Requirements AVERAGE NEED INCREASED HIGH Phytoestrogens

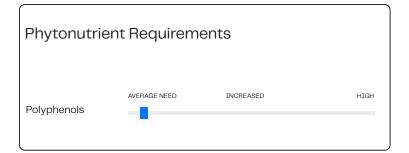
Phytoestrogens are plant derived compounds found in a wide variety of foods. There are pros and cons to phytoestrogen intake that appears to have a genetic, age, and gut health connection for determining optimal intake.

 Your genotype combinations are associated with a low to average need for phytoestrogens for healthy hormones



A recommended daily intake of lutein and zeaxanthin hasn't been established. Lutein and zeaxanthin can help protect your eyes from harmful high-energy light waves like UV sunlight.

- Your genotype is associated with an increased need for foods high in lutein and zeaxanthin to support eye health
- Around 700 carotenoids have been discovered and only lutein and zeaxanthin are found in the eye
- American adults typically consume 1–3 mg/day of lutein and zeaxanthin, the Spanish consume 3.5 mg/day, the Germans consume 5.33 mg/day, and older Australians consume 0.9mg per day
- For reducing the risk of eye disorders, the estimated target is 6mg or more of lutein and zeaxanthin daily
- The foods highest in lutein and zeaxanthin include cooked spinach (1/2 cup, 12.64 mg lutein), raw spinach (1/2 cup, 6.6mg lutein), cooked kale (1/2 cup, 8.88mg lutein), egg yolks (1 egg, 237mcg lutein and 216mcg zeaxanthin), and orange peppers (208mcg lutein and 1665mcg zeaxanthin)



Research strongly suggests that long term consumption of diets rich in plant polyphenols offer protection against development of cancers, cardiovascular diseases, diabetes, osteoporosis and neurodegenerative diseases.

 Your genotype is associated with a slower metabolism of certain polyphenols, which means you have a higher benefit with a lower intake of green tea, coffee, berries, and chocolate



 $\label{thm:causing hypoglycemia and increases satiety.} Cinnamon lowers blood glucose usually without causing hypoglycemia and increases satiety.$

 Your genotype is associated with a higher need for cinnamon to control blood sugar and satiety

Phytonutrient Requirements AVERAGE NEED INCREASED HIGH Cruciferous Vegetables

Isothiocyanates from cruciferous vegetables are known for their anti-cancer activity. Certain genotypes require higher levels of this anti-cancer activity.

- Your genotype combinations are associated with a higher requirement of cruciferous vegetables
- Cruciferous vegetables include broccoli, Brussels sprouts, cabbage, cauliflower, radishes, turnips, Bok choy, and watercress
- · Aim for 1-2 cups of cruciferous vegetables per day



Lycopene is found in tomatoes, watermelon, guava and pink grapefruit, and has unique benefits for the heart, breast, prostate and skin.

 Your genotype combinations are associated with a lower to average requirement for lycopene



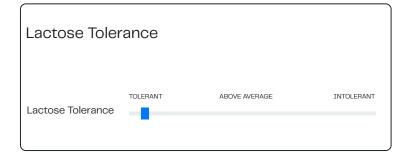
Apigenin is a flavonoid that possess anti-inflammatory, antioxidant and anticancer properties. Certain genotypes require higher levels for breast health.

 Your genotype is associated with an average need for apigenin for breast health



Resveratrol is a plant chemical produced in response to stress from the elements, and has been found to protect against heart disease and potentially extend life.

 Your genotype combinations are associated with an average requirement for resveratrol and heart health



Lactose is the major carbohydrate in milk. The arrival of farming in Europe around 8,500 years ago necessitated adaptation to new environments, pathogens, diets, and social organizations. One of the best examples of genetic dietary changes to this is the lactase enzyme in northern Europeans that only dates to the last 4,000 years.

- Your LCT genotype is associated with lactose tolerance
- The ability to digest lactose is much more common in people of European ancestry
- Approximately 32 percent of the world's population is lactose tolerant
- · Since this gene only looks at lactose, sensitivities to dairy can still exist



Variants in the CYP1A2 gene determine the rate at which you metabolize caffeine.

 You are a fast metabolizer of caffeine, meaning that you feel the effects of caffeine more quickly, but the effects also wear off quickly





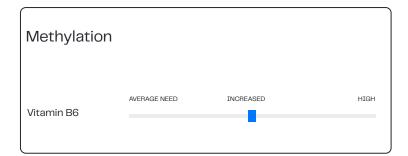
MTHFR 677 and MTHFR 1298 genotypes determine your folate requirements to assist normal homocysteine levels.

- Your genotype combination is associated with a higher than average requirement for folate to maintain healthy homocysteine levels
- If your homocysteine is elevated, check that you are getting enough folate
- High homocysteine has been implicated in amyloid buildup, DNA damage and cancer, mitochondrial dysfunction, cardiovascular disease, agerelated macular degeneration, apoptosis of neurons and depression



Folinic acid is a second type of folate found in folate-rich food.

 You have a higher than average requirement for folinic acid to maintain healthy methylation and homocysteine levels



Vitamin B6 plays an important role in homocysteine metabolism and CBS gene function.

- A combination of your genotypes related to vitamin B6 serum levels and methylation requirements are associated with a higher than average requirement for B6 to maintain healthy methylation and homocysteine levels
- B6 is highest in wild salmon, wild cod, pistachios, avocados, Yukon gold or red potatoes, taro root, sweet potatoes, spinach, cauliflower and unfiltered fermented drinks
- Many medications deplete B6 including antibiotics, oral contraceptives,
 ACE inhibitors, antacids, and proton pump inhibitors

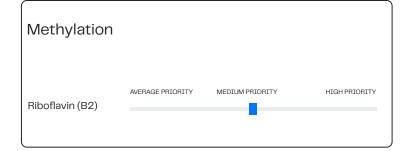
Methylation

AVERAGE NEED INCREASED HIGH

Vitamin B12

Vitamin B12 plays an important role in homocysteine metabolism.

- You have a higher than average requirement for B12 to maintain healthy methylation and homocysteine levels
- B12 is highest in animal foods and seafood
- B12 is depleted by antacids, antibiotics, proton pump inhibitors, Metformin, oral contraceptives, and yeast overgrowth



Vitamin B2 plays a special role in stabilizing the MTHFR gene for homocysteine metabolism.

- Your genotype is associated with a higher than average requirement for riboflavin to maintain healthy methylation and homocysteine levels
- B2 is highest in liver, lamb, fish, yogurt and mushrooms



Choline and betaine play a crucial role in homocysteine metabolism, especially for those with variants in MTHFR.

- Your genotype is associated with a higher than average requirement for choline and betaine to maintain healthy methylation and homocysteine levels
- Low choline intake can manifest as memory issues, NAFLD, anxiety, neurological disorders, breast cancer, histamine issues, gallbladder issues, and SIBO
- Choline may be depleted by nighttime pain relievers, antihistamines, sleep aids, antidepressants, incontinence drugs and narcotic pain relievers
- Intense endurance exercise depletes choline levels, and increasing phosphatidylcholine has been found to improve exercise capacity during high-intensity cycling and running, as well as reduce muscle soreness



Certain genotypes in the folate metabolism pathway can affect the metabolism of synthetic folic acid, leading to high circulating levels.

- Your genotype combinations may decrease the metabolism of synthetic folic acid, which can increase circulating levels and affect overall folate metabolism
- When possible, avoid foods and drinks fortified with folic acid, and supplements that use folic acid
- Supplements that use methylfolate or folinic acid are better options





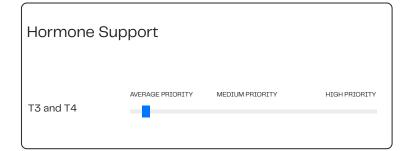
Certain glutathione SNPs are associated with breast protection.

- Your genotypes for multiple genes are associated with slightly lower glutathione protection for breast health
- Boosting glutathione can be accomplished with selenium, glycine, cysteine, vitamin C, and cruciferous vegetables



There are multiple genes in the estrogen detoxification pathway that have a cumulative value on the ability to properly detoxify estrogen.

- Your combination of numerous genotypes in the estrogen pathway are associated with reduced estrogen detoxification
- To reduce the risk of harmful estrogen metabolites, you should avoid xenoestrogens, manage stress levels, and focus on gut health
- Increasing prebiotic fiber, polyphenols, magnesium and bifidobacteria may improve breast health by reducing the amount and activity of harmful estrogen metabolites



 $\mathsf{T3}$ and $\mathsf{T4}$ level variations have been associated with variants in the $\mathsf{DIO1}$ gene.

- Your genotype is associated with normal T3 and T4 levels
- T3 and T4 can still be out of range based on other epigenetic factors



ADIPOQ encodes for adiponectin, a protein secreted by fat cells that affect insulin and glucose metabolism. Low levels of adiponectin play a role in obesity, insulin resistance and Type 2 diabetes.

 Your genotype is associated with normal adiponectin levels, which can increase the effect of insulin, improve glucose metabolism and assist a healthy body weight

Hormone Support

AVERAGE PRIORITY MEDIUM PRIORITY HIGH PRIORITY

Ghrelin

Variants in genes related to ghrelin levels and dopamine receptor density have been shown to create a larger appetite and the potential for overeating in multiple populations.

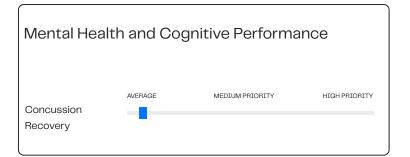
- Your genotypes are not associated with higher ghrelin levels
- You are at a decreased risk for overeating and abdominal weight gain



Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Brain Repair and Maintenance

Multiple genes are responsible for daily neural repair and maintenance, and a combination of genotypes are associated with decreased neural repair.

- Your genotype combination is associated with average to improved neural repair
- You can be proactive for neural repair with eight hours of sleep per night,
 DHA, B-vitamins, Lion's Mane mushrooms, zinc, vitamin C, and vitamin E



A combination of genotypes in the pathways responsible for glutamate transport and modulation, BDNF levels, neural repair, and inflammation during a concussion are associated with delayed or improved recovery.

 Your genotype combination is associated with improved recovery from concussions

Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Episodic Memory

The 5-HT2A gene is associated with episodic memory, which is the ability to recall details of an event.

• Your 5-HT2A genotype is associated with an improved episodic memory

Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Cardio, Mood and Cognitive Fitness

Cardiovascular exercise has a tremendous influence on neurotransmitter balance, memory and cognitive fitness.

 You have an average requirement for cardiovascular exercise to improve mood and cognitive fitness

Mental Health and Cognitive Performance Mood (Folate) AVERAGE MEDIUM PRIORITY HIGH PRIORITY HIGH PRIORITY

MTHFR genotypes determine folate requirements for healthy BH4 levels responsible for neurotransmitter balance. The current daily value for folate is 400mcg DFE.

- Your genotype may require 400-500mcg (or more) for healthy BH4 levels responsible for neurotransmitter balance
- · Foods high in folate include:
- · Liver (215mg) 3 oz.
- Spinach (131mg) 1/2 cup cooked
- · Asparagus (89mg) 4 spears
- Brussels sprouts (78mg) 1/2 cup
- Broccoli (52mg) 1/2 cup

Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Anxiety (Choline)

The PEMT gene is associated with your required choline intake to support memory, anxiety and REM sleep.

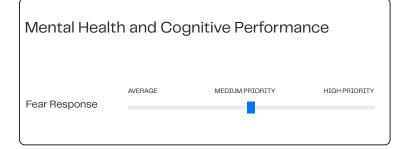
- Your PEMT genotype is associated with a higher need for choline (550mg or more) to support memory, anxiety and REM sleep
- Research has shown that uridine, DHA, and choline combined increases levels of phosphatidylcholine in the brain more than each on their own
- · Foods high in choline include:
- · Liver (356mg) for 3 oz.
- · Egg (294mg) for 2 eggs
- · Beef top round (234mg) for 6 oz.
- Chicken breast (144mg) for 6 oz.
- Chicken thigh (120mg) for 6 oz.
- Edamame (107mg) for 1/2 cup

Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Anxiety (Glutamate and GABA)

Anxiety is linked to altered levels of one or multiple neurotransmitters.

Understanding the genetic link to specific levels of neurotransmitters can help you be precise in your approach to reduce anxiety.

- Your genotype combination is associated with a reduced modulation of glutamate levels that could cause high glutamate and low GABA that could lead to anxiety
- Too much sugar and caffeine can create a high glutamate and low GABA mental state
- Cardio exercise, HIIT training and yoga have all been found to balance glutamate and GABA levels
- B6 and magnesium help convert excess glutamate to GABA (the calming neurotransmitter)
- Increase prebiotic intake to increase GABA levels and slow down an overactive mind at night to assist sleep



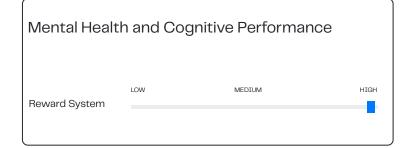
The FAAH gene is associated with anandamide levels, correlated with a heightened fear response to potential threats, while BDNF variants affect the ability to extinguish the fear response.

- Your genotype combination is associated with a heightened fear response that may affect your ability to extinguish fear memories
- Getting 30 minutes or more of aerobic exercise per day (especially in altitude), CBD, and hops help increase anandamide – known as the "bliss molecule" – to reduce the fear response

Mental Health and Cognitive Performance AVERAGE MEDIUM PRIORITY HIGH PRIORITY Addiction

ANKK1 modulates the density of dopamine receptors in the brain and is the most-studied genetic variant related to addictions. Variants have been associated with alcoholism, opioid addiction, sugar addictions, compulsive eating, obesity and Internet addiction.

 Your genotype is associated with a improved density of dopamine receptors for the ANKK1 gene, increasing dopamine targets within the striatum of the brain and reducing addiction susceptibility



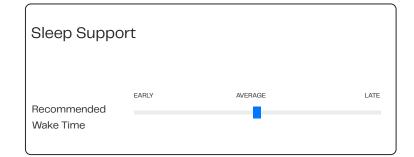
COMT rs4680 has been linked in a meta-analysis to variations in the reward response based on genotypes associated with low and high dopamine levels.

 Your genotype is associated with a higher dopamine response to reward processing that could improve motivation as well as decision making



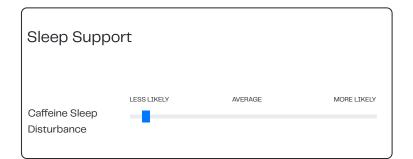
Your COMT genotype is associated with the "Strategist" that has the highest dopamine levels and may thrive more in low-pressure environments combined with complex problem-solving.

- If your levels of dopamine get too high and you find yourself irritable, impulsive, and stressed, add strength training 3–5 times a week and increase your magnesium and vitamin C intake for balance.
- Low catecholamine (coffee, green tea, red wine, chocolate) intake recommended due to their effect on dopamine
- For men and premenopausal women, avoid IPA beers due to a higher estrogenic effect that can slow COMT down further



Research has found that MTNR1B G allele carriers had a significant association with delayed melatonin release in the evenings and a substantially longer duration of elevated melatonin levels in the morning.

- Your genotype in the melatonin receptor gene is related to average wake times (6:30am or later) due to a slight delay of melatonin cessation
- To wake up earlier than 6:30am, you require more light exposure to assist ending melatonin release
- Melatonin supplementation is best avoided for G carriers due an increased risk of impaired glucose tolerance and elevated blood sugar



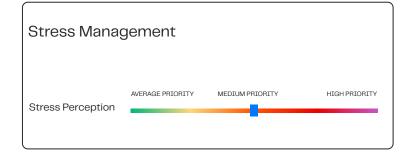
The rate at which caffeine is metabolized genetically is associated with variations of sleep disturbance.

 You are a fast metabolizer of caffeine, which means that you are less likely to have your sleep affected by caffeine



Acetylcholine plays a role in promoting REM sleep, the phase that occurs while we dream and where memory consolidation occurs.

- Your genotype is associated with increased sensitivity to not meeting your daily choline requirements for acetylcholine production and REM sleep
- You may be more sensitive to anticholinergic drugs, which block acetylcholine and have been found in research to cause cognitive decline
- Make sure you are getting at least 550mg of choline per day, walking 45 minutes or more per day, and if consuming alcohol, you will sleep better if you consume it before 6:00pm and limit the quantity



Your perception of stress is unique to your genotypes and life experience. Variants in 5-HT2A are associated with perceived stress, low vagal tone, anxiety, depression, OCD, and IBS, especially in females.

- · Your genotypes are associated with a higher perception of stress
- Moderate intensity aerobic exercise, meditation and yoga are recommended for stress relief
- Tryptophan, green or black tea, prebiotics, probiotics, B2, B6, B12, and folate all target the 5-HT2A gene to help lower stress perception



The G allele carriers of ADRB2 were associated with a higher percentage of IBS cases, twice the rates of anxiety, and functional chest pain diagnoses.

 Your genotype is associated with a reduced percentage of digestive disorders, IBS, and anxiety related to adrenaline levels



The 5-HT2A gene for the serotonin has a role in BDNF regulation – which helps regulate the excitatory neurotransmitter glutamate and calming neurotransmitter GABA. The A allele for 5-HT2A rs6314 was found to cause a lower expression of BDNF, and can compound the effect for those with variants in BDNF for mood and behavior in response to stress.

 Your genotype combination is associated with reduced susceptibility to low BDNF levels in response to stress, helping balance glutamate and GABA



Weight lifting has a higher impact on hormonal pathways that may provide higher levels of stress relief based on the speed of these pathways.

- Your genotype is associated with higher dopamine levels and a reduced clearance of adrenaline
- Weight lifting helps speed up the pathway responsible for clearing excess dopamine and adrenaline, and therefore is a useful tool for you to use for chronic stress





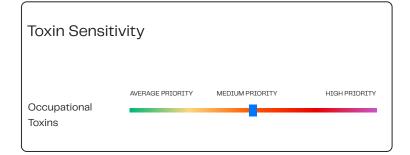
Mycotoxins are toxic compounds that are naturally produced by certain types of fungi. Research suggests that mycotoxins can decrease the formation of glutathione due to decreased gene expression of the enzymes needed to form glutathione.

- Your genotype is associated with lower glutathione levels which may cause glutathione depletion to occur at a faster rate and decrease mycotoxin detoxification
- The highest exposure to mycotoxins can be in foods grown or stored in damp conditions
- This may include grains, nuts, corn, coffee, wine, beer, grape juice, sorghum, rice, dried beans, apples, pulses, cacao products, and spices
- Boosting glutathione can be accomplished with selenium, glycine, cysteine, alpha lipoic acid, vitamin C, and cruciferous vegetables



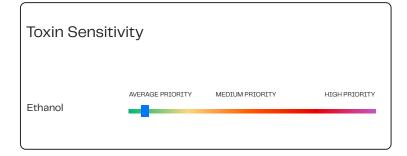
 $\label{thm:constraints} \mbox{Xenoestrogens are synthetic hormone disruptors found in plastics and pesticides.}$

- Your genotype is associated with a slower metabolism of xenoestrogens, and therefore the damage may be greater from xenoestrogen exposure
- Increasing magnesium targets the enzyme responsible for assisting xenoestrogen detoxification



Workers exposed to certain chemicals over a long period in the metalworking, petroleum, agricultural industries and in glass factories are at increased risk for occupational skin cancers.

- Your genotypes are associated with an increased sensitivity to these toxins
- Focus on zinc, selenium, niacin, and vitamin C to improve DNA protection for skin health
- Ellagic acid, lutein, zeaxanthin, cocoa polyphenols, chaga tea, green tea and citrus have all been found to help protect against skin damage and cancerous growth



ALDH2 encodes for aldehyde dehydrogenase, and variants can affect the levels of acetaldehyde and therefore the carcinogenic effect of alcohol.

 Your genotype is not associated with a higher risk of alcohol-related adverse reactions including flushing, palpitation, nausea, headache, drowsiness, breathlessness, and general discomfort



The International Agency for Research on Cancer has classified formaldehyde as carcinogenic to humans. Sources of formaldehyde in the home include building materials, smoking, household products, gas stoves, kerosene space heaters, as a food preservative, permanent-press clothes, and draperies, as a component of glues and adhesives, and as a preservative in some paints and coating products. Variants in the XRCC3 DNA repair gene have been associated with higher DNA damage from formaldehyde.

 Your genotype is not associated with an increased sensitivity to formaldehyde



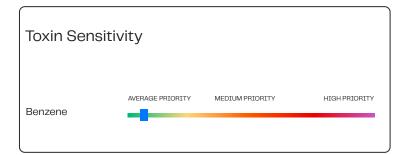
Benzo(a)pyrene is a carcinogenic compound produced from the burning of wood or trash, tobacco smoke, asphalt, coal, diesel exhaust, charred meat, and gas cooking.

- Your genotype combinations are associated with decreased detoxification of benzo(a)pyrene
- It is recommended to increase your intake of cruciferous vegetables, vitamin C, vitamin E, vitamin A, resveratrol, curcumin, green tea, and white tea to protect and detoxify benzo(a)pyrene



Aromatic amines are found in cigarettes, rubber factories, hair dyes that contain 4-aminobiphenyl, and meat cooked at high temperatures.

- Your genotype combinations are associated with a poor detoxification ability of aromatic amines
- If your exposure is higher to aromatic amines, increase cruciferous vegetable intake, carotenoids, vitamin C, and use marinades for meat when barbecuing



Sources of benzene include cigarette smoke, gasoline, exhaust, fires, industrial emissions, paint, detergent, glue, furniture wax, building materials, paint, petrochemical manufacturing, oil storage tanks, urban-industrial areas, service stations, certain foods, certain creams, groundwater contamination, and underground gasoline leaks.

• Your genotype is associated with an average sensitivity to benzene



Aspartame is an artificial sweetener that has been linked to behavioral, neurological and cognitive problems, increased blood sugar, thyroid issues, and certain types of cancer.

- Your genotype combinations may lower the sensitivity to aspartame
- Possible neurophysiological symptoms include learning problems, headache, seizure, migraines, irritable moods, anxiety, depression, and insomnia
- Artificial sweeteners in general increased waist circumference 500 percent while aspartame increased blood sugar in diabetes-prone mice
- Aspartame has been found to contribute to the formation of tumors in the CNS such as gliomas, medulloblastomas, and meningiomas, increased lymphoma and leukemia and, is an excitotoxin to brain neurons
- Aspartame in the body further metabolizes to formaldehyde, and rat studies found that formaldehyde (as a metabolite of aspartame) caused increased TSH levels and worsens the capacity of the gland leading to thyroid failure



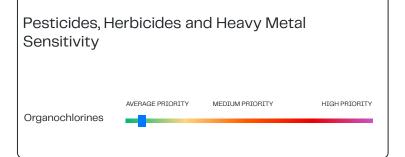
Food dyes have been found to inhibit mitochondrial respiration: the ability of the powerhouse of your cells to convert nutrients to energy. They have also been found to especially affect those with ADHD.

- Your genotype combination is associated with a higher sensitivity to food dyes
- Avoid foods and drinks that use food dyes when possible

Pesticides, Herbicides and Heavy Metal Sensitivity AVERAGE PRIORITY MEDIUM PRIORITY HIGH PRIORITY Glyphosate

Glyphosate is an herbicide that has been found to be highly toxic.

- Your genotype is associated with potentially more cellular damage from exposure to the herbicide glyphosate
- The highest glyphosate levels have been found in non-organic wheat and non-organic pulses like beans, lentils, and peas
- A meta-analysis of human epidemiological studies suggests a link between exposures to glyphosate and an increased risk for non-Hodgkin's lymphoma
- An association between glyphosate and thyroid disease comes from plots over time of the usage of glyphosate in the U.S. on corn and soy time-aligned with plots of the incidence rate of thyroid cancer in the U.S.
- Manganese deficiency and toxicity can occur simultaneously from glyphosate exposure due to a disruption in liver enzymes, causing transportation of manganese through the vagus nerve to the brainstem where excess manganese can lead to Parkinson's disease
- The gut bacterium Lactobacillus is negatively impacted by glyphosate and the depletion in associated with celiac disease
- Humic acid from Shilajit has been shown in vivo to reduce glyphosate concentration, inhibit the destructive effect of glyphosate on beneficial bacteria, and protect and repair against tight junction injury of the digestive system



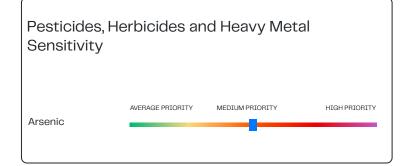
Organochlorines are found in certain pesticides, PCBs and sucralose.

Your genotype is associated with improved protection against organochlorines

Pesticides, Herbicides and Heavy Metal Sensitivity Organophosphate Insecticides AVERAGE PRIORITY MEDIUM PRIORITY HIGH PRIORITY

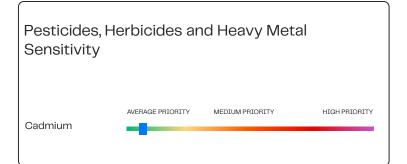
PON1 plays a large role in removing pesticides and is also involved with supporting HDL function and LDL oxidation. Organophosphates are a class of insecticides, including parathion and chlorpyrifos, that were among the most widely used insecticides available until the 21st century.

 Your PON1 genotype is associated with improved PON1 levels and detoxification of organophosphate insecticides



Arsenic is a heavy metal that is damaging at high levels. The highest dietary sources of inorganic arsenic include contaminated groundwater and rice. A consistent and growing body of evidence has shown that people who metabolize arsenic poorly may be at two to four times the risk of developing certain cancers and other arsenic-related diseases than people who are better metabolizers.

- Your genotype combination is associated with a reduced detoxification of arsenic
- The highest dietary sources of inorganic arsenic include contaminated groundwater and rice.
- Optimal selenium and folate intake have been found to improve arsenic detoxification and mitigate toxicity
- Chlorogenic acid a polyphenol highest in light roast coffee has been found to inhibit arsenic-induced neurotoxicity in mice
- Chlorogenic acid is also found in moringa tea, strawberries, cherries, bilberries, and wild blueberries



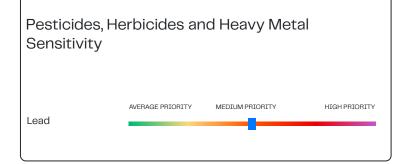
Chemical agriculture uses high amounts of synthetic organophosphates, creating a very high phosphorus content. Synthetic phosphorus concentrates the amounts of heavy metals, like cadmium in non-organic soils and food. Choosing organic produce is one of the best ways to avoid excess cadmium.

 Your genotype is associated with average detoxification of the heavy metal cadmium

Pesticides, Herbicides and Heavy Metal Sensitivity AVERAGE PRIORITY MEDIUM PRIORITY HIGH PRIORITY Mercury

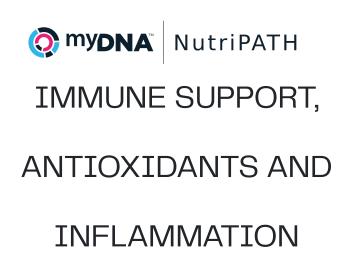
Mercury is a neurotoxin linked to neurological and behavioral disorders including tremors, insomnia, memory loss, neuromuscular effects, headaches, and cognitive and motor dysfunction. Burning coal for power and heat is a major source of mercury exposure. Glutathione is responsible for protecting against and detoxifying heavy metals like mercury.

- Your glutathione genotypes are associated with reduced protection against mercury toxicity
- Mercury is found in many pharmaceutical drugs, dental amalgams, and large fish including swordfish, ahi tuna, and halibut
- Selenium blocks mercury uptake, folate decreases mercury levels, and magnesium and holy basil protect against mercury toxicity



Lead-based paint, lead-based dust in older buildings, contaminated water, and air pollution are the major sources of lead. Exposure to lead over time may cause abdominal pain, constipation, depression, distraction, forgetfulness, irritability, and nausea.

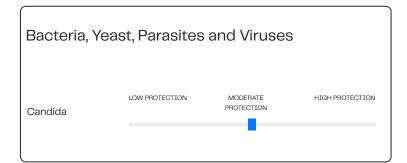
- Your genotype is associated with reduced detoxification and more toxic effects from elevated lead levels
- You may require more vitamin C and calcium if you are exposed to excess lead



Bacteria, Yeast, Parasites and Viruses H. Pylori AVERAGE PROTECTION HIGH PROTECTION

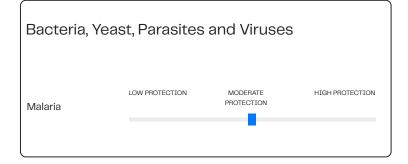
The inactive "non-secretor" genotype for FUT2 confers resistance to H. Pylori. H. Pylori is present in approximately 50% of the population in developed countries.

- You do not have the non-secretor genotype for FUT2, associated with an average susceptibility to H. Pylori
- H. Pylori inhibition has been demonstrated with alcohol extracts of the mushroom Lion's Mane



The inactive "non-secretor" genotype for FUT2 decreases resistance to Candida overgrowth.

 You have the secretor genotype for FUT2, giving you an average susceptibility to Candida overgrowth



Research has shown that MTHFR genotypes influence T-lymphocytes, natural killer cells, and protection against malaria.

- Your genotype is associated with moderate protection against malaria
- The malaria parasite needs higher amounts of folate to survive and replicate
- For malaria-endemic regions, your genotype does not provide as much protection as the homozygous genotype, but you have more protection than the wild-type genotype

Bacteria, Yeast, Parasites and Viruses

AVERAGE PROTECTION

HIGH PROTECTION

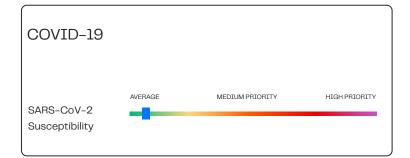
The inactive "non–secretor" genotype for FUT2 confers resistance to the Norovirus.

 You do not have the non-secretor genotype for FUT2, associated with an average susceptibility to the Norovirus

Bacteria, Yeast, Parasites and Viruses AVERAGE MODERATE PROTECTION DNA Viruses HIGH PROTECTION

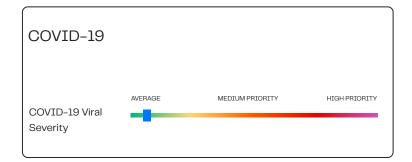
DNA viruses include HPV, Epstein Barre, herpes, and smallpox. Folate is a precursor to BH4 to produce nitric oxide. Nitric oxide acts as an antiviral that is more potent against DNA viruses.

- Your genotype combination is associated with slightly lower BH4 levels with insufficent folate, lowering protection against DNA viruses
- Low BH4 affects the aggressiveness of DNA viruses
- To increase BH4, include foods high in folate, vitamin C, L-arginine, B6, magnesium, and selenium for healthy nitric oxide levels and DNA virus support
- BH4 is depleted by high blood sugar, high omega-6 intake, chronic stress, high levels of mercury, arsenic, lead and aluminum, aspartame, and oxidative stress



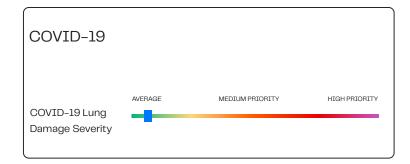
Genome-wide association studies have identified a region of chromosome 3p21.31 as the for conferring susceptibility to infection with LZTFL1 as the candidate gene. ApoE-e4, ACE2 and TMPRSS2 polymorphisms have been shown to be strongly associated with the susceptibility, severity, and clinical outcomes of COVID-19.

- Your genotype combination is associated with a reduced probability to SARS-CoV-2 infection
- Advanced age, obesity, and being male are considered the top risk factors for SARS-CoV-2 susceptibility, especially when combined with Type 2 diabetes, high blood pressure, and cardiovascular disease
- Research has shown that CBD, Chaga mushroom, birch bark and olive oil may stop SARS CoV-2 entry by helping block the "lock" for viral entry
- The flavonols kaempferol, quercetin, myricetin, fisetin and their derivatives were the most documented molecules with antiviral activities against SARS-CoV-2
- Propolis has antiviral activity and inhibitory effects on ACE2, TMPRSS2 and PAK1 signaling pathways used by SARS-CoV-2, while promoting immunoregulation of pro-inflammatory cytokines, and reducing the risk of cytokine storm syndrome



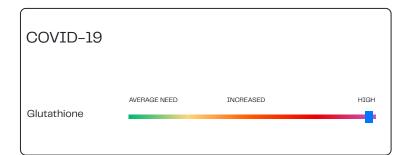
ACE2 (the receptor for SARS-CoV-2) is reduced in individuals that are carriers of ApoE4. Once the virus fuses with the cellular membranes, it takes control, shuts down more ACE2 receptors, closing the door behind it. Fewer ACE2 receptors lead to elevated angiotensin II levels, which increases the viral load. This leads to a more severe infection, NF-kb activation, lung damage, and viral replication.

- An analysis of your ApoE genotype and numerous genes involved in the viral inflammatory pathways show an association with a reduced risk of viral severity
- The risk of viral severity can still be increased by existing health issues including Type 2 diabetes, obesity, high blood pressure, and cardiovascular disease
- In September 2020, Boston University found that SARS-CoV-2 is highjacking the NF-kB pathway and should be a target for suppressing viral aggressiveness
- In addition to vaccination, NF-Kb inhibitors can be both preventative for inflammation and successful with slowing viral aggressiveness
- These include vitamin C, vitamin D, zinc, selenium, magnesium, cordyceps mushrooms, resveratrol, triterpenoids (Chaga, reishi, olive oil, holy basil), caffeic acid (coffee, Chaga, elderberry), and anthocyanins (elderberry, goji berries, cacao)
- The dietary flavonols kaempferol, quercetin, myricetin, fisetin were the
 most documented molecules with antiviral activities against SARS-CoV2, and had a broad spectrum of biological activities that could reduce
 the severity of infection symptoms and enhance the immune response
- The strong association between air pollution exposure and COVID-19 mortality suggests that inhaled noxious particles influence COVID-19 outcomes



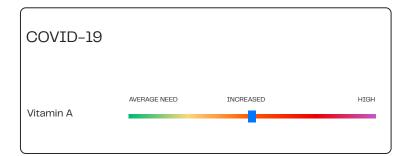
Scientists at the University of Oxford published results in November 2021 that variants in the LZTFL1 gene doubles the risk of lung failure and death from COVID-19. Approximately 60% of people with South Asian ancestry, 15% of people with European ancestry, 2% of people with African–Caribbean ancestry and 1.8% of people with East Asian ancestry carry the high-risk variant. Scientists found that the LZTFL1 gene high-risk variant affects the lungs, but does not have an impact on the immune system.

 Your genotype is not associated with an increased risk COVID-19 related lung damage severity



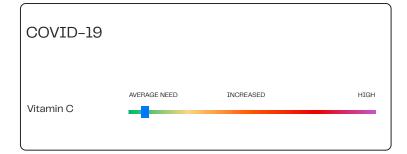
Glutathione is the master antioxidant system involved in oxidative stress, detoxification, and immunity. The functional capacity of immune cells and the ability to cope with oxidative stress has been proposed as one of the significant markers of health and longevity. In both animals and humans, those who reach exceptionally old age have immune markers the same as young adults.

- Your genotype combination is associated with decreased baseline glutathione levels
- Glutathione decreases with age, and low levels of glutathione are associated with chronic exposure to chemical toxins, heavy metals and excess alcohol, immunocompromised conditions, and neurodegenerative disorders
- Glutathione has been found to increase by 20% with deep breathing practices like Tai Chi or yoga
- For exercise, a combination of aerobic exercise and circuit weight training produced the highest glutathione effect
- Selenium, glycine, cysteine, vitamin C, and cruciferous vegetables all improve glutathione levels
- Chicken or bone broth, herbs, and spices are some of the best dietary ways to maintain higher levels of glutathione
- Some of the all-stars include cinnamon, anise, sage, and thyme due to also containing the antiviral compound caffeic acid



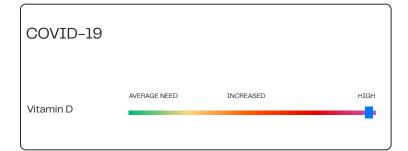
Vitamin A and some other retinoids show important immunomodulatory properties, including the ability to increase the efficiency of actions of type 1 interferons, an important antiviral cytokine released by the innate immune system against viral infections. Coronaviruses similar to SARS-CoV-2 can suppress the host IFN-I-based antiviral response as part of their infection mechanism.

- Your genotype is associated with a 32% lower conversion rate of betacarotene to vitamin A, making it important to include more animal-based vitamin A to hit your daily target
- Vitamin A intake by country shows that Spain is the country with the lowest population meeting nutritional requirements for vitamin A, followed by Belgium and Finland
- Germany and Portugal show the best, and with the exception of Finland, countries with suboptimal Vitamin A status are correlated (although not significantly) with their COVID-19 incidence and mortality



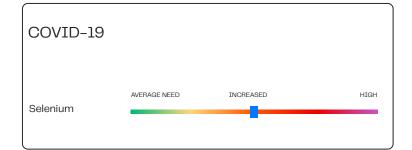
Optimal status of vitamin C plays an important role in the proper working of the immune system.

- Your genotype is associated with average serum vitamin C levels
- Countries such as the UK, France, Netherlands, and Belgium do not reach optimal dietary intake of vitamin C
- Germany stands out for its level of vitamin C intake in comparison with other countries
- Despite suboptimal vitamin C intake correlating weakly with COVID-19 incidence, it correlates strongly with deaths percentage, which could suggest a positive effect to fight infection once the individual has already been infected with SARS-CoV-2



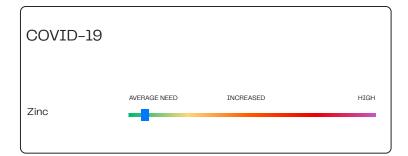
Vitamin D plays a key role in modulating the immune system, and suboptimal or deficient consumption of vitamin D is associated with various conditions related to a malfunction of the immune system and dysregulations in inflammatory status.

- Your genotype is associated with below average circulating levels of vitamin D
- A 2021 study found that those with the GG genotype for CYP2R1 were associated with 5.9 higher odds of experiencing severe COVID-19
- Vitamin D intake is deficient in all countries studied with COVID severity, with Spain, France, and Italy as the countries with the lowest intake
- A meta-analysis of the studies appears to show that vitamin D is only useful for those who are clinically low (below 20 ng/ml), with moderate doses daily or weekly to raise levels being more effective than periodic large doses



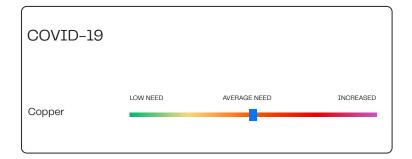
Suboptimal or deficient levels of selenium are associated with decreased cytotoxicity of NK cells, decreased antibody titers, and impaired cellular immunity. Supplementation is commonly related to improvements in cellular immunity and an improved optimal immune response against viruses, including an inhibitory effect on the development of the poliovirus and influenza.

- Your genotype is associated with a higher than average need for selenium
- Glutathione peroxidase 1 (GPX1) is a selenoenzyme with described antioxidant and antiviral properties that depends on nutritional selenium status
- Spain is at the top for meeting selenium requirements while Denmark is at the bottom
- The only two populations above the median of the countries analyzed included Finland and France, while the rest of the countries are below the general median



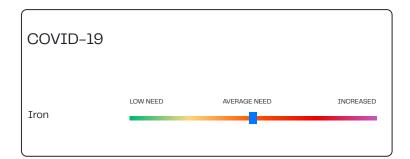
Optimal status of zinc is essential for the proper operation of the immune system and regulates NF-kb, where zinc deficiency in the setting of severe infection provokes a systemic increase in NF-kB activation. In vitro approaches have shown that zinc can inhibit SARS-CoV-1 replication.

- · Your genotype is associated with an average serum zinc
- Scientific evidence supports that optimal zinc intake or supplementation should be considered part of the strategy to reduce COVID-19 effects, with early reports finding that 15-23mg a day show significant improvement in symptoms



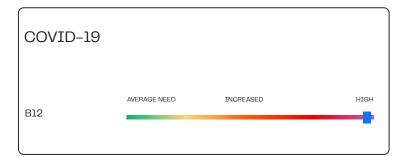
Copper plays a key role in optimal performance of relevant components of the immune system, such as NK cells, macrophages, neutrophils, and monocytes. A deficiency has been related to less effective immune responses against infections, vulnerability for the heart and blood vessels to damage, and increased virulence. Excessive intake is also associated with negative immune function.

· Your genotype is associated with lower serum copper



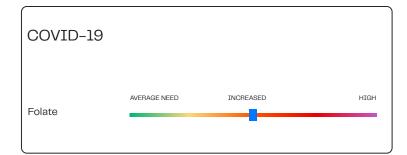
Suboptimal levels of iron are associated with decreased killer efficiency of NK cells and lymphocytes as well as with compromised cytokine production. Both iron uptake disturbances and metabolism are implicated in virulence of airway hospital-acquired infection and chronic respiratory infections. In contrast, excessive iron levels can generate harmful cellular toxicity, so their serum levels must be well regulated.

- Your genotype combination is associated with average serum iron levels
- A retrospective study based on 50 hospitalized Chinese subjects with confirmed COVID-19 demonstrated that 90% of these subjects had abnormally low serum iron concentrations
- Populations with lower iron status could be more prone to suffer a mild to severe (or critical) symptomatology of COVID-19 and the fact of monitoring patient iron levels has been proposed as a potential early marker to predict COVID-19 severity and mortality
- Iron levels should always be monitored by your practitioner



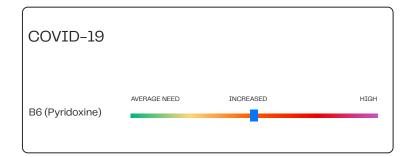
Sufficient vitamin B12 intake is essential for antibody production and a deficiency is related to a lower concentration of circulating lymphocytes and altered antibody-based responses. SARS CoV-2 infection is related to an aggravation of the cellular metabolism and the homocysteine pathway causing severe complications from COVID-19, and the correct supply of vitamin B12, folate and B6 may be crucial for COVID-19 patients.

- Your genotype is associated with low B12 levels
- Some of the countries least affected by SARS-CoV-2 show the highest levels of vitamin B12 intake (Portugal and Finland)
- Some of the countries most affected by SARS-CoV-2 (Belgium and Spain) have intakes below the median



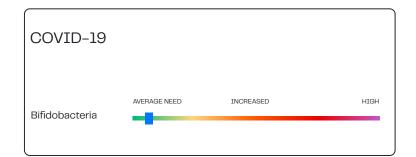
Folate is crucial for optimal Th-1 mediated immune response and proper antibody production. Suboptimal levels of folate may trigger imbalances in T and NK cell mediated immune responses and decrease the amount of antibody production.

- Your genotype combination is associated with a higher than average need for folate
- The correct intake of vitamin B6, folate and B12 in patients affected by COVID-19 has been proposed as part of the disease treatment, even by supplementation formulas, in an attempt to regulate the disruption of cellular metabolism of the homocysteine pathway caused by the SARS-CoV-2 infection



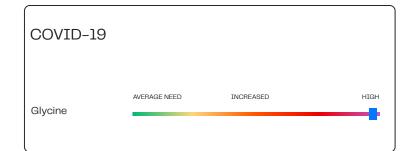
Vitamin B6 is essential for maintaining cytotoxic activity of NK cells, lymphocyte development, and B-cell antibody production. Suboptimal intake is associated with lower concentrations of circulating lymphocytes, impaired lymphocyte maturation, and decreased antibody-based responses.

- Your genotype is associated with low serum B6 levels, requiring a higher than average intake of B6
- The correct intake of vitamin B6, folate and B12 in patients affected by COVID-19 has been proposed as part of the disease treatment, even by supplementation formulas, in an attempt to regulate the disruption of cellular metabolism of the homocysteine pathway caused by the SARS-CoV-2 infection



Approximately 80% of your immune system is in your gut. The good bacteria bifidobacterium is highest in breast-fed infants and has been found to be lower in the higher-risk demographics for COVID-19 including those with diabetes, obesity, asthma and the elderly. Bifidobacteria populations have been found to vary based on the FUT2 genotype.

- Your genotype is associated with improved bifidobacteria levels in the gut, helping to protect against lower and upper respiratory infections
- Prebiotics found in foods like bananas, garlic, leeks, barley, asparagus, pistachios, onions, and polyphenol-rich foods – have been found in human trials to increase bifidobacteria levels



Glycine is one of the three major amino acids for glutathione production, protecting the body from oxidative damage during the immune response, and supporting T-cell proliferation.

- Your genotype is associated with low type 1 collagen production, increasing your glycine requirement
- Type I collage is a major structural protein in the lung and is stimulated during certain inflammatory reactions in the lung
- Collagen protein, bone, or chicken broth that gelatinizes, gelatin, meat with the skin, ribs, shanks, and drumsticks are all ways to increase dietary glycine
- Baobab is considered an exceptionally good source of plant-based glycine found in the hunter-gatherer Hazda diet





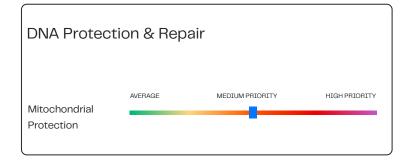
Glutathione is the master antioxidant system involved in oxidative stress, detoxification, and immunity. Glutathione status parallels telomerase activity, an important indicator of lifespan.

- Your genotype combinations are associated with decreased baseline glutathione levels
- Glutathione decreases with age, and low levels of glutathione are associated with chronic exposure to chemical toxins, heavy metals and excess alcohol, immunocompromised conditions, and neurodegenerative disorders
- Glutathione has been found to increase by 20% with deep breathing practices like Tai Chi or yoga
- For exercise, a combination of aerobic exercise and circuit weight training produced the highest glutathione effect
- Selenium, glycine, cysteine, vitamin C, and cruciferous vegetables all improve glutathione levels
- Chicken or bone broth, herbs, and spices are some of the best dietary ways to maintain higher levels of glutathione
- Some of the all-stars include cinnamon, anise, sage, and thyme due to also containing the antiviral compound caffeic acid



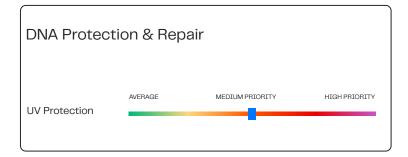
CAT makes an enzyme called catalase, which helps reduce oxidative stress. CAT is present in all aerobic cells while research has found the highest correlation to prostate, breast, liver, and blood health.

· Your genotype is associated with improved catalase levels



The SOD2 gene is responsible for superoxide dismutase levels, an important protector of the mitochondria, the powerhouse of the cell.

- Your genotype is associated with slightly reduced mitochondrial protection
- Manganese, boron, vitamin A, C, E, omega-3 fatty acids, CoQ10, lutein, lycopene, milk thistle, cordyceps, holy basil, reishi and cryotherapy all increase mitochondrial protection



One hypothesis for variants in MTHFR 677 is that they were selected based on higher folate intake and UV exposure, both common in Mediterranean climates. What happens in the body when MTHFR enzymatic function is reduced is that thymidine production increases. Thymidine enhances the repair of UV-induced DNA damage to help quickly repair sun damage.

- Your MTHFR genotype is associated with moderate UV protection from the sun
- To improve UV protection, increase your intake of folate-rich greens, blackberries, wild salmon, cacao powder, schisandra, reishi, dill and dried parsley

DNA Protection & Repair AVERAGE MEDIUM PRIORITY HIGH PRIORITY Skin Protection

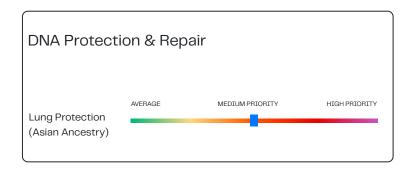
GPX1 activity is considered to be the most important antioxidant enzyme defense mechanism in the skin.

- Your genotype is associated with slighly lower antioxidant protection for the skin
- Vitamin C, niacin, zinc, selenium, dandelion root, ashwagandha, and dark roast coffee have all been found to reduce the risk of melanoma



Variants in the GSTP1 gene have been associated with lower antioxidant support in the lungs when exposed to environmental pollution.

 Your genotype is associated with increased lung protection against environmental pollutants



Glutathione levels and pro-inflammatory cytokines, such as TNF-a, are found in airways and environmental pollutants are known to induce inflammatory responses.

- Your genotype combination is associated with reduced lung protection against environmental pollutants
- It is recommended to increase your intake of cruciferous vegetables, vitamin C, vitamin E, vitamin A, milk thistle, resveratrol, curcumin, green tea, and white tea to target the GSTM1 gene
- Additional support includes cold water immersion, breathing exercises, cordyceps, and ginger to target the TNFA gene



The MLH1 gene codes for a DNA repair enzyme linked to colon health.

 Your genotype is associated with improved DNA protection for colon health



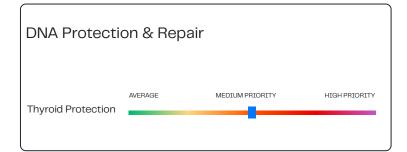
A large-scale genome-wide analysis of over 18,000 people from the U.S., Canada, Australia and Europe found that variants in GATA3 were associated with an increased risk of colon cancer for those eating processed meat compared to those with the normal genotype.

 Your genotype is not associated with an increased risk of colon cancer from cured meat consumption

DNA Protection & Repair AVERAGE MEDIUM PRIORITY HIGH PRIORITY Eye Protection

Blue light is a high-energy or short-wavelength visible light from your phone and computer that induces inflammation and retinal diseases such as agerelated macular degeneration and retinitis pigmentosa.

- Your genotype is associated with higher requirements for foods high in lutein, zeaxanthin, and anthocyanins for eye health
- A meta-analysis found that the rates of myopia (nearsightedness) will increase 140% by 2050 due to our increased time in front of a screen
- Research has found that bilberry and lingonberry exert protective effects against blue LED light-induced retinal photoreceptor cell damage due to their polyphenol content
- Increase your dietary intake of dark purple berries, dark leafy greens, summer squash, green peas, broccoli and Brussels sprouts



Up to 60 percent of those with a thyroid disorder are unaware of their condition. The cause is considered largely unknown and occurs 10 times more in women than in men. Hashimoto's disease runs in the family and 70%–80% of susceptibility to autoimmune thyroid disease is based on genetics.

- Your genotype combination is associated with reduced thyroid protection and a slightly increased risk of Hashimoto's disease
- · Autoimmune thyroid disease is associated with celiac disease
- A deficiency in selenium is associated with celiac disease and thyroid disease, and plays a significant role in thyroid hormone synthesis, secretion and metabolism
- Sucralose and glyphosate destroy gut flora like lactobacillus, which disturbs selenocysteine levels present in the catalytic center of enzymes that protect the thyroid from free radical damage
- The artificial sweetener aspartame in the body further metabolizes to formaldehyde, and rat studies found that formaldehyde (as a metabolite of aspartame) caused increased TSH levels and worsens the capacity of the gland leading to thyroid failure



Multiple genes are linked to DNA protection for pancreatic health.

 Your genotype combination is associated with increased DNA protection for pancreatic health



Certain gene combinations have been found to decrease the detoxification ability of certain toxins found to be carcinogenic for the bladder.

- Your genotype is associated with decreased DNA protection for bladder health
- Avoid tobacco smoke, commercial hair dyes, working in industrial and manufacturing plants, charred meat, and diesel exhaust
- Increase your cruciferous vegetables, carotenoid, and vitamin C intake





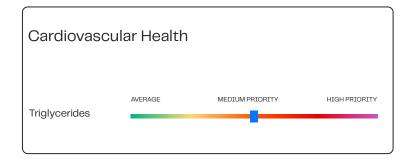
ApoE is connected to HDL and LDL levels, while PON1 is involved with supporting HDL function and LDL oxidation, an important mechanism in atherosclerosis and heart disease.

 Your genotype combination is associated with a higher likelihood of good HDL levels and a lower likelihood of higher levels of LDL, oxidized LDL, and total cholesterol



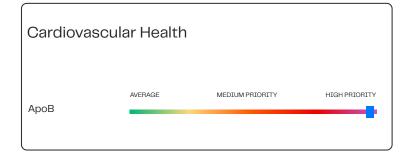
Subjects with variants in PPAR-alpha have been found to have a larger waist circumference and a higher proportion of small, dense LDL particle size.

 Your genotype is associated with a higher proportion of small, dense LDL particle size if following a high saturated fat and low polyunsaturated fat diet



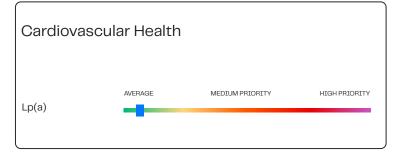
Variants in the FADS1 SNP (rs174546) are associated with elevated trigly ceride levels. $\label{eq:control}$

- · Your genotype is associated with elevated triglycerides
- Numerous studies have found that omega-3 fatty acids administered as fish oil supplements lowers plasma triglyceride levels by 25% to 34%



ApoB is a protein that is involved in the metabolism of lipids and is the main protein constituent of lipoproteins. High levels of ApoB, especially with the higher LDL particle concentrations, are the primary driver of arterial plaque. The PPAR-alpha polymorphism has been associated with ApoB in many populations such as Caucasians, Indians, and African-Americans.

- Your genotype is associated with elevated $\ensuremath{\mathsf{ApoB}}$ levels
- Lowering saturated fat intake and increasing polyunsaturated fat intake is recommended
- PPAR-alpha can be targeted with astaxanthin (high in wild salmon), pterostilbene (blueberries, mulberries, cranberries, raw almonds), genistein (fermented soy), tomatoes, cinnamon, zinc, Lion's Mane mushroom, Gynostemma tea and L-carnitine



 $\label{local_local_local} \mbox{Lp(a) is a sticky form of LDL that appears to affect plaque growth, LDL particle size and increase the risk of plaque rupture and blood clotting.}$

· Your genotype is not associated with elevated Lp(a) levels



The risk of heart attacks and cardiovascular disease conferred by the 9p21 gene appears to be modified by a prudent diet high in raw vegetables and fruits for South Asian, Latin American, Arab, Chinese and European populations for variants in rs4977574.

 You have a higher than average requirement for raw fruits and vegetables to maintain a healthy heart



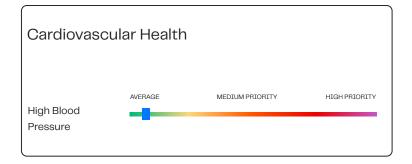
The uncoupling of nitric oxide has been linked to play an essential role in cardiovascular pathologies including dilated cardiomyopathy, ischemia-reperfusion injury, endothelial dysfunction, atherosclerosis, and hypertension.

- Your genotype combinations are associated with a higher than average need for folate to produce adequate BH4, the precursor to nitric oxide
- BH4 is depleted by high blood sugar, high omega-6 intake, chronic stress, high levels of mercury, arsenic, lead and aluminum, aspartame, and oxidative stress
- Other strategies to increase BH4 include vitamin C, L-arginine, B6, magnesium, and selenium



Homocysteine is a non-protein amino acid that is produced from methionine, can be recycled back into methionine and converted into cysteine in the methylation cycle. High homocysteine levels have been connected to depression, blood clots, inflammation, macular degeneration, dementia, and cancer.

 You have a higher than average need for folate to maintain healthy homocysteine levels



Age-related increases in blood pressure have been observed in almost every population, except among hunter-gatherers and farmers. High physical activity, low-stress levels, and potentially protective diets high in fruits, vegetables, potassium, lower in calories, salt, and alcohol are the major contributing factors to the stark differences.

 Your genotype combination is associated with normal blood pressure levels



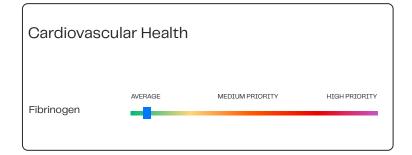
Deep vein thrombosis is a condition that occurs when a blood clot forms in a vein deep inside a part of the body and is most common for those over 60. Variants in F5 increase the risk of deep vein thrombosis.

Your genotype is not associated with deep vein thrombosis



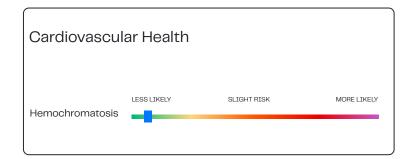
Variants in the ABCG2 gene have been associated with elevated uric acid levels and an increased risk of gout in Asians, Europeans, African Americans, Mexican Americans, and American Indians. Epidemiological studies have shown that uric levels are positively correlated with gout, hypertension, atherosclerosis, atrial fibrillation, and heart failure.

- · Your genotype is associated with increased uric acid levels
- The association with your genotype and elevated uric acid levels is significantly stronger in men, postmenopausal women, and hormone therapy users
- Your genotype increases your sensitivity to sugar, alcohol intake, highpurine foods like liver, and dehydration with the development of gout
- Avoid refined sugar, increase water intake, parsley, quercetin, kaempferol, olive oil and vitamin C



Fibrinogen is a soluble protein in the plasma that is broken down to fibrin by the enzyme thrombin to form clots. Fibrinogen serves to stop excessive bleeding, but in high amounts can cause blot clots.

• Your genotype is not associated with elevated fibrinogen levels



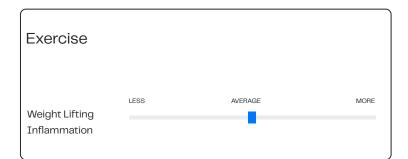
A homozygous HFE C282Y may lead to an iron overload due to increased iron absorption and disrupted metabolism. People who are homozygous for the HFE C282Y gene mutation comprise 85 to 90 percent of those with hemochromatosis in whites of western European descent.

 Your wild-type HFE genotype is associated with a reduced likelihood of genetically linked hemochromatosis



ACTN3 is currently the most promising gene for predicting the likelihood of becoming an Olympic level sprint and power athlete in males and females. The RR (CC) genotype expresses the ACTN3 protein found in Type II muscle fibers, which produces explosive and powerful contractions.

- You have the RX genotype for the ACTN3 gene associated with enhanced improvements in strength, power and muscle hypertrophy
- · More powerful muscle contractions
- · Higher muscle hypertrophy response
- Faster recovery



Weight lifting leads to a variation in muscle inflammatory markers based on genetics and intensity.

- Your genotype combination is associated with average levels of muscle inflammation (creatine kinase) for weight lifting
- To accelerate recovery, ice baths, whey protein, American ginseng, curcumin, vitamin C, and collagen protein have all been found to attenuate creatine kinase levels



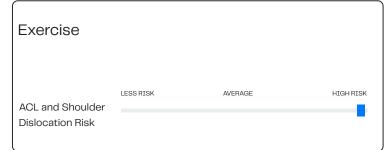
Endurance training leads to a variation in creatine kinase levels based on genetics.

 Your genotype combination is associated with average levels of muscle inflammation (creatine kinase) for endurance exercise



High-intensity exercise is defined as 70% to 85% of your maximum heart rate, and inflammation variation has been associated with the SOD2 gene.

 Your SOD2 genotype is associated with less muscle inflammation in response to high-intensity exercise



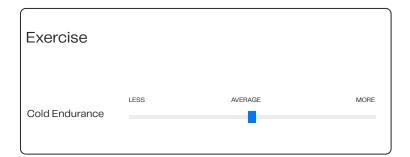
The COL1A1 gene is associated with ACL and shoulder injury risk.

- Your COL1A1 genotype is associated with an increased need for dietary collagen to prevent ACL and shoulder injuries
- Vitamin C, zinc, copper, glycine, proline, lysine, and B6 are all precursors to collagen production



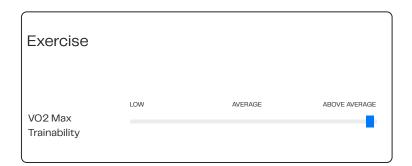
The ACTN3 gene is linked to increased or decreased risk of ankle and hamstring injuries.

 Your ACTN3 genotype is associated with an average risk of ankle and hamstring injuries



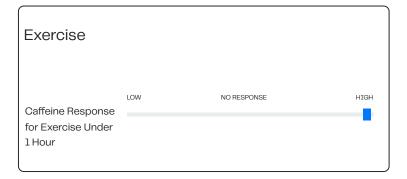
The ACTN3 gene is associated with a lower or higher adaptation rate to cold endurance.

 You have the ACTN3 RX genotype, associated with an average adaptation rate to cold endurance



VO2 max is measurement for oxygen capacity and endurance training. The most recent research has shown that 97 genes predicted VO2 max trainability. The phenotype is dependent on several of these variants combined together, which may contribute to approximately 50% of an individual's VO2 max trainability.

 Research has shown that your genotypes for multiple genes are associated with a higher VO2 max training response



The CYP1A2 gene is associated with caffeine response for improving or decreasing exercise performance.

 Fast metabolizers have been found to show an increase in exercise performance for activities under 1 hour from caffeine consumption



The CYP1A2 gene is associated with caffeine response for improving or decreasing exercise performance.

 Your CYP1A2 genotype is associated with a decrease in exercise performance activities over 1 hour from caffeine consumption due to being a fast metabolizer

MY HEALTH REPORT: STRENGTHS

This section is a thorough overview of your individual gene function across the entire analysis in just a few pages. If you are looking for a brief summary of the most important parts of your report without doing a deep dive into the genotype tables and clinical research sections, this is the place to start. Be proud of your inherent genetic strengths!

NUTRIENT METABOLISM & DIGESTION

- — **Prebiotics, Probiotics and B12-FUT2** The GG FUT2 genotype in European, African, and Indian populations is associated with improved bifidobacteria populations in the gut compared to the AA genotype, increasing immune function against respiratory infections.
- **Vitamin C-SLC23A1** Your genotype is associated with improved whole-body vitamin C homeostasis through dietary absorption and renal reabsorption.
- Adiponectin-ADIPOQ Your genotype is associated with a higher probability of normal adiponectin levels, linked to improved bodyweight, insulin, and glucose levels.
- Iron Your genotype is associated with a lower risk of iron overload for the HFE C282Y gene.
- **Saturated Fat-PPAR-alpha** While your heterozygous genotype is associated with abnormal lipid metabolism and a poor response to the ketogenic diet, it has also been found to have lower C-reactive protein levels, a marker of inflammation.
- **Ghrelin and Appetite-FTO** Your genotype is associated with normal ghrelin levels (hunger hormone), decreasing the risk for overeating and abdominal weight gain.
- Saturated Fat-APOA2 Your genotype is associated with a reduced likelihood of saturated fats causing weight gain.
- Lactose You have the heterozygous AG genotype that is associated with a lower probability of lactose intolerance.
- Histamines-APB1 You have the wild-type genotype that is associated with improved histamine breakdown in the digestive tract.
- **Ethanol Metabolism-ALDH2** Your genotype is less likely to experience facial flushing from alcohol due to improved acetaldehyde metabolism.
- **Stress and IBS-ADRB2** You have the ADRB2 CC wild-type genotype that is associated with a reduced percentage of digestive disorders, IBS, and anxiety related to adrenaline levels.

METHYLATION

- **Folate-MTHFR 1298** Your genotype is associated with improved BH4 levels and neurotransmitter function. Healthy BH4 levels assist in the management of cardiovascular health, mental health, and digestive health.
- **Folate-DHFR** Your genotype is associated with an improved breakdown of synthetic folic acid at the beginning of the folate cycle. However, variants in MTHFR 677 can also affect folic acid metabolism.
- B12, B2 and Zinc-MTR You may have improved MTR function, assisting homocysteine metabolism.
- •= B12-MTRR Your genotype is associated with improved gene function, assisting B12 and homocysteine metabolism.
- **B12-TCN2** Your genotype is associated with improved B12 transportation.
- **Arsenic-CBS** Your genotypes are associated with improved arsenic metabolism and detoxification for the CBS genes.

HORMONE SUPPORT

- Thyroid-DI01 Your genotype is associated with average DI01 gene function for T3 and T4 thyroid function, however other epigenetic factors should be assessed.
- Thyroid-DI02 Your genotype is associated with average T3 and T4 thyroid function in the brain for the DI02 gene. However, other factors can affect T3 and T4 levels including thyroid surgeries.
- **Estrogen Metabolism-CYP1A1** Your CYP1A1 wild-type genotype is improved for the beginning phase of estrogen metabolism. Please review all genes involved in estrogen metabolism for a complete picture of the process.
- **Estrogen Metabolism-CYP1A2** For men and women with the CYP1A2 AA genotype, coffee intake was found to be more protective against estrogen receptor-positive breast cancer and prostate cancer.
- **Estrobolome-FUT2** Your wild-type genotype is associated with improved bifidobacteria gut bacteria, assisting the gut phase of estrogen detoxification.

MENTAL HEALTH, NEUROTRANSMITTERS & COGNITIVE PERFORMANCE

- Serotonin Receptor-Memory You have the wild-type genotype that is associated with an improved episodic memory,
 which is the ability to recall details regarding personal experiences, names of people, specific events, and what exactly
 occurred.
- **Dopamine, Adrenaline and Estrogen-COMT** The homozygous COMT AA V158M genotype is associated with increased creativity, divergent thinking, problem-solving, better memory (in men only), executive function, and overall cognitive function.
- **Dopamine Receptors-ANKK1** Your genotype is associated with an improved density of dopamine receptors for healthy dopamine levels.
- Histamines and Migraines-DAO The wild-type CC genotype for DAO rs1049793 is associated with a reduced risk of histamine-induced migraine headaches.
- **Glutamate-BDNF** Your genotype is associated with improved glutamate modulation, brain repair, spatial learning, memory, and adaptability.
- **Glutamate Transport-SLC17A7** Your genotype is associated with improved recovery from head injuries. However, your APOE and BDNF genotype should also be assessed because these all have a cumulative impact.
- **Cholesterol-APOE** You have the ApoE e3/e3 genotype, improving cholesterol transport and the maintenance of brain neurons. The ApoE e3 allele improves cognitive fitness, HDL and LDL profiles, viral protection, and the response to plant bioactive compounds.

IMMUNE SUPPORT, ANTIOXIDANTS AND INFLAMMATION

- **Cell Protection-CAT** Your genotype is associated with improved catalase levels, mitigating damage to your cells.
- Glutathione-GSTM1 While the GSTM1 null genotype has been associated with a greater sensitivity to benzo(a)pyrene, there is also a benefit to this genotype. The benefit is that the null genotype may retain a higher level of isothiocyanates, the anti-cancer compounds found in cruciferous vegetables that may also be required in higher amounts for this genotype.
- Glutathione-GSTP1 While the homozygous GG genotype for GSTP1 rs1695 is associated with a higher sensitivity to
 heavy metals, one advantage may be an increased VO2 max response from endurance training compared to the wildtype genotype.
- Heavy Metals-GSTP1 You have the wild-type CC genotype for GSTP1 rs1138272 that is associated with improved glutathione antioxidant protection against heavy metals, pesticides, and air pollution for colon, prostate, lung, throat, and fertility health. Your GSTP1 rs1695 genotype may increase or decrease this effect.
- Nitric Oxide-NOS1 Your genotype is associated with an average required intake of red, yellow, and orange vegetables to modulate the inflammatory process for NOS1.
- **Nitric Oxide-NOS2** Your NOS2A gene is functioning optimally for reducing the probability of age-related macular degeneration from cigarette smoke.
- **Eye Health-ARMS2** Your genotype is associated with a lower sensitivity to the negative effects of smoking on eye health.

DETOXIFICATION

- **Liver Enzyme-CYP1A1** Your genotype is associated with improved detoxification of benzopyrene from cigarette smoke and will assist the function of your GSTM1 gene.
- **Liver Enzyme-CYP2D6** Your genotype is associated with improved metabolism of certain drugs associated with CYP2D6 rs1065852. However, more CYP2D6 SNPs are needed for a complete panel. Please talk to your doctor about further testing for CYP2D6 and drug metabolism.
- **Liver Enzyme-CYP2E1** Your genotype is associated with improved metabolism of benzene and acrylamide for colon health.
- **Liver Enzyme-CYP3A4** Your genotype is associated with normal metabolism of certain drugs that use this enzyme. We recommend further pharmacogenomic testing with your doctor for more information regarding CYP3A4.

DNA DAMAGE, PROTECTION AND REPAIR

- __ DNA Repair-ATM Your genotype is associated with improved DNA repair function for pancreatic health.
- Breast-ESR2 For women with the AA ESR2 rs2987983 genotype, your genotype is associated with improved tumor suppression function for breast health. All genes related to breast health should be analyzed to better determine the cumulative value for breast protection.
- DNA Repair-MDM2 Your MDM2 genotype is associated with improved DNA repair for sun damage if you are female.
- **DNA Repair-MLH1** Your genotype is associated with improved DNA repair for colon, endometrium, lung, and brain protection.
- **Processed Meat and Colon Cancer-GATA3** You have the wild-type genotype that is associated with a reduced risk of processed meat consumption and colon cancer.
- Longevity-SIRT1 Your SIRT1 genotype is associated with normal SIRT1 activity for longevity. While not a weakness, you may want to increase SIRT1 activity epigenetically to increase the probability of longevity, especially if you have the APOE-e4 allele. A sedentary lifestyle, aging, poor diet, and obesity lowers SIRT1 activity. Exercise, fasting, 7-8 hours of sleep per night, saunas, polyphenols, vitamin D, omega-3 fatty acids, resveratrol, magnesium, and melatonin have all been found to increase SIRT1 activity.

CARDIOVASCULAR HEALTH & EXERCISE

- Power and Recovery-ACTN3 You have the RX genotype associated with enhanced improvements in strength, muscle
 hypertrophy, sprint times, protection from eccentric training-induced muscle damage, and a reduced risk of sports injury.
 The RX genotype may represent the best of both ACTN3 genotypes for strength training, maintaining lean muscle mass
 later in life, and longevity.
- **Lung Cytokines-TNFA** If you have Asian ancestry, your genotype is associated with improved TNF-a gene function for lower inflammation in the lungs.
- **Pesticides, HDL and LDL-PON1** You have the wild-type genotype associated with improved PON1 activity for pesticide detoxification and protection against LDL oxidation.
- **LDL-LPA** Your genotype is associated with healthy Lp(a) levels, a sticky form of LDL that affects plaque levels.
- **Caffeine-CYP1A2** You have the homozygous AA genotype and are a "rapid metabolizer" of caffeine. This means that caffeine will quickly be metabolized from your body and the effects lasting a shorter period of time. Variants in COMT can increase the sensitivity to catecholamines in coffee, and oral contraceptives can slow down caffeine metabolism.
- Fibrinogen-ESR2 Your genotype is associated with improved fibrinogen levels.
- **Blood Clots-F5** Your genotype is associated with improved gene function for a lower probability of deep vein thrombosis.
- **Blood Pressure-ACE1** Your genotype is associated with intermediate baseline ACE levels. If you are female, ACE levels may be lower. Depending on ACE2 levels, you may have a more balanced renin-angiotensin system for blood pressure.
- **Blood Pressure-ACE2** Your genotype is associated with higher baseline ACE2, improving the balance between ACE1 and ACE2 for blood pressure, and potentially lowering the risk of COVID-19 severity. Other dietary habits and health issues could affect this result.
- **Phytoestrogens-TMPRSS2** You have the AG genotype that is associated with a lower expression of TMPRSS2 and may decrease the susceptibility to viral infections and prostate cancer (men).

MY HEALTH REPORT: WEAKNESSES

Genes are not your destiny - they are your blueprint. Please understand that these weaknesses can be turned into strengths based on the personalized recommendations given below. Making strategic changes to diet, environment, stressors, and even relationships can have a profound effect on optimizing gene function. Aim to turn every weakness into a strength by giving attention to the proactive, customized dietary and lifestyle modification recommendations in this section!

NUTRIENT METABOLISM & DIGESTION

- Beta Carotene to Vitamin A Conversion Rate-BCMO1 Your BCMO1 genotype combination is associated with a reduced conversion rate of plant-based beta carotene (squash, sweet potatoes, carrots) to vitamin A. This increases your need for foods higher in vitamin A like eggs, cod liver oil, wild salmon oil and organ meats for skin, digestion, healthy eyes, lungs, and immunity.
- ALA to EPA and DHA Conversion-FADS2 Your genotype is associated with a reduced conversion of plant-based omega-3 ALA (walnuts, flax seeds, and pumpkin seeds) to EPA and DHA. Increased EPA and DHA intake may be needed.
- **Prebiotics, Probiotics and B12-FUT2** The rs601338 FUT2 GG genotype has been associated with lower B12 levels in European, Indian and African populations.
- **B6-NBPF3** You are more likely to have low B6 levels due to variants in the NBPF3 gene, increasing the sensitivity to medications that deplete B6 (oral contraceptives, antibiotics, ACE inhibitors, antacids, proton pump inhibitors and more). You need to focus on increasing foods high in B6 like wild salmon, pistachios, avocados and potatoes.
- **Saturated Fat-PPAR-alpha** You have the heterozygous genotype that is associated with abnormal lipid metabolism and a poor response to the ketogenic diet. This is more pronounced if you are male and have multiple variants in the other fat metabolism genes. You may need to lower your saturated fat intake, increase your polyunsaturated fat intake, and focus on PPAR-alpha activators including astaxanthin, pterostilbene, cinnamon, tomatoes, zinc, Lion's Mane mushroom, Gynostemma tea, and increasing L-carnitine production.
- **Fat Metabolism-ACSL1** Your genotype is associated with higher fasting glucose levels from a higher saturated fat intake. If your fasting glucose is high and you have variants in the other fat metabolism genes, fatty red meat and dairy should be reduced and more focus should be on monounsaturated and polyunsaturated fats.
- **Carbohydrates-TCF7L2** Your genotype is associated with an increased probability of elevated blood sugar from refined sugar and grains. A diet low in refined sugar and flour, higher in protein and omega-3 fatty acids, glycine, diversified prebiotic foods, olive oil, cinnamon, turmeric, dark roast coffee and cordyceps mushrooms may be more beneficial.
- **Uric Acid-ABCG2** Your genotype is associated with a higher probability of chronically elevated uric acid levels, increasing the sensitivity to sugar consumption and dehydration with the development of gout. Avoid refined sugar, increase water intake, flavonoids, olive oil, and vitamin C.

METHYLATION

- **Folate-MTHFR 677** You have the heterozygous genotype that is associated with a reduced function of approximately 30%. This increases the need for riboflavin and methylfolate for normal homocysteine levels.
- •= Folate-MTHFD1 G1958A Your genotype is associated with an increased need for folinic acid, the second most common type of folate after methylfolate.
- **Choline-PEMT** Your genotype is associated with an increased need for dietary choline for liver health, normal homocysteine levels, breast health for women, and a healthy pregnancy for women.
- •= **B6-CBS** Your genotype is associated with reduced CBS gene function for homocysteine levels, gut repair, and brain health, increasing your need for B6.

HORMONE SUPPORT

- **Sex Hormone Binding Globulin** If you are female, your genotype is associated with a sensitivity to oral contraceptives and hormone replacement therapy in relation to sex hormone binding globulin (SHBG) levels for healthy hormone levels. If you are male, higher SHBG levels could affect bone mineral density and SHBG may need to be tested. However, variants may also positively lead to higher testosterone levels.
- **Vitamin D-CYP2R1** Your genotype is associated with low circulating vitamin D levels that can affect immunity, breast health in women, and testosterone levels in men. Check your vitamin D levels and make sure you are in range.
- **Estrogen Metabolism-CYP2C19** Individuals with the CC genotype for CYP2C19*17 are considered the normal metabolizer phenotype, which may lack the estrogen metabolism benefits of the ultra-rapid metabolizer phenotype. Please review all genes involved in estrogen metabolism for a complete picture of the process.
- **Estrogen Metabolism-COMT** For estrogen metabolism and detoxification, those with the slow AA COMT V158M genotype may have an increase in harmful estrogen metabolites that can cause DNA damage. To reduce the risk of these metabolites, you should avoid xenoestrogens, manage stress levels, maintain gut health, increase magnesium intake, and consume green tea polyphenols.
- MTNR1B-Melatonin You have the CG MTNR1B genotype, which is associated with delayed melatonin release, a longer duration of morning melatonin levels, and less insulin release during late night and early morning feeding. It is recommended to eat dinner early, avoid late night snacking and consume breakfast later in the morning for better glycemic control.

MENTAL HEALTH, NEUROTRANSMITTERS & COGNITIVE PERFORMANCE

- Serotonin Receptor-Stress The 5-HT2A heterozygous genotype may be more impactful in females who also have variants in the BDNF gene. Chronic stress may increase the susceptibility to anxiety, depression, OCD, and IBS for these genotypes. If you experience higher perceived stress and chronic stress levels, you may require more aerobic exercise, cognitive behavioral therapy, mindfulness training, meditation, yoga, singing, prebiotics, lactobacillus helveticus, bifidobacterium longum, tryptophan, green or black tea, and B-vitamins.
- **Dopamine, Adrenaline and Estrogen-COMT** The homozygous AA COMT V158M genotype is associated with a slower breakdown of dopamine, adrenaline and estrogen, creating higher circulating levels in response to stress. This may increase your need for magnesium, vitamin C, strength training, and sprints to reduce stress levels.
- Histamines and Hyperactivity-HNMT You have the AA genotype that is associated with increased hyperactivity in response to food dyes and sodium benzoate. Excess histamine is lowered by vitamin C, choline, folate, magnesium, chamomile, basil, stinging nettle, echinacea, fennel, ginger and wild oregano.
- Anandamide-FAAH You have the common CC genotype that encodes for the fast activity of FAAH. This is associated with naturally lower anandamide levels that could increase anxiety, pain, pesticide sensitivity and a heightened stress response to threatening situations. You may benefit from aerobic exercise over 30 minutes (especially in altitude), CBD oil, red clover tea (women), kaempferol (raspberries, capers, cumin, cloves, almonds, cherry tomatoes, red wine), cacao, echinacea, rosemary, and hops to increase anandamide levels.
- •= Brain Health-PEMT Your genotype is associated with an increased need for dietary choline and daily walks for memory, anxiety, and REM sleep.

IMMUNE SUPPORT, ANTIOXIDANTS AND INFLAMMATION

- **Cell Protection-SOD2** You have the heterozygous AG genotype for SOD2. Your mitochondria (powerhouse of the cell) may have a higher sensitivity to glyphosate, fluoridated water, chronic stress, poor sleep, and shallow breathing. Increase foods that contain manganese, lycopene, and vitamin C, milk thistle, mushrooms like reishi and cordyceps, and exercise that encourages deep breathing.
- Glutathione-GSTM1 You have the null genotype that is associated with a higher sensitivity to benzo(a)pyrene from the burning of wood or trash, tobacco smoke, asphalt, coal, diesel exhaust, charred meat, and gas cooking. If you have the GSTM1 null and NAT2 slow acetylator combination, that may affect lung, breast, bladder, skin, colon, and kidney health. It is recommended to increase your intake of cruciferous vegetables, vitamin C, vitamin E, vitamin A, milk thistle, resveratrol, curcumin, green tea, and white tea.
- Glutathione-GSTP1 You have the homozygous GG genotype for GSTP1 rs1695 that is associated with a higher sensitivity to mercury, cadmium, arsenic, pesticides, and air pollution for breast, prostate, urinary, esophagus, and skin health. Your GSTP1 rs1138272 genotype may increase or decrease this sensitivity. Selenium, vitamin C, vitamin E, milk thistle, and cruciferous vegetables all assist GSTP1 gene function.
- **Glutathione-GPX1** Your genotype is associated with a higher need for selenium to combat oxidative stress and less tolerance to heat stress. Lower glutathione peroxidase increases the sensitivity to oxidative stress from low or high iron levels, statin drugs, thyroid damage, sun damage, and dietary or environmental lead exposure. Selenium, cold exposure, optimizing testosterone levels in men and estrogen in women, and adequate vitamin C, vitamin E, milk thistle, ginger, cumin, anise, fennel, caraway, and cardamom intake are all ways to assist GPX1.
- Glutathione-CTH Your genotype may increase your need for cysteine-rich foods including eggs, meat, yogurt, garlic, and sunflower seeds for healthy homocysteine and glutathione levels. Other genes related to homocysteine and glutathione should also be reviewed.
- •= Eye Health-CFH Your genotype is associated with an increased need for lutein, zeaxanthin, bilberry, lingonberry, vitamin C, and vitamin E for healthy eyes.

DETOXIFICATION

- **Liver Enzyme-CYP1A2** You have the AA genotype for CYP1A2 that is associated with a higher sensitivity to heterocyclic amines (fried meat) depending on if you have the homozygous null GSTM1 genotype or the NAT2 slow acetylator genotype. Marinades, unfiltered fermented drinks (Kombucha, beer, wine), cruciferous vegetables, parsley, and spinach have all been found to reduce the carcinogenic effect of heterocyclic amines.
- **Liver Enzyme-CYP1B1** You have the CG genotype for CYP1B1 that is associated with a slightly reduced detoxification of polycyclic aromatic hydrocarbons (highest in vegetable oils), oral contraceptives, cigarette smoke, an increased sensitivity to excessive sun exposure, and high-dose biotin supplementation. You can assist CYP1B1 with seaweed, celery, berries, rooibos tea, red wine, and dark roast coffee.
- **Liver Enzyme-THC and CYP2C9** You have the heterozygous genotype associated with a reduced clearance of THC, the psychoactive compound in cannabis. While the heterozygous genotype does not affect clearance as much as the homozygous genotype, this could create a higher sensitivity to THC, especially in the edible form.
- Aromatic Amines-NAT2 You have the slow acetylator genotype for the NAT2 gene. This is associated with reduced detoxification of aromatic amines found in tobacco smoke, commercial hair dyes, industrial and manufacturing plants, charred meat, and diesel exhaust for bladder, prostate and breast health. Cruciferous vegetables, carotenoids, and vitamin C all assist NAT2 detoxification.
- **Vitamin K2-VOKRC1*2** Your genotype is associated with a higher sensitivity to vitamin K2 induced deficiency from antibiotics and the blood thinner Warfarin.
- = Statins-COQ2 Your genotype is associated with a higher likelihood of statin drug-induced muscle pain.

DNA DAMAGE, PROTECTION AND REPAIR

• **DNA Repair-TP53** - You have the heterozygous CG genotype that may be advantageous for fertility in cold climates, but also increases the need for selenium, zinc, vitamin C, reishi, and niacin for DNA repair against chemical toxicity to the thyroid gland and skin.

CARDIOVASCULAR HEALTH & EXERCISE

- **VO2 Max-PPARGC1A** Your genotype is associated with a higher need for more strategies to increase oxygen capacity for aerobic exercise, including a structured endurance program, cold exposure, and adaptogens. Your genotype in the GSTP1 rs1695 gene can also influence this result.
- **Muscle Recovery-IL6** You have the CG genotype that is associated with higher levels of creatine kinase a marker of muscle damage from workouts. To accelerate recovery, whey protein, cold water immersion, American ginseng, curcumin, allicin, optimal testosterone levels, vitamin C, and collagen protein have all been found to attenuate creatine kinase levels.
- **Muscle Injury-COL1A1** You have the wild-type CC genotype that is associated with an increased need for dietary collagen for healthy skin, tendons, corneas, lungs, and bones. Vitamin C, zinc, copper, glycine, proline, lysine, and B6 are all precursors to collagen production.
- •= Raw Fruit and Vegetable Intake-9p21 You have the homozygous genotype that is associated with an increased need for phytonutrients from a higher raw fruit and vegetable intake for a healthy heart.
- **Triglycerides-FADS1** Your genotype is associated with a higher need for EPA and DHA omega-3 fatty acids to maintain healthy triglyceride levels.
- **Stress-ADRB2** You have the homozygous AA genotype for ADRB2 rs104271 that is associated with a higher adrenaline response from stress that could affect heart and GI health, and has been linked to TMJ and fibromyalgia. COMT gene function should also be reviewed. Optimize COMT function and ADRB2 with foods that contain magnesium and vitamin C, deep breathing, and consider adaptogens to lower the stress response.
- **Potassium and Magnesium-ADD1** If you have Asian ancestry, your genotype is associated with an increased risk of a higher sodium intake causing elevated blood pressure. Increasing potassium, vitamin D, magnesium, calcium, garlic, and omega-3's all lower blood pressure.

YOUR PERSONALIZED DNA-BASED GROCERY LIST

This section of the report represents the most expansive, actionable summary of what you can do, right now, to dramatically up-regulate gene function, building a happier, healthier you! No technical expertise is required - just make these recommendations non-negotiable when you visit the grocery store.

Your grocery list is generated based on a combination of unique gene variants that require an increased intake of the following vitamins, minerals, phytonutrients, amino acids, fiber and more. This list generates the foods and drinks based on the highest levels for each section and does not take into account any food allergies or sensitivities.

|--|--|--|

B12

Seafood, meat, dairy (if consuming dairy) and unfiltered fermented drinks



В2

Lamb, salmon, yogurt, liver and oyster mushrooms



В6

Wild salmon, yellowfin tuna, liver, chicken breast, unfiltered fermented drinks, pistachios, avocado, sweet potatoes, and spinach



Beta-Carotene

Sweet potatoes, carrots, spinach, squash, cantaloupe, and broccoli



Betaine

Spinach, shrimp, beets, and whole grain sourdough bread



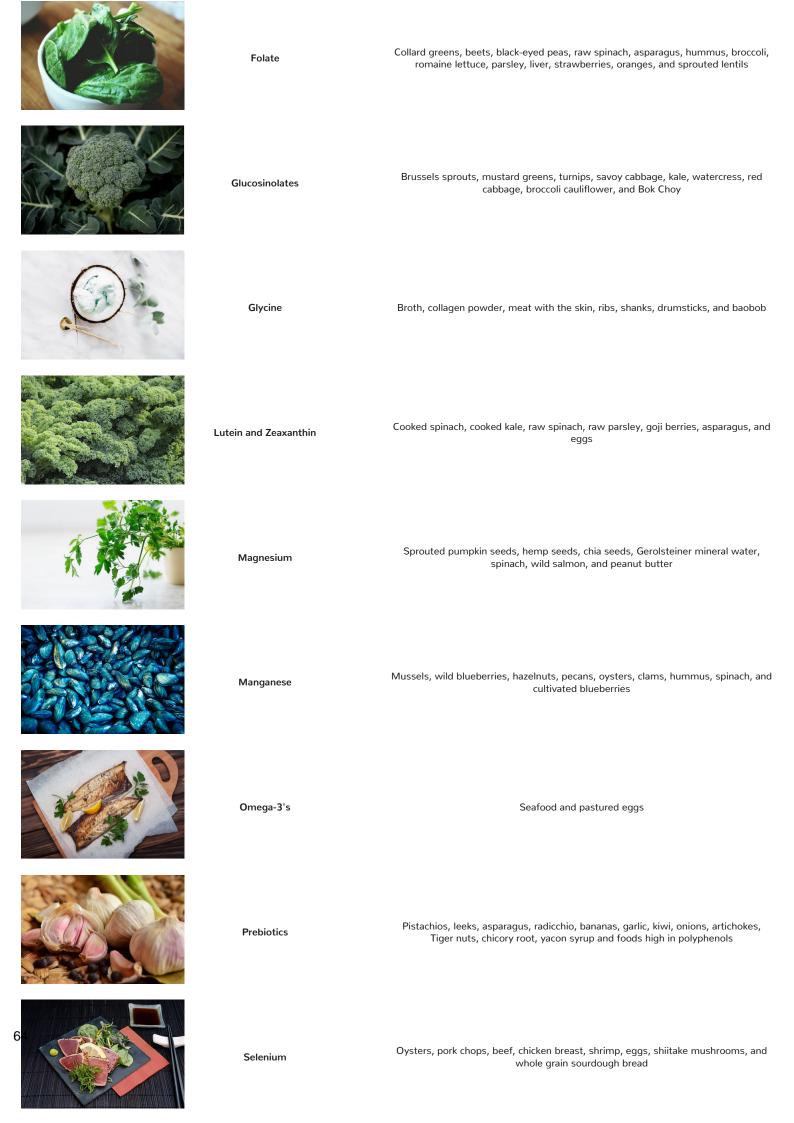
Choline

Pastured eggs, beef round, liver, heart, chicken, wild cod, bacon, and edamame



Copper

Potatoes, shiitake mushrooms, cashews, sunflower seeds, dark chocolate, and shellfish





Vitamin A

Liver, pastured eggs, cod liver oil, wild salmon oil, eel, and sockeye salmon



Vitamin C

Bell peppers, guava, black currants, strawberries, oranges, and broccoli



Vitamin D

Sockeye salmon, cod liver oil, canned tuna, wild herring, and sardines



Vitamin E

Sunflower seeds, almonds, avocado, spinach, butternut squash soup, and olive oil

PERSONALIZED BLOOD WORK

These results are generated based on a combination of gene variants unique to you. These biomarkers may not be out of range based on your diet and lifestyle habits, but they may be the ones for you to monitor to ensure you are making the right choices based on your genetic results (your predispositions).

For example, if vitamin D comes up in this section, it does not mean that your current levels of vitamin D are actually low. What we are saying is that based on a variety of genetic factors, your variants could make it more difficult to obtain recommended levels of circulating vitamin D, so it might be prudent to further monitor to ensure that you are taking the necessary steps to turn genetic weaknesses into strengths and maintain correct levels.



Apolipoprotein B

ApoB should be checked if cardiovascular disease runs in your family



B12

If poor B12 status is suspected, methylmalonic acid (MMA) levels may be needed to accurately assess B12 status, absorption, and requirements



Fasting Glucose and HbA1C

Check both fasting glucose and HbA1C



Homocysteine

Homocysteine should be between 7-9



LDL

Test LDL-P, LDL-C, and small dense LDL



Mercury

If mercury levels are a concern talk to your doctor about the best testing options $% \left\{ 1,2,...,n\right\}$



Triglycerides

Triglycerides should be <150



MY CLINICAL RESEARCH SUMMARY: NUTRIENT METABOLISM & DIGESTION

Beta Carotene to Vitamin A Conversion Rate-BCMO1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
BCMO1 R267S rs12934922	Heterozygous AT
BCMO1 A379V rs7501331	Wild Type CC

Recap





BETA CAROTENE TO VITAMIN A CONVERSION RATE-BCMO1

Research: If you are heterozygous or homozygous for BCMO1 A379V or BCMO1 RS267S, you have a reduced conversion of beta-carotene to vitamin A. If you have a heterozygous or homozygous BCMO1 RS267S and BCMO1 RS267S, the reduction is even more dramatic. Many nutrition labels will have beta-carotene listed as vitamin A, however this is not true vitamin A.

The normal conversion for beta-carotene (carrots, sweet potatoes) to retinol is 1:6 and 1:12 for other carotenoids. Female volunteers carrying the T variant of rs7501331 (379V) had a 32% lower ability to convert beta-carotene, and those carrying at least one T in both SNPs (379V and R267S) show a 69% lower ability to convert beta-carotene into retinol.

In a cohort study of 48,400 US men and 75,170 US women, during a follow-up period of more than 26 years, a higher total vitamin A intake was associated with a reduction in cutaneous squamous cell carcinoma risk.

You want to make sure you consume animal based vitamin A (pastured egg yolks, wild salmon oil, cod liver oil, butter) along with zinc for digestive lining repair, oral health, eye health, iron mobilization, mitochondria health, skin health (sunburns deplete vitamin A in the skin, and acne responds to vitamin A), healthy lung function, and increased immunity.

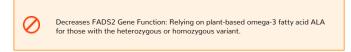
ALA to EPA and DHA Conversion-FADS2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
FADS2 rs1535	Heterozygous AG
FADS2 rs174575	Heterozygous CG

Recap





ALA TO EPA AND DHA CONVERSION-FADS2

Research: You may have a decreased conversion rate of the plant based omega-3 fatty acid ALA to DHA and should choose DHA sources for sufficient omega-3's.

FADS1 and FADS2 are enzymes that are involved in converting omega-3 and omega-6 fatty acids for brain development and inflammation control. Like the lactase gene, FADS1 is likely to be a critical gene of adaptation. In this case, it was in response to a plant-based diet versus a meat and fish based diet depending on migration routes and food availability.

It has been hypothesized that populations that began to rely more on plant-based diets adapted with the selected allele in FADS2 to synthesize more EPA and DHA from plants. The Inuit populations of Greenland, for example, who rely heavily on seafood with very little plant intake, have a deleted allele showing an opposite adaptation to a diet without plants.

A meta-analysis has found an association between variants in FADS2 in European heritage and a low conversion rate of ALA (plant-based omega-3) to DHA. There is also evidence for gene variants in those with African, Chinese, and Hispanic ancestry having a reduced conversion rate.

Children who had a higher dietary ratio of omega-6 to omega-3 were vulnerable for developing colitis if they also presented specific variants in FADS2.

A higher need of animal-based EPA and DHA may be needed for those with variants in FADS2.

B6-NBPF3

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE		
NBPF3 rs4654748	Heterozygous CT		

Recap





Decreases NBPF3 Gene Function: Sugar, stress, high intake of alcohol and refined flour based carbohydrates, antibiotics, oral contraceptives, ACE inhibitors, antacids, proton pump inhibitors, Phenytoin, bronchodilators, Digoxin, diuretics, hormone replacement therapy, Estradiol, MAO inhibitors, St. John's Wort and Parnate.

68 B6-NBPF3

Research: You may require a higher intake of B6. Heterozygotes have a 1.45 ng/mL lower Vitamin B6 blood concentration than the wild-type genotype.

Vitamin B6 plays a major role in neurotransmitter health. B6 deficiency can manifest as anorexia, irritability, anxiety, depression, muscle pain, bad PMS/low progesterone, nausea, seizures, migraines, dermatitis, age related macular degeneration (with low folate and B12) and lethargy.

Researchers have found an inverse association between ovarian cancer risk and vitamin B6 intake. Subjects with the highest vitamin B6 intake showed a 24 percent decrease in the likelihood of developing ovarian cancer compared to the individuals with the lowest intake.

Women of reproductive age, especially current and former users of oral contraceptives, teenagers, male smokers, non-Hispanic African-American men, and men and women over age 65 are most at risk of B6 deficiency. Data suggests that oral contraceptive users have extremely low plasma PLP levels. Three quarters of the women who reported using oral contraceptives, but not vitamin B6 supplements, were vitamin B6 deficient.

Saturated Fat-PPAR-alpha

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE		
PPAR-alpha rs1800206	Heterozygous CG		

Recap



Improves PPAR-alpha Gene Function: Higher polyunsaturated fat intake and lowersaturated fat intake, astaxanthin, pterostilbene, terpenoids, genistein, tomatoes, cinnamon, zinc, Lion's Mane mushroom, Gynostemma tea, and L-carnitine.



SATURATED FAT-PPAR-ALPHA

Research: The PPAR-alpha gene is mainly expressed in tissues with extensive fatty acid metabolism and its activation leads to changes in the transcription of multiple genes that regulate lipid and lipoprotein metabolism. PPAR-alpha also regulates the metabolism of amino acids in the liver, influences the apolipoprotein (Apo) genes, HDL genes, LDL genes and ketone bodies during fasting.

The PPAR-alpha polymorphism (rs1800206) has been associated with increased levels of triglycerides, total cholesterol levels, and ApoB in many populations such as Caucasian, Indian and African-Americans, however variants may be more relevant in men. The positive trade-off is that variants in the PPAR-alpha genotype may also have lower C-reactive protein levels (CRP), a marker of inflammation.

Subjects with variants in PPAR-alpha have been found to have a larger waist circumference and a higher proportion of small, dense LDL particle size (risk factor for heart health) with a higher saturated fat intake and low polyunsaturated fat intake.

In mice missing the PPAR-alpha gene, fasting decreased the plasma level of the amino acid alanine and the ketogenic amino acid tyrosine. Those with variants in this gene have been found to have a poor response to fasting due to low ketone bodies and the research has stated that this genotype may explain diverse reactions to the ketogenic diet.

An agonist is a chemical that binds to a receptor and activates the receptor to produce a biological response. PPAR-II agonists activate the gene, promoting the uptake, utilization, and breakdown of fatty acids. The activation of PPAR-II has also been demonstrated to inhibit tumor growth and angiogenesis.

Animal models have shown that PPAR-alpha activation reduces body mass, and treats insulin resistance and non-alcoholic fatty liver. Research has found that polyunsaturated fats (fish, nuts, seeds), astaxanthin (high in wild salmon), pterostilbene (blueberries, mulberries, cranberries, raw almonds), genistein (fermented soy), tomatoes, cinnamon, zinc, Lion's Mane mushroom, Gynostemma tea and L-carnitine all activate PPAR-alpha gene expression. Human studies have shown that increasing polyunsaturated fat and decreasing saturated fat in those with variants in PPAR-alpha improved triglycerides levels, LDL particle size, and body weight.

One exception for saturated fats may be coconut oil. Rat studies found that virgin coconut oil decreased tissue lipid levels,

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reduced the activity of the enzymes involved in lipogenesis and enhanced the rate of fatty acid catabolism, which was mediated at least in part via the PPAR-alpha dependent pathways.

Fat Metabolism-ACSL1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
ACSL1 rs9997745	Wild Type GG

Recap



Improves ACSL1 Gene Function: Total fat intake under 35%, lower saturated fat intake and increased PUFA intake.



FAT METABOLISM-ACSL1

Research: If you have the GG genotype, it may be beneficial for fat intake to be below 35% of your total calories or have a higher intake of polyunsaturated fat from fish, nuts and seeds if you struggle with weight and high glucose.

The GG genotype had higher fasting glucose and insulin concentrations compared with the minor A allele carriers from saturated fat intake, with the result that the GG genotype was more insulin resistant. Among individuals within the top 50th percentile of PUFA intake, the metabolic syndrome risk associated with GG genotype was eliminated.

Foods that are higher on the insulin index include dairy and red meat, and insulin inhibits fat breakdown. Fat should come primarily from nuts, seeds, olive oil, avocados, poultry and fish if there are issues with fasting glucose, insulin or weight.

Carbohydrates-TCF7L2

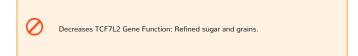
Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE	
TCF7L2 rs7903146	Heterozygous CT	

Recap



Improves TCF7L2 Gene Function: A Paleolithic and low-carb Mediterranean diet, glycine, omega-3 fatty acids, olive oil, turmeric, cinnamon, prebiotics, organic dark roast coffee, and cordyceps mushrooms.



CARBOHYDRATES-TCF7L2

Research: The TCF7L2 gene has become the strongest indicator of Type 2 diabetes and gestational diabetes risk for multiple ethnicities in studies. A meta-analysis also found an association with breast, prostate and colon cancer risk, all of which are connected to blood sugar levels and the risk is reduced by many of the same nutrients that improve this gene's function. Other genes and family history need to be assessed for cancer risk and prevention.

This gene is unique in its relation to Type 2 diabetes because people with variants in TCF7L2 may not exhibit risk signs like obesity. In fact, they may have a low body mass index (BMI) and low triglycerides. The increased risk is hypothesized to be due to the effect of TCF7L2 on the sensitivity of the pancreatic ß-cells to incretins, not overall insulin sensitivity.

Incretins are hormones that are released from the gastrointestinal tract after a meal and regulate the amount of insulin secreted. The two most important incretin hormones are GLP-1 and GIP. Researchers believe that increasing incretin sensitivity may decrease the risk of type 2 diabetes.

One study found that the consumption of meals based on the Paleolithic diet (no grains or dairy) focusing on fish, polyphenol-rich foods, fiber-rich vegetables, and spices high in phytochemicals resulted in significant increases in incretin and increased perceived satiety (feeling full). All three test meals were normalized to contain 50 grams of carbohydrates. Sufficient protein in particular shows promise in the management of Type 2 diabetes by stimulating incretin, insulin secretion, and slowing gastric emptying.

Two clinical studies have demonstrated that plasma GLP-1 levels rise following the ingestion of gelatin, a protein extraordinarily rich in glycine. Another study found that higher levels of indolepropionic acid produced by good bacteria due to a diet higher in prebiotic fiber-rich food decreased the risk of Type 2 diabetes.

Spices also appear very effective. Turmeric significantly increases the secretion of the incretin GLP-1. Cinnamon lowers blood glucose usually within physiological levels without hypoglycemia and increases satiety, showing it may act by potentiating the effects of incretin hormones.

There is a progressive deterioration of beta-cell function in patients with Type 2 diabetes. In vitro studies demonstrated that pancreatic beta-cell viability increased dramatically with cordyceps extract treatment, implying that cordyceps protect beta cells. This is crucial for the TCF7L2 gene due to the communication between pancreatic beta cells and incretins. The researchers concluded that "the potential ability of cordyceps to preserve beta-cell function may afford a promising therapy for diabetes."

Uric Acid-ABCG2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
ABCG2 (Q141K) rs2231142	Heterozygous GT

URIC ACID-ABCG2

Research: Uric acid is produced during the breakdown of purines formed in the body and in certain foods and drinks. Uric acid is then carried by your blood and passed through your kidneys where it is excreted in the urine. High serum levels of uric acid is defined as greater than 7.0 mg/dL in men and 6.0 mg/dL in women. Very low and very high levels appear to be clinically relevant.

Variants in the ABCG2 gene was correlated with a 53% reduced urate transport rate compared to the normal genotype. Data from a population-based study of 14,783 individuals support rs2231142 as the causal variant in the region and show highly significant associations with urate levels and gout. Data indicates that at least 10% of all gout cases in Caucasions are attributable to this causal variant.

Uric acid has a paradoxical function as both an antioxidant in blood plasma (preserving SOD3 function) and a pro-oxidant in cells. An acute rise in uric acid may show the body trying to protect against oxidative stress in the presence of vitamin C. Uric acid is capable of binding iron and inhibits iron-dependent ascorbate oxidation, preventing oxidative stress. It also stimulates expression of a glutamate transporter in astroglia, by which it protects neurons from glutamate-induced toxicity.

Very low levels of uric acid are being investigated for glutamate toxicity related disorders like Alzheimer's disease, Huntington's disease, Parkinson's disease, and Multiple Sclerosis. In a treatment of patients with a uric acid precursor, inosine, it prevented progression of multiple sclerosis in all 11 patients tested and even improved the symptoms of some patients.

However, chronically high uric acid levels are strongly associated with gout, hypertension, obesity, insulin resistance, elevated triglycerides, type 2 diabetes, kidney disease, uric acid kidney stones, oxidized LDL and cardiovascular disorders.

Gout is caused by an excess of uric acid in the blood and deposits of uric acid salts in the tissue around the joints, especially in the fingers and toes. One study found that men who drank two or more sugar-sweetened beverages a day have an 85% higher

chance of developing gout than those who drank such beverages infrequently.

Cherries are one of the best additions for lowering uric acid levels. In one study, cherry intake over a 2-day period was associated with a 35% lower risk of gout attacks compared with no intake. When cherry intake was combined with allopurinol use, the risk of gout attacks was 75% lower than during periods without either exposure.

Ten healthy subjects who swim regularly in ice-cold water during the winter (winter swimming), were evaluated before and after this short-term whole body exposure. A drastic decrease in plasma uric acid concentration was observed during and following the exposure to the cold stimulus.

Another study found that cold exposure allowed men to clear sugar from their blood 43 percent more efficiently than when they started.

A family history of exceptional longevity was associated with lower serum uric acid levels in Ashkenazi Jews.

More research is needed to eludicate the paradoxical role of uric acid.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
HLA DQ2.5	HLA DQ2.5- rs2187668	СС		
HLA-DQ8	HLA-DQ8- rs7454108	тт		
BCMO1 R267S BCMO1 encodes the	BCMO1 R267S- rs12934922		АТ	
conversion rate from beta- carotene to vitamin A.	BCMO1 A379V- rs7501331	СС		
FADS2 The FADS2 gene encodes the	FADS2-rs1535		AG	
conversion of plant based omega-3 fatty acid alpha linolenic acid (ALA) to EPA.	FADS2-rs174575		CG	
FUT2 The FUT2 gene controls prebiotic production, B12 absorption and how much bifidobacteria you carry in your digestive tract. The rs601338 SNP is found in European, African and Indian populations.	FUT2-rs601338	GG		
NBPF3 NBPF3 has been associated with vitamin B6 levels.	NBPF3-rs4654748		СТ	

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
SLC23A1 Solute carrier family 23 member 1 (SLC23A1) is one of the two transporters which aids in the absorption of vitamin C into the body. Polymorphisms in the gene are associated with reduced plasma vitamin C levels in the body.	SLC23A1- rs33972313	СС		
ACAT1-02 The ACAT gene converts protein and fat to ATP (energy) in the mitochondria, and plays an important role in cellular cholesterol homeostasis.	ACAT1-02- rs3741049	GG		
ADIPOQ ADIPOQ encodes for adiponectin, a protein secreted by fat cells that affect insulin and glucose metabolism. Low levels of adiponectin play a role in obesity, insulin resistance and Type 2 diabetes.	ADIPOQ- rs2241766	ТТ		
HFE-C282Y A homozygous HFE C282Y may lead to an iron overload due to increased iron absorption and disrupted metabolism. Compound heterozygotes C282Y/H63D and single C282Y heterozygotes carry a very low risk of hemochromatosis, making the homozygous C282Y the most clinically relevant.	HFE-C282Y- rs1800562	GG		
PPAR-alpha The PPAR-alpha gene plays a vital role in fatty acid metabolism and ketosis, and is considered one of the most critical targets for ameliorating abnormalities with triglycerides, HDL, LDL, VLDL, and ApoB.	PPAR-alpha- rs1800206		CG	

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
ACSL1 Long-chain acyl CoA synthetase 1 (ACSL1) plays an important role in fatty acid metabolism and triglyceride synthesis. Disturbance of these pathways may result in dyslipidemia and insulin resistance, hallmarks of the metabolic syndrome.	ACSL1-rs9997745	GG		
FTO Polymorphisms in the FTO genes have been shown to cause higher ghrelin levels	FTO-rs9939609		АТ	
(hunger hormone) in many populations, which can create a larger appetite and the potential for overeating.	FTO-rs17817449		GT	
APOA2 The APOA2 gene contains instructions for making a protein called apolipoprotein A-II, which is found in HDL cholesterol particles. The homozygous genotype has been linked to saturated fat intake and weight gain.	APOA2-rs5082	AA		
TCF7L2 TCF7L2 polymorphisms have been associated with low incretin hormones and impaired insulin secretion.	TCF7L2-rs7903146		СТ	
LCT LCT is the gene connected with the ability to breakdown lactose in dairy.	LCT-rs4988235		AG	
APB1 APB1 is encodes for the DAO enzyme to breakdown histamines primarily in the digestive tract. The homozygous genotype may increase the risk of migraines from histamines in women or a hypersensitivity to Aspirin in men.	APB1-rs10156191	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
ABCG2 (Q141K) The ABCG2 (Q141K) gene is located at the membrane of kidney proximal tubule cells, where it mediates renal urate secretion. Variants in this gene are linked to reduced uric acid excretion.	ABCG2 (Q141K)- rs2231142		GT	
ALDH2 Alcohol metabolism in the liver most commonly involves the enzymes alcohol dehydrogenase and aldehyde dehydrogenase, metabolizing alcohol to acetaldehyde, and then to acetate. ALDH2 encodes for aldehyde dehydrogenase, and variants can affect the levels of acetaldehyde and therefore the carcinogenic effect of alcohol.	ALDH2-rs671	GG		
ADRB2 The beta-2-adrenergic receptor (ADRB2) is the main target of the catecholamine epinephrine, and a primary mediator of the stress response. ADRB2 is widely expressed both in the gastrointestinal tract and in the CNS.	ADRB2-rs1042714	CC		
PPCDC PPCDC is necessary for the biosynthesis of coenzyme A and variants in this SNP are associated with serum zinc levels.	PPCDC-rs2120019	тт		
SELENBP1 The Protein Selenium Binding 1 gene codes for an integral membrane protein involved in antigen presentation and serum copper levels.	SELENBP1- rs2769264		GT	

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
TFR2 The TFR2 gene provides instructions for making a protein called transferrin receptor 2 to help iron enter liver cells. The receptor on the surface of liver cells binds to transferrin, which transports iron through the blood to tissues throughout the body. When transferrin binds to transferrin receptor 2, iron is allowed to enter the cell.	TFR2-rs7385804		AC	

MY CLINICAL RESEARCH SUMMARY: METHYLATION CYCLE

Folate-MTHFR 677

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
MTHFR 677 rs1801133	Heterozygous AG

Recap





FOLATE-MTHFR 677

The frequency of the 677 polymorphism of MTHFR in the Caucasian population is up to 50% heterozygous.

The heterozygous MTHFR 677 has a 30% reduced function, potentially creating a higher need for dietary methylfolate depending on climate, skin tone and sun exposure, and dietary B2, choline, betaine, B6 and B12 intake. Variants in MTHFR 677, especially the homozygous genotype, are higher in Mediterranean climates and malaria-endemic regions like Southeast Asia. Researchers believe these variants were selected to protect against UV-induced DNA damage and malaria.

While a heterozygous MTHFR 677 and MTHR 1298 have been associated with higher homocysteine levels, not all people will develop high homocysteine levels.

Homocysteine is a non-protein amino acid that is created and recycled in the methylation cycle. Sluggish enzymes in the cycle can cause elevated levels in the blood, which can cause inflammation in the blood vessels. High homocysteine has been implicated in amyloid buildup, DNA damage and cancer, mitochondrial dysfunction, cardiovascular disease, age related macular degeneration, apoptosis of neurons and depression. BH4 structurally resembles folate and has been described to be reduced in endothelial cells when increased levels of homocysteine are present. Stabilizing MTHFR with B2 and targeting

One study in 259 post-menopausal women found that for those with variants in CYP1B1 (rs1056836), KRAS (rs61764370) and MTHFR (rs1801133 and rs1801131), oral contraceptives and hormone replacement therapy was associated with shorter leukocyte telomere length. Shorter leukocyte telomeres are connected to premature aging, and may increase the risk of cancer, cardiovascular disease, obesity, diabetes, chronic pain, and sensitivity to perceived psychological stress.

A large meta-analysis showed the lack of statistically significant association between MTHFR mutations and coronary heart disease except in the Middle East and Japan, where it portrayed statistical significance.

It is important to consider riboflavin intake, PEMT, MTR/MTRR, and CBS activity to assess overall homocysteine metabolism. Too high or too low levels of B12, B6, folate or their co-factors may cause dysregulation of methyl donor activity.

Folate-MTHFD1 G1958A

GENE	GENOTYPE
MTHFD1 G1958A rs2236225	Heterozygous AG



Improves MTHFD1 Gene Function: 5-formyl-tetrahydrofolate (folinic acid) and chaling



Decreases MTHFD1 Gene Function: Folate and choline deficiency, proton pump inhibitors, oral contraceptives, NSAIDs, anticonvulsants, antivirals, antibiotics, and acid blockers/antacids.

FOLATE-MTHFD1 G1958A

Research: A meta-analysis strongly suggests that the MTHFD1 G1958A polymorphism might be associated with maternal risk for neural tube defects in Caucasian populations. However, the evidence of this association should be interpreted with caution due to the selective nature of publication of genetic association studies. Another study found that the polymorphism decreases enzyme stability and increases risk of congenital heart defects.

5-formyl-tetrahydrofolate is the second most common type of folate after methylfolate in the certain foods. This is why dietary folate is optimal because it addresses both upstream and downstream folate gene polymorphisms in the methylation cycle.

Checking MTHFR and PEMT genes along with MTHFD1 helps you determine your requirements for folinic acid, methylfolate and choline to help stabilize enzymatic function.

Choline-PEMT

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
PEMT rs7946	Homozygous TT
PEMT rs12325817	Heterozygous CG

Recap



Improves PEMT Gene Function: Choline, vitamin C and estrogen.



Decreases PEMT Gene Function: Nighttime pain relievers, antihistamines, antiseizure medications, sleep aids, antidepressants, incontinence drugs and narcotic pain relievers.

CHOLINE-PEMT

Improves PEMT Gene Function: Choline, vitamin C and estrogen.

Decreases Gene Function: Nighttime pain relievers, antihistamines, anti-seizure medications, sleep aids, antidepressants, incontinence drugs and narcotic pain relievers.

Research: Phosphatidylethanolamine-N methyltransferase (PEMT) catalyzes the synthesis of phosphatidylcholine.

Choline is responsible for shuttling fat out of the liver, aiding the gallbladder, healthy cell membranes to protect against inflammation, lowering anxiety, preventing damage from glutamate spikes, deep sleep, healthy DNA, healthy pregnancy and breast health. Non-Alcoholic Fatty Liver Syndrome occurs mainly from a choline deficiency. Choline deficiency also increases sensitivity to carcinogenic chemicals, mycotoxins and vegetable oils due to poor cell membrane health.

Research shows that the highest dietary intake of choline is found from people in the Northern countries, whereas Mediterranean countries had the lowest intake. Worldwide, total choline intake in adults ranges from 284 mg/day to 468 mg/day for men, from Taiwan and Sweden, respectively; and from 263 mg/day to 374 mg/day for women, from Mexico and Sweden. Major food sources of dietary choline vary by country. For example, eggs, meat, and dairy are the major sources of total dietary choline in New Zealand, while eggs, seafood, meats, and soy products are the predominant sources in Japan and China.

Having one or more T alleles at rs7946 is associated with having lower phosphatidylcholine production in the liver.

More than 40% of women have a genetic polymorphism in PEMT (rs12325817) that makes this gene unresponsive to estrogen, which creates the same high choline requirement as men. These women may be especially sensitive to dietary choline variations during pregnancy. One study found that the highest quintile of choline consumption was associated with a lower risk of breast cancer compared with the lowest quintile.

Eighty percent of the women who were homozygous for the rs12325817 SNP manifested signs of choline depletion (liver or muscle dysfunction), relative to 43% of subjects carrying one copy of the variant allele and 13% of subjects without the SNP. Almost 75% of the North Carolina population in the United States has one variant allele.

B6-CBS

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CBS A13637G rs2851391	Heterozygous CT

Recap



Improves CBS Gene Function: B6 and SAMe as co-factors, selenium and folate to



Decreases Gene Function: Antibiotics, arsenic, birth control, ACE inhibitors, antacids, proton pump inhibitors, Phenytoin, bronchodilators, Digoxin, diuretics hormone replacement therapy, Estradiol, MAO inhibitors, St. John's Wort, high cysteine and Parnate.

B6-CBS

Research: CBS is an important enzyme in the transsulfuration pathway that catalyzes the conversion of homocysteine (HCY) to cystathionine, a substrate for glutathione synthesis.

The CBS gene requires B6 and healthy SAMe production to regulate function. Deficiencies in CBS activity are the most frequent cause of familial high homocysteine and the underlying cause of the CBS genetic disorder homocystinuria, which is characterized by severe high homocysteine levels.

Research has hypothesized that rs2851391 variants might reduce the activity of CBS, and thus was positively associated with homocysteine levels and a marginal association with decreased plasma B12 levels.

One study demonstrated a significant association of both elevated homocysteine levels and low vitamin B6 levels with CBS polymorphisms in the presence of nonvalvular atrial fibrillation.

Hydrogen sulfide (H2S) may also need to be the focus with CBS and homocysteine levels. H2S is produced in the brain, pancreas, liver, reproductive tissues. Low levels of HS2 affect repair of the GI tract and disrupted levels of HS2 can lead to cognitive deficits or excitation in the brain.

Reduced CBS activity could cause low H2S concentrations, affecting mitochondrial health and the gut/brain axis. Abnormalities of hydrogen sulfide in the body have been identified in several disorders including ulcerative colitis, Alzheimer's disease, Down's syndrome, and possibly in diabetes.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
MTHFR 677 The MTHFR 677 gene encodes the MTHFR gene to convert folate into the active form, methylfolate. Variants in this gene slow down enzymatic function.	MTHFR 677- rs1801133		AG	
MTHFR 1298 MTHFR 1298 is involved in converting 5-methylfolate (5MTHF) to tetrahydrofolate (THF). Unlike MTHFR 677, the 1298 variant does not lead to elevated homocysteine levels unless paired with a heterozygous MTHFR 677.	MTHFR 1298- rs1801131	тт		
MTHFD1 G1958A (Methylenetetrahydrofolate dehydrogenase 1) encodes a protein that possesses three distinct enzymatic activities in the interconversion of 1-carbon derivatives of tetrahydrofolate.	MTHFD1 G1958A- rs2236225		AG	
DHFR A20965G Dihydrofolate reductase (DHFR) catalyzes the	DHFR A20965G- rs1643659		СТ	
reduction of dihydrofolate to tetrahydrofolate (THF) and affect synthetic folic acid metabolism.	DHFR C19483A- rs1677693		GT	
MTR A2756G MTR (methionine synthase) combines folate, methyl B12 and homocysteine into methionine.	MTR A2756G- rs1805087	AA		
MTRR A66G MTRR attaches a methyl group to B12 and variants here will slow the process. When both MTR and MTRR exist, dysfunction can occur.	MTRR A66G- rs1801394		AG	

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
TCN2 C766G Transcobalamin II (TCN2, or holotranscobalamin when bound) transports B12 to peripheral tissues. Variants in this gene may affect B12 transport.	TCN2 C766G- rs1801198	СС		
PEMT Variants in PEMT may increase the need for choline	PEMT-rs7946			π
and increase the sensitivity to anticholinergic drugs.	PEMT-rs12325817		CG	
CBS A13637G The Cystathione Beta- Synthase (CBS) enzyme pulls homocysteine to hydrogen sulfide (H2S) and glutathione, requiring B6 and SAMe as a modulator.	CBS A13637G- rs2851391		СТ	
CBS The Cystathione Beta- Synthase (CBS) enzyme pulls homocysteine to hydrogen	CBS-rs234709	СС		
sulfide (H2S) and glutathione, requiring B6 and SAMe as a modulator. CBS rs234709 and rs4920037 assists in arsenic detoxification.	CBS 191150T- rs4920037	GG		
ВНМТ	BHMT-rs3733890	GG		

MY CLINICAL RESEARCH SUMMARY: HORMONE SUPPORT

Sex Hormone Binding Globulin

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
SHBG rs1799941	Homozygous AA

Recap



Improves SHBG Gene Function for Women: SHBG levels that are too low will benefit from a high-fiber, low-fat diet, coffee, no sugar, and exercise 3-5 times a week. SHBG levels that are too high may be caused from SHBG variants combined with estrogen therapy (oral contraceptive or hormone replacement therapy). Improves SHBG Gene Function and Testosterone for Men: Magnesium, zinc, vitamin D, fish or fish oil, boron, adequate protein and a higher healthy fat intake (if testosterone is low and other fat genes are working optimally).



Decreases SHBG Gene Function for Men and Women: Anorexia, fatty liver, obesity, Type 2 diabetes, high fructose corn syrup, agave and crystalline fructose.

SEX HORMONE BINDING GLOBULIN

Research Women: Sex Hormone Binding Globulin (SHBG) is synthesized in the liver, and in the blood it transports and regulates the access of sex steroids to their target tissues. Serum levels of SHBG are influenced by hormonal as well as nutritional and metabolic status.

In a study of Italian women free of diabetes, serum SHBG levels showed a U-shaped trajectory with age, declining from age 20 to age 60, and increasing after the age 60 progressively after each decade. These changes mirror the age-related changes in BMI and fasting insulin, suggesting that BMI and insulin negatively influence SHBG concentration.

The SHBG levels in AA homozygotes for rs1799941 were 39% higher than in GG homozygotes in post-menopausal women. Subjects with the A allele (GA+AA) for rs1799941 had a trend for lower free estradiol index compared to the GG genotype. They also had a significantly lower bone mineral density (BMD) at the intertrochanter of the hip and trend for lower BMD at the total hip.

Changes in SHBG concentration will also affect the levels of bioavailable testosterone in women. Elevations in estradiol (as occurs during pregnancy), oral contraceptives, hormone therapy, anorexia and hyperthyroidism cause a marked increase in SHBG levels with a subsequent decrease in serum free testosterone levels. Levels of SHBG that are too high could affect mood, lean muscle mass, bone strength and sex drive.

Hypothyroidism, Type 2 diabetes, fatty liver and obesity are associated with SHBG levels that are too low, and therefore very low SHBG can be a biomarker for these disorders. A low-fat and high-fiber diet alone or combined with exercise reduces insulin, BMI levels and increases SHBG levels.

Research Men: Sex Hormone Binding Globulin (SHBG) is synthesized in the liver, and in the blood it transports and regulates the access of sex steroids to their target tissues. Serum levels of SHBG are influenced by hormonal as well as nutritional and metabolic status. In men, SHBG levels increase with age as testosterone levels decline.

Only a small fraction of the total testosterone - from 1% to 2% - is free in the blood and biologically active. About 40% to 70% of total testosterone travels around with SHBG and may not available to your cells. This means a large part of total testosterone may not be biologically active and available to your cells if SHBG is too high even though your testosterone is in a healthy range.

One study showed that serum SHBG concentration is increased in middle-aged men with primary or secondary osteoporosis and is correlated with bone remodeling markers, hip bone mineral density, and vertebral fracture risk. Serum SHBG level was significantly higher (+42.2%), whereas free androgen index was lower (-24.8%) in patients with primary or secondary osteoporosis. Testosterone and estradiol levels did not correlate with any bone resorption or bone formation markers for men. Another study found that osteoporotic Chinese men had lower free testosterone (FT) and higher levels of SHBG.

You can also go too far in the other direction. A study of men in the U.S indicated that men with lower concentrations of total testosterone and SHBG had a higher likelihood of having metabolic syndrome than those with higher concentrations.

The associations of rs12150660 and rs6258 were confirmed in the three replication cohorts showing that men with the GT and TT genotype for rs12150660 had higher levels of testosterone, free testosterone, and SHBG, while the TC genotype for rs6258 had lower testosterone, calculated free testosterone and SHBG compared to the wild-type CC genotype. Not enough subjects had the homozygous TT genotype to produce data.

The rs6258 SHBG gene was found to substantially affect SHBG binding affinity by lowering free testosterone levels. The lowest testosterone levels were found in those with the GG genotype of rs1210660 and the TC or TT genotype of rs6258. Therefore variants in rs12150660 may benefit free testosterone levels even though SHBG is higher, however this may depends on your rs6258 genotype.

Another study found that individuals with the AA genotype for rs1799941 were associated with decreased sperm motility compared to GG genotypes. Research has found that vitamin C supplementation might improve sperm count, sperm motility, and sperm morphology.

Vitamin D-CYP2R1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CYP2R1 rs10741657	Homozygous GG

Recap





VITAMIN D-CYP2R1

Research: Studies confirm that CYP2R1 is the principal 25-hydroxylase in humans and demonstrates that CYP2R1 alleles have dosage-dependent effects on vitamin D homeostasis.

A 2018 meta-analysis of sixteen articles with a total of 52,417 participants was reviewed for rs10741657. The GG genotype was associated with a clear descending trend of 25(OH)D levels when compared with the AA genotype in Caucasian and Asian populations.

Research has shown that oral administration of vitamin D led to negligible increases in serum 25-hydroxy-vitamin D for homozygotes, and significantly lower increases in serum 25-hydroxy-vitamin D in heterozygous subjects than in control subjects. The heterozygous effect may only be relevant in Caucasian populations.

Vitamin D can influence the expression of more than 1,000 genes and vitamin D deficiency has been linked to fatty liver, seizures, infertility, osteoporosis, cancer, autism (mother deficient), depression, heart attacks, Alzheimer's, dementia, high blood pressure, low testosterone in men, autoimmune disorders and more.

The literature is mixed on optimal vitamin D levels, which most likely vary based on your heritage, skin color and current health issues. The most well documented cause of Vitamin D deficiency is inadequate sunlight exposure such as high latitude countries. Paradoxically, despite its high sunlight hours, vitamin D deficiency is well recognized in Middle Eastern women, inner city young adults in America, athletes and dancers in Israel, elite gymnasts in Australia, young Hawaiian surfers, and adolescent girls in England.

For athletes, vitamin D deficiency has long been associated with muscle weakness and suboptimal muscle function. A positive relationship between serum vitamin D level and jump height, jump velocity and power was found in young women.

Clinical vitamin D deficiency is below 20 ng/ml. There is little evidence to prove there is a benefit for levels above 50 ng/ml. The

latest cancer research has found that women with 25(OH)D concentrations greater than 40 ng/ml had a 67% lower risk of cancer than women with concentrations less than 20 ng/ml. Pesticides have been linked to suppressing vitamin D levels and creating a vitamin D deficiency. Your PON1 gene function should also be assessed.

Research has found that sunlight is the optimal way to optimize vitamin D levels along with exercise, vitamin D rich foods and vitamin D cofactors, however supplementation may be necessary.

Estrogen Metabolism-CYP1A2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CYP1A2 rs762551	Homozygous AA

Recap



Improves CYP1A2 Gene Function: Unfiltered fermented drinks (Kombucha, beer, wine), hops, marinades, cruciferous vegetables, blueberries, blackberries, red grapes, kiwi, watermelon, parsley, and spinach.



Decreases CYP1A2 Gene Function: Heterocyclic amines, nitrosamines, aflatoxin B1, polycyclic aromatic hydrocarbons, dioxins, and I⊩naphthoflavone. Omeprazole and primaquine are inducers. Caffeine and Tylenol combined with these compounds can make the effect worse.

ESTROGEN METABOLISM-CYP1A2

CYP1A2 is a key enzyme in caffeine metabolism and the 2-hydroxylation of the main estrogens, estrone, and estradiol. 2-hydroxylation and 16a-hydroxylation are two mutually exclusive pathways in estrogen metabolism. 2-hydroxyestrone acts as a weak estrogen or anti-estrogen. 16I-OHE1 acts as a procarcinogen.

Coffee may protect against breast cancer by altering estrogen metabolism. Women with higher coffee intake and the CYP1A2 homozygous AA fast metabolizer genotype have a ratio of high 2-hydroxyestrone to low 16\(\text{\text{\text{-}OHE1}}\). Researchers found that higher coffee intake was more protective against ER-positive breast cancer.

In men, a 2019 study found that low to moderate coffee intake and the AA fast caffeine metabolizer genotype were less likely to experience prostate grade cancer progression than non-consumers. In a large, pooled cohort of men with prostate cancer, coffee intake of more than 2.5 cups per day was associated with longer survival with the AA fast metabolizer genotype.

Estrogen Metabolism-COMT

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
COMT rs4680	Homozygous AA

Recap



Improves COMT Gene Function: Vitamin C, magnesium, and copper (copper should not be too low or too high).



Decreases Gene Function: Chronic stress, sugar, proton pump inhibitors, aspartame, low magnesium levels, low vitamin C levels, low and high copper levels, constipation, xenoestrogens, high homocysteine levels, high SAH levels, estrogen-based medications, and mercury toxicity.

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ESTROGEN METABOLISM-COMT

COMT is a phase II enzyme involved in the inactivation of catechol estrogens that can otherwise lead to cancerous growth, while also increasing 2-methoxyestradiol, a metabolite that has been shown to inhibit the growth of breast cancer cells.

Variants in COMT V158M have been shown to decrease enzymatic activity and consequently increases the risk of carcinogenesis due to the accumulation of estrogen metabolites. COMT has been extensively investigated for correlation with different cancer risks including esophageal cancer, colorectal cancer, hepatocellular, carcinoma, lung cancer, breast cancer, ovarian cancer, endometrial cancer, testicular germ cell tumor, and bladder cancer with mixed results.

Due to the COMT V158M heterozygous and homozygous genotypes potentially having reduced estrogen clearance, slowing this pathway down further with chronic stress and a high catecholamine intake combined with poor gut health and low magnesium intake may affect the level of harmful estrogen metabolites.

However, this doesn't mean catecholamines should be avoided. It simply means that the dosage should be altered. For example, green tea has been found to be beneficial for breast cancer prevention in the COMT heterozygous and homozygous genotype because these individuals retained the polyphenols the longest. The wild type may need more to achieve the same benefit. Less is more for COMT variants.

MTNR1B-Melatonin

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
MTNR1B rs10830963	Heterozygous CG

Recap





MTNR1B-MELATONIN

Research: Melatonin is a hormone that helps to maintain our circadian rhythm such as the sleep-wake cycle, neuroendocrine rhythms or body temperature cycles through its action on melatonin receptors. The physiological effects of melatonin are various and include detoxification of free radicals and antioxidant actions, the activation of brown adipose tissue, bone formation and protection, reproduction, and cardiovascular, immune and body mass regulation. However, melatonin also affects glucose levels and insulin release.

In humans, melatonin release starts soon after sundown, reaches a peak between 2am and 4am and decreases gradually after that. However, in approximately one-third of individuals, there is a delay in melatonin release and stays elevated longer in the morning.

Dim light melatonin onset is defined as the start of the melatonin production in the evening during dim light conditions and has become a reliable phase marker of the circadian clock. One study found that MTNR1B rs10830963 G allele carriers had a significant association with delayed circadian phase of dim-light melatonin offset (1.37 hours) and a substantially longer duration of elevated melatonin levels in the morning (41 minutes).

MTNR1B rs10830963 has been associated with one of the strongest effects on insulin secretion and insulin sensitivity out of over 90 common variants identified for Type 2 diabetes and has been associated with gestational diabetes. Variants increase the amount of MTNR1B protein on the surface of insulin-producing cells, making the cells more sensitive to the effects of melatonin, which results in less insulin. Subjects carrying one or two G alleles showed a 2 to 4-fold increase in MTNR1B mRNA expression in human pancreatic islets, respectively, compared with the non-carriers.

The individuals with G allele of rs10830963 have been associated with increased plasma glucose level, decreased serum insulin level and an increased risk of Type 2 diabetes in Caucasians, Asians, African Americans and Hispanics. The researchers suggest that an increase of food intake to coincide with elevated melatonin levels in the evening and early morning lead to decreased glucose tolerance.

In a randomized, cross-over trial to compare glucose tolerance in the presence (late dinner 1 hour before bedtime) or absence (early dinner, 4 hours before bedtime) of elevated physiological melatonin concentrations, researchers compared the results

between homozygous carriers and non-carriers of the MTNR1B risk allele. The concurrence of meal timing with elevated endogenous melatonin concentrations resulted in impaired glucose tolerance. This effect was stronger in MTNR1B risk-carriers than in non-carriers. Furthermore, eating late significantly impaired glucose tolerance only in risk-carriers and not in the non-risk carriers.

Results have also found that in carriers of the MTNR1B risk variant, melatonin supplementation (5 mg) significantly impaired glucose tolerance, with no effect in non-carriers. These results have been recently replicated, and are consistent with our findings even after chronic melatonin administration.

Oral contraceptives have been found to increase nighttime melatonin levels due to inhibiting catalyzing enzymes in the liver, and therefore could theoretically create a higher impact on insulin release and glucose tolerance in G carriers.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
SHBG Sex Hormone Binding Globulin (SHBG) is synthesized in the liver, and in the blood it transports and regulates the access of sex steroids to their target tissues.	SHBG-rs1799941			AA
DI01 DI01 is connected to thyroid health and is responsible for the deiodination of T4 into T3.	DI01-rs2235544		AC	
DI02 DI02 is connected to thyroid health and is responsible for the deiodination of T4 into T3. D2 is the only activating deiodinase in the brain.	DI02-rs225014		СТ	
CYP2R1 Vitamin D is technically a hormone, and CYP2R1 is connected to circulating vitamin D levels.	CYP2R1- rs10741657			GG
CYP1A1 CYP1A1 is in the estrogen metabolism pathway along with CYP1B1, CYP1A2, CYP31A, SULT's and COMT.	CYP1A1-rs1048943	тт		
CYP2C19*17 Genetic variability impacts expression and activity of CYP2C19 and therefore can influence drug metabolism and catabolism of estrogens.	CYP2C19*17- rs12248560	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
CYP1A2 CYP1A2 is a key enzyme in caffeine metabolism and the 2-hydroxylation of the main estrogens, estrone, and estradiol.	CYP1A2-rs762551			AA
COMT COMT is involved in catecholamine, dopamine, adrenaline, and estrogen metabolism through the inactivation of the catechol estrogens.	COMT-rs4680			AA
FUT2 The FUT2 gene controls prebiotic production, B12 absorption, and how much bifidobacteria you carry in your digestive tract.	FUT2-rs601338	GG		
MTNR1B The MTNR1B gene encodes for the melatonin receptor 1B.	MTNR1B- rs10830963		CG	
GC	GC-rs2282679	тт		
CYP27B1	CYP27B1- rs4646536		AG	
VDR-FOK	VDR-FOK- rs2228570			GG
DHCR7	DHCR7-rs12785878	тт		

MY CLINICAL RESEARCH SUMMARY: MENTAL HEALTH & COGNITIVE PERFORMANCE

MAO-Serotonin

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
MAO-A rs6323	Heterozygous GT

Recap



Improves MAO-A Gene Function: Vitamin B6, folate, B12, B2, magnesium, vitamin C and probiotics.



Decreases Gene Function: Antibiotics, aspartame, oral contraceptives, proton pump inhibitors, high estrogen levels, constipation and deficiencies in the vitamins and minerals above.

MAO-SEROTONIN

Research: MAO-A (Monoamine oxidase A) is a critical enzyme involved in breaking down important neurotransmitters such as serotonin, estrogen, norepinephrine, and dopamine. Normal variants for men may not be as relevant as they are for women due to the role of estrogen.

The heterozygous genotype of MAO-A does not have a major impact on MAO-A function, however, MAO-A can still be disturbed by high estrogen levels, constipation, antibiotics, certain medications and vitamin deficiencies.

Serotonin Receptor-Stress

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
5-HT2A rs6311	Heterozygous CT
5-HT2A rs6313	Heterozygous AG

Recap



Improves Gene Function: Moderate intensity aerobic exercise, cognitive behavioral therapy, mindfulness training, meditation, yoga, tryptophan, green or black tea, prebiotics, probiotics, B2, B6, B12, and folate.



Decreases 5-HT2A Gene Function: Chronic stress, poor gut flora, high-dose lithium, cannabis abuse, and excessive smartphone use.

SEROTONIN RECEPTOR-STRESS

Research: The serotonin 2A receptor (5-HT2A) has been implicated in mental disorders with complex etiologies that are still not clearly understood, in processes such as learning and memory, and also in neurogenesis. Although the functional significance of 5-HT2A polymorphisms are not entirely understood, there is evidence that rs6311 modulates transcription factor binding and promoter methylation, affecting gene transcription (the first step of gene expression).

The T allele of the 5-HT2A gene rs6311 has been shown to increase the 5-HT2A expression in vitro and is associated with anxiety, IBS and depressive disorders. It has also been hypothesized that 5-HT2A variants may influence resting vagal activity among persons with chronically high levels of perceived stress.

One meta-analysis showed that the T allele of rs6311 or the linked A allele of rs6313 was significantly associated with obsessive compulsive disorder (OCD). This result was confirmed in the author's subsequent comprehensive meta-analysis in 2016 with a larger dataset. Multiple studies in this analysis indicated that the rs6311 T allele was more abundant in females with OCD compared to control females.

Another meta-analysis of 37 twin samples suggests that obsessions and compulsions arise from a combination of genetic factors and non-shared environment. OCD might be shaped by a large number of genes of modest impact, which combine to influence the risk for developing OCD. Polymorphisms in genes related to BDNF, GABA, glutamate, serotonin, acetylcholine, glycine, ubiquitin, bradykinin, myelinization, TNFA, gender and environmental trauma may all have a cumulative effect on whether or not someone develops OCD.

Psoriasis is a chronic inflammatory skin disease affecting about 2-4% of the population worldwide, and is thought to be a multifactorial disease with both genetic and immunogenic backgrounds. Psoriasis occurs in connection with stress and mood disorders and is apparently induced in patients who have been treated with antidepressants. The serotonergic system, which consists of serotonin-producing cells, serotonin receptors and serotonin transporters, may play a significant role in psoriasis.

Theanine, a component of green tea and black tea, has been shown to increase BDNF levels, modulate serotonin and dopamine levels, and improve learning and memory. It has shown promise as an adjunct therapy for schizophrenia and depression, and researchers believe there may also be an application for anxiety disorders, panic disorder, OCD, and bipolar disorder.

Vagus nerve stimulation may be a promising add-on treatment for anxiety, depression, PTSD, seizures, and inflammatory bowel disease. Natural ways to stimulate the vagus nerve and increase vagal tone include singing, deep breathing, meditation and yoga. Another way is to make a dietary shift towards good gut bacteria, shown to influence the activity of the vagus nerve.

In human volunteers as well as in a rat model, administration of a probiotic formulation consisting of Lactobacillus helveticus R0052 (traditionally used in the manufacture of Swiss-type cheeses and long-ripened Italian cheeses such as Emmental, Gruyere, Grana Padano and Parmigiano Reggiano) and Bifidobacterium longum R0175A (colonizes at birth, but levels vary genetically) significantly attenuated psychological distress and reduced anxiety-like behavior. Research has also found that prebiotics can improve non-REM sleep as well as REM sleep after a stressful event.

One pilot study found that a 12-week moderate intensity aerobic exercise program reduced OCD symptoms and the reductions lasted 6 months later.

Another study combined cognitive behavioral therapy and a 12-week moderate intensity aerobic exercise program with tremendous results, exceeding effects typically observed with individual and group-based cognitive behavioral therapy for OCD based on leading meta-analytic reviews.

Dopamine, Adrenaline and Estrogen-COMT

GENE	GENOTYPE
COMT V158M rs4680	Homozygous AA
COMT rs4633	Homozygous TT



Research: COMT (catecholamine methyltransferase) shares a pathway with MAO-A and is the gene for dopamine, estrogen, adrenaline and catecholamine metabolism. This pathway requires magnesium, vitamin C and copper as co-factors.

Studies have found that the A allele in COMT V158M (rs4680) results in a 40% decrease in COMT enzyme activity, leading to naturally higher dopamine and adrenaline levels. There are both benefits and detrimental aspects to this combination.

Research has shown that those with the AA COMT V158M genotype may have increased creativity, divergent thinking (rs4680 and rs5993883), problem solving, better memory (in men only), executive function and overall cognitive function. Females with the AA genotype have also been found to be higher risk-takers, but this was not true in men. It should be noted that females also naturally have higher dopamine due to the effect of estrogen on COMT, and therefore the AA genotype may not be as impactful for women as it is for men.

The downside of the A allele in COMT V158 is that the body overreacts to stress and trauma that can lead to anxiety, depression, impulsiveness, obsessive behavior, irritability, ADHD and abnormal behavior. The "inverted U" hypothesis suggests that when dopamine levels are either too high or too low, cognition is adversely affected and therefore stress could be negatively affect cognitive performance for the AA genotype.

The AA genotype can also create a sensitivity to catecholamines (coffee, black tea, green tea, red wine, chocolate), especially in a stressed state. However, green tea has been found to be beneficial for breast cancer prevention in the AG and AA genotype because these individuals retained the polyphenols the longest. Other genetic variants involved in dopamine transport and receptor function also influence this magnitude.

One study found the A allele carriers performed better than G carriers in executive function tasks, being statistically significant in the adult group and more emphasized in men due to COMT activity being higher in the prefrontal cortex for men versus women.

Further research showed that individuals carrying A allele of rs4680 or T allele of rs4633 scored significantly higher on insight problem-solving tasks, and for the COMT H62H rs4633 gene, the homozygous TT and heterozygous TC carriers had higher insight problem-solving scores than those with CC genotype.

Having a homozygous variant in COMT V158M may increase your need for magnesium, vitamin C, and healthy copper levels (not too high or low). Compound weight lifting (squats, bench press, deadlift), sprints, and chopping wood can assist a slow COMT enzyme by increasing testosterone levels, which speed up the pathway and lower the stress response. Supplementation of magnesium and vitamin C may be essential to modulate COMT due to low magnesium levels in the water and soil, or lack of freshly picked fruits and vegetables for vitamin C, and chronic stress levels.

Histamines and Hyperactivity-HNMT

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
HNMT rs1050891	Wild Type AA

Recap



Improves HNMT Gene Function: Vitamin C, choline, folate and magnesium, chamomile, basil, stinging nettle, echinacea, fennel, ginger and wild oregano.



Decreases HNMT Gene Function: Food dyes, poor gut flora, gluten sensitivity, too many fermented foods, sodium benzoate, and deficiencies in vitamin C, choline, folate and magnesium.

HISTAMINES AND HYPERACTIVITY-HNMT

Research: HNMT stands for histamine methyltransferase. HNMT is the primary enzyme responsible for histamine metabolism in the brain. Dysfunction of the histaminergic nervous system is associated with various neuropsychiatric disorders including narcolepsy, Alzhemer's disease, Tourette's syndrome, eating disorders, and depression.

This gene requires adequate methyl donors from methionine and choline. If you do not have enough methyl groups available, you may be more prone to high histamine levels. HNMT polymorphisms differ considerably between Chinese and American populations.

In a 2019 mice study, results demonstrated that HNMT played an essential role in regulating brain histamine concentration, controlling aggression and sleep—wake cycles. HNMT disruption did not affect histamine concentration of the skin and stomach.

In children with ADHD, the adverse effect of food dyes and sodium benzoate on ADHD symptoms was determined by histamine degradation in the rs1050891 AA HNMT polymorphism. Histamine is lowered by vitamin C, choline, folate, magnesium, chamomile, basil, stinging nettle, echinacea, fennel, ginger and wild oregano.

Anandamide-FAAH

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
FAAH rs324420	Wild Type CC

Recap



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Improves FAAH Gene Function: Exercise over 30 minutes, red clover tea (women), kaempferol, cacao, genistein (fermented soy), Echinacea, 7-hydroxyflavone (parsley, onions, berries, tea, and citrus fruits), \(\Pi\)-caryophyllene (cloves, rosemary, hops).



ANANDAMIDE-FAAH

Anandamide is a neurotransmitter and endogenous cannabinoid, and is known as the "bliss" molecule that targets the endocannabinoid system.

The endocannabinoid system is involved in many physiological processes including reward, addiction, fertility, pain and energy regulation. This system was named from the cannabis plant, such as marijuana and hemp. THC closely resembles anandamide.

The endocannabinoids play a significant role in pain modulation and inflammation, and have been demonstrated to relieve pain by activating the CB1 and CB2 receptors.

The wild-type genotype (CC) encodes for the fast activity of FAAH, and therefore naturally leads to lower anandamide levels. Those with the homozygous genotype (AA), have the slow-activity of FAAH and naturally higher levels of anandamide. This means that the CC individuals may have more anxiety and have to work harder to achieve higher levels of happiness, while the AA individuals have less anxiety and naturally higher levels of the "bliss" molecule that stimulate feelings of happiness.

Low levels of anandamide have been linked to slower extinction of fear memories and a heightened stress response to threatening situations than those with higher anandamide levels. Healthy volunteers who carried the rs324420 "A" allele (low FAAH activity, high anandamide levels) had much less amygdala activation when placed in a threatening situation. They also had a weaker correlation between amygdala activation and trait anxiety, which is a general tendency to perceive situations to be threatening and to respond to such situations with subjective feelings of apprehension and tension.

Pesticides such as chlorpyrifos and diazinon alter the endocannabinod system and researchers have hypothesized that eating organic foods lacking pesticide residues may promote endocannabinoid balance. Phthalates are plasticizers added to water bottles, tin cans, food packaging, and even the enteric coating of pharmaceutical pills. Phthalates may act as endocrine disruptors and carcinogens, and have been found to block CB1 receptors, found in the brain.

However, there are also ways for people to lower excessive levels of chronic stress and anxiety by increasing anandamide levels in the body. One of best ways to do this is with exercise. Endorphins (endogenous opioids) enhance the effects of cannabinoids and what has been known as the "runner's high" may in fact be the increase of anandamide. Research found that running and biking over 30 minutes, along with strenuous hiking at high altitude significantly increased anandamide.

Clinical anecdotes suggest that stress-reduction techniques, such as meditation, yoga, and deep breathing exercises impart

Brain Health-PEMT

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
PEMT rs7946	Homozygous TT
PEMT rs12325817	Heterozygous CG

Recap



Improves PEMT Gene Function: Choline, vitamin C, and estrogen.



Decreases Gene Function: Nighttime pain relievers, antihistamines, sleep aids, antidepressants, incontinence drugs and narcotic pain relievers.

BRAIN HEALTH-PEMT

Research: Choline is required for acetylcholine, a neurotransmitter of the vagus nerve that innervates multiple organs including the lungs, heart, liver, stomach, ovaries, and temporal lobe of the brain. A deficiency could affect all of these, especially memory. Acetylcholine also plays a role in promoting REM sleep.

Having one or more T alleles at rs7946 is associated with having lower phosphatidylcholine production in the liver.

Eighty percent of the women who were homozygous for the rs12325817 SNP manifested signs of choline depletion (liver or muscle dysfunction), relative to 43% of subjects carrying one copy of the variant allele and 13% of subjects without the SNP.

Vitamin C has been shown to induce the release of acetylcholine from synaptic vesicles of neurons and increase acetylcholine levels in the brain.

Possible drugs that can cause memory loss include antidepressants, antihistamines, anti-anxiety medications, anti-seizure drugs, muscle relaxants, tranquilizers, sleeping pills, and pain medications given after surgery. Why? The majority of these are in a class called anticholinergic drugs and block acetylcholine.

A French study looking at 4,128 women and 2,784 men that reported taking anticholinergic drugs showed a greater decline over four years in verbal fluency scores and in global cognitive functioning than women not using anticholinergic drugs. In men, an association was found with a decline in visual memory and to a lesser extent in executive function. Significant interactions were observed in women between anticholinergic use and age, APOE genotype, or hormone replacement therapy. A significantly 1.4–2 fold higher risk of cognitive decline was observed for continuous anticholinergic users.

These drugs could be especially theoretically problematic for those with poor PEMT function, low estrogen (in women) and a family history of dementia and Alzheimer's disease.

Research shows that only 15% of women get enough choline, and one study found that those with lowest choline have the highest anxiety.

Panic and PTSD-GAD1

GENE	GENOTYPE
GAD1 rs3749034	Heterozygous AG



Probiotics, B6, B2, taurine, magnesium, lithium, choline, vitamin C, zinc, vitamin D, progesterone (women), CBD, lemon balm, ashwagandha, high intensity exercise for 8-20 minutes, endurance exercise, yoga, meditation, and deep sleep.



Antibiotics, caffeine, high estrogen, excess wheat, excess sugar, broth cooked over 24 hours, low blood sugar, poor sleep, manganese deficiency, boron deficiency, chronic stress, proton pump inhibitors, diuretics, hormone replacement therapy, MAOI's, fibrates, MSG, low progesterone, sucralose and aspartame.

PANIC AND PTSD-GAD1

GAD1 stands for "Glutamate Decarboxylase 1" and is responsible for the conversion of glutamate to GABA. GABA and glutamate account for 80% of brain activity. Glutamate is excitatory while GABA is calming. In the right amounts, glutamate helps focus, cognitive function and productivity. Too much, however, can be excitatory and detrimental.

The GAD system influences mood stability and the pathophysiology of mood and anxiety disorders. To date, GAD1 genetic variants have been associated with mood disturbance, and panic disorder. GAD1 SNPs may impact both mood and anxiety-like traits, and may also be relevant following stress or trauma exposure in influencing risk for PTSD as well as depression.

The subjects carrying A allele of rs3749034 were associated with an increased risk of Posttraumatic stress disorder when compared to subjects with the "G" allele in the dominant model.

GABA levels in various brain regions are reduced in panic patients possibly due to impaired GAD function. Further studies in patients with major depression found reduced GABA levels to be accompanied by increased glutamate concentrations strengthening the link between anxiety and mood disorders and GAD.

Following a trauma, individuals at higher genetic risk with certain genotypes in GAD1 may experience physiological effects of anxiety, overconsolidation of the fear memory, and negative thoughts about the event, decreasing their ability to extinguish fear responses when reminded of the trauma and increasing the likelihood of mood-related disturbances. Therefore the correlation with a genetic predisposition to a higher trauma response may require variants in GAD1, an environmental trauma, and gender to due the influence of estrogen on GAD.

Estrogen and progesterone decrease GAD expression in the amygdala and the hippocampus (which both are involved in regulating fear), which provides a link between hormone levels and anxiety as well as mood changes during menstruation in women. Natural progesterone in women (B6 helps produce progesterone) has powerful effects on enhancing GABA activity in the brain. When progesterone is too low, it causes elevated glutamate levels.

Abnormalities in the GABA neurotransmitter system have been noted in subjects with mood and anxiety disorders, which is why anticonvulsants are also marketed for mood disorders. Lithium and the drug Lamictal has been shown to help regulate the neurotransmitter glutamate by keeping the amount of glutamate between brain cells at a stable, healthy level. The anticonvulsant drug Topamax is used for migraines by lowering glutamate and raising GABA levels.

Excess glutamate is supposed to convert to GABA with B6 and magnesium. GAD1 variants slow down the conversion of glutamate to GABA and increase the need for B6/magnesium to make it run normally. Studies have found that exercise helps the brain direct excess glutamate to be used as an energy source and prevent toxic build-up.

GABA requires adequate probiotics (bifidobacterium produces large amounts of GABA, so the FUT2 gene function should also be assessed) zinc, B2, B6, vitamin C, vitamin D and deep sleep to keep glutamate in check. Taurine (found in grass-fed animal protein, wild fish and eggs) appears to increase the levels of GAD1 to reduce glutamate and help bind to GABA receptors in brain cells.

One study found that neuronal excitability from glutamate appears to be attenuated when eating or supplementing with the mushroom Lion's Mane. Research on Lion's Mane also shows that the hot water extract stimulates Nerve Growth Factor (part of a family of similar proteins that serve to promote the health and normal function of the brain and nervous system) and accelerates the growth of the myelin sheath. This has exciting potential for those with neurodegenerative disorders from high glutamate levels.

The artificial sweetener aspartame is especially troubling for those with GABA and glutamate imbalances. The lowered levels of serotonin due to aspartame consumption might cause lowered activity of the GABA transporters.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
MAO-A MAO-A (Monoamine oxidase A) is a critical enzyme involved in breaking down important neurotransmitters such as serotonin, estrogen, norepinephrine, and dopamine.	MAO-A-rs6323		GT	
5-HT2A The 5-HT2A gene encodes for serotonin receptors found in the brain and central nervous system and is concentrated in the brain region essential for learning and cognition. Polymorphisms in rs6314 may result in reduced episodic memory in young and middleaged individuals.	5-HT2A-rs6314	GG		
5-HT2A The 5-HT2A gene encodes for serotonin receptors found in the central nervous system. Polymorphisms in rs6311 and	5-HT2A-rs6311		СТ	
rs6313 may contribute to a reduced capacity to regulate stress, low vagal tone, anxiety, depression, OCD, and IBS, especially in females.	5-HT2A-rs6313		AG	
COMT V158M COMT is connected to dopamine, adrenaline,	COMT V158M- rs4680			AA
estrogen and catecholamine metabolism.	COMT-rs4633			тт
ANKK1 ANKK1 modulates the density of dopamine receptors in the brain.	ANKK1-rs1800497	GG		
DAO C2029G DAO participates in the degradation of extracellular histamine. This gene is connected to migraines.	DAO C2029G- rs1049793	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
HNMT C314T Histamine N- methyltransferase (HNMT) is a histamine-metabolising enzyme expressed in the brain. This gene is connected to migraines.	HNMT C314T- rs11558538	CC		
HNMT Histamine N- methyltransferase (HNMT) is a histamine-metabolising enzyme expressed in the brain. This gene is connected to hyperactivity and food dyes.	HNMT-rs1050891	AA		
FAAH FAAH (fatty acid amide hydrolase) is a gene that encodes for anandamide breakdown, a neurotransmitter and endogenous cannabinoid.	FAAH-rs324420	CC		
PEMT Choline is required for acetylcholine, a	PEMT-rs7946			тт
neurotransmitter of the vagus nerve that enervates numerous organs.	PEMT-rs12325817		CG	
GAD1 GAD1 stands for "Glutamate Decarboxylase 1" and is responsible for the conversion of glutamate to GABA.	GAD1-rs3749034		AG	
BDNF BDNF is a synaptic modulator of glutamate while GABA synapses are also regulated by BDNF.	BDNF-rs6265	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
SLC17A7 SLC17A7 mediates the uptake of glutamate into synaptic vesicles at presynaptic nerve terminals of excitatory neural cells in the brain. Polymorphisms are associated with delayed recovery time from head injuries.	SLC17A7- rs74174284		CG	
APOE Apolipoprotein E (APOE) is a lipid binding protein that	APOE-rs429358	тт		
transports triglycerides and cholesterol in multiple tissues, including the brain.	APOE-rs7412			СС
	GAD1-rs3791851	тт		
GAD1 GAD1 stands for "Glutamate	GAD1-rs2241165		СТ	
Decarboxylase 1" and is responsible for the conversion of glutamate to GABA.	GAD1-rs3791850		AG	
	GAD1-rs769407	GG		

MY CLINICAL RESEARCH SUMMARY: DETOXIFICATION

Liver Enzyme-CYP1A2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CYP1A2 C164A rs762551	Homozygous AA

Recap



Improves CYP1A2 Gene Function: Unfiltered fermented drinks (Kombucha, beer, wine), hops, marinades, cruciferous vegetables, blueberries, blackberries, red grapes, kiwi, watermelon, parsley, and spinach.



Decreases CYP1A2 Gene Function: Heterocyclic amines, nitrosamines, aflatoxin B1, polycyclic aromatic hydrocarbons, dioxins, and \(\begin{array}{c} -naphthoflavone. Omeprazole and primaquine are inducers. Caffeine and Tylenol combined with these compounds can make the effect worse.

LIVER ENZYME-CYP1A2

Research: Approximately 200 polymorphisms exist in CYP1A2 gene region, with numerous studies focusing on rs762551. You have the homozygous (AA) rs762551 genotype, which is the rapid metabolizer.

The cytochromes P450 liver enzymes play an important role in the development of various cancers since they are involved in the metabolic transformation of numerous endogenous and exogenous compounds including carcinogens. CYP1A2 is a key factor in the metabolic activity of carcinogenic aromatic and heterocyclic amines, and researchers have found that the inhibition activity of this enzyme may represent a logical strategy for preventing the development of human cancers induced by the aromatic and heterocyclic amines. Further research has shown a cumulative value of phase I (CYP-450 enzymes) and phase II enzymes (GSTM1, GSTP1 and NAT2) in determining individual carcinogenic potential of compounds.

Heterocyclic amines (HCAs) are created by high heat reacting with the proteins. The way to reduce HCAs is to use marinades. Marinades reduce HCAs by up to 90 percent. For further protection, pair with cruciferous vegetables (especially fermented like sauerkraut) and an unfiltered beer or Kombucha due to the protection of the yeast. Red wine, blueberries, blackberries, red grapes, kiwi, watermelon, parsley, and spinach all inhibit the mutagenic activity of certain HCAs in vitro.

High antioxidant fruits, lemon juice, herbs, and spices help keep meat fresh and juicy while protecting against HCAs and reducing AGEs.

Grass-fed meat is higher in vitamin E, and in a study adding concentrations of vitamin E to the surface of ground beef reduced HCA production by 70%. Aim for medium to medium-rare for red meat, flip often and avoid burning. The darker the color the higher the HCA concentrations.

Nitrosamines are used in pesticides, created by frying meat, and from a conversion in the gut by nitrites from cured meats. Vitamin C prevents nitrites from becoming nitrosamines. Limit cured meat consumption using nitrites and take vitamin C when needed.

Aflatoxin B1 is the most common in food and amongst the most potent genotoxic and carcinogenic. It can occur in grain-fed milk, nuts/grains stored in hot conditions or bins, vegetable oils, cocoa or coffee beans stored in warm conditions, and dried fruit. We don't recommend Brazil nuts because they are prone to aflatoxin contamination. Choose nuts and seeds in sealed bags, preferably sprouted. You also want to minimize or avoid oats (unless tested free of ochratoxin). Low protein diets may increase the toxicity of aflatoxin and promote cancerous growth.

Hops in beer contain a flavonoid called xanthohumol, which strongly inhibits CYP1A2. Xanthohumol has anti-carcinogenic properties and has been found to scavenge reactive oxygen species, including hydroxyl- and peroxyl radicals, and to inhibit superoxide anion radical and harmful nitric oxide production.

Liver Enzyme-THC and CYP2C9

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CYP2C9*3 A1075C rs1057910	Heterozygous AC

Recap



Improves CYP2C9 Gene Function: Avoiding vegetable oils and high dosing of drugs that could slow or inhibit this enzyme, especially if using THC.



Decreases Gene Function: Linolenic acid from vegetable oils (soy, corn, canola, sunflower, safflower) generates superoxide (inflammatory) via induction of CYP2C9 and should be avoided. Monounsaturated fats like olive oil and avocados are professed.

LIVER ENZYME-THC AND CYP2C9

Research: The heterozygous AC and homozygous CC genotype of rs1057910 have been found to be a poor metabolizers of THC, which could cause a sensitivity to marijuana, especially in the edible form. The effect is much more pronounced in the CC homozygotes.

Numerous studies have found that THC is immunosuppressive and reduces T-cell activity. In vitro studies have found that for the influenza virus, THC caused higher hemagglutinin 1 expression levels, and diminished CD4 and CD8 lymphoctye and macrophage recruitment into the lungs. For respiratory viruses, slow metabolizers of THC could be susceptible to viral activity if using THC.

Aromatic Amines-NAT2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
NAT2 rs1495741	Homozygous AA

Recap



Improves NAT2 Gene Function: Cruciferous vegetables, unfiltered fermented drinks, meat and fish marinades, blueberries, blackberries, red grapes, kiwi, watermelon, rosemary, parsley, carotenoids, and vitamin C.



Decreases NAT2 Gene Function: Smoking, commercial hair dyes, industrial and manufacturing plants, charred meat, and diesel exhaust.

AROMATIC AMINES-NAT2

Research: N-acetyltransferase 2 (NAT2) could influence the detoxification of numerous drugs, and chemical carcinogens including aromatic amines. Aromatic amines are chemicals found in industrial and manufacturing plants, tobacco smoke, commercial hair dyes, and diesel exhaust.

Generally, the NAT2 phenotype can be classified into slow, intermediate, and rapid acetylator. The AA genotype is the slow acetylator, and numerous studies have associated the NAT2 slow acetylator phenotype with bladder cancer risk in smokers found in America, Europe, and Asia. However, in nonsmokers, rs1495741 AA did not increase susceptibility to bladder cancer when compared to GG and AG genotypes.

Exposure to aromatic amines has been found to increase the risk of breast cancer in those that work in rubber factories, use hair dyes that contain 4-aminobiphenyl (which also affects Tp53), and consistently consumed meat cooked at high temperatures. Research has shown the aromatic amine formed with meat cooked at high temperatures may cause both DNA

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damage and cause the proliferation of estrogen-sensitive cancer cells.

Heterocyclic aromatic amines, known mutagens formed in cooked meat and fish at high temperatures, are considered the causative agents for the association between meat intake and prostate cancer risk. Researchers found that a high heterocyclic aromatic amine intake was significantly associated with an increased risk of prostate cancer among Japanese men with the NAT2 slow acetylator phenotype, CYP1A1 rs1048943 TC and CC genotype, and CYP1A2 AC and AA genotype.

Marinades, cruciferous vegetables, unfiltered fermented drinks, blueberries, blackberries, red grapes, kiwi, watermelon, rosemary, and parsley all help reduce the carcinogenic risk posed by heterocyclic amines in meat cooked at high temperatures.

Vitamin K2-VOKRC1*2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
VKORC1*2 rs9923231	Heterozygous CT

Recap





VITAMIN K2-VOKRC1*2

Research: Vitamin K2 is produced by intestinal microbiota and is usually enough to cover the daily requirements. It is fat soluble and stored in the liver. If your gut flora is disturbed by FUT2 genes, elevated glutamate, gluten sensitivity, and you have used the drug Warfarin or antibiotics, your K2 requirements may be higher.

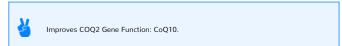
Vitamin K2 deficiency is linked to arterial calcification, osteoporosis and poor dental health. This is why long-term use of anticoagulants like Warfarin are linked to accelerated bone loss and bone mass. Recent research has shown that vitamin K2 plays a role in having an inhibitory effect on breast cancer cells.

Polymorphisms in VOKRC1 have been linked to higher rates of arterial calcification and may increase the need for vitamin K2. To paraphrase one study, "A lifelong decreased activity of the VKORC1 enzyme may increase the risk of vascular calcification and could be further worsened by reduced intake of vitamin K2."

Polymorphisms in VOKRC1*2 may increase the sensitivity to Warfarin dosing and vitamin K recycling. VKORC1*2 appears to be the most important in relation to the variability in response to oral anticoagulants and the risk of excessive bleeding. Vitamin K2 has also been found in studies to be inhibited by statin drugs.

Statins-COQ2

GENE	GENOTYPE
COQ2 rs4693596	Homozygous CC





Research: Statin drugs deplete CoQ10 and therefore may affect people more with variants in this pathway. One study found that people with the homozygous CC genotype were the most at risk for statin induced myopathy (muscle cramps, stiffness, and spasm). However, a study done in the Czech population in 2017 did not find an association with polymorphisms in COQ2 and low-dose statin drug therapy.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
CYP1A1*2C 4889 CYP1A1 is in the estrogen metabolism pathway along with CYP1B1, CYP1A2, CYP31A, SULT's and COMT. CYP1A1 is involved in the metabolism of benzopyrene.	CYP1A1*2C 4889- rs1048943	TT		
CYP1A2 C164A CYP1A2 metabolizes various environmental procarcinogens, such as heterocyclic amines, nitrosamines, aflatoxin B1 and ochratoxin A.	CYP1A2 C164A- rs762551			AA
CYP1B1*6 L432V The CYP1B1 gene metabolizes pro-carcinogens such as polycyclic aromatic hydrocarbons and 17 beta-estradiol.	CYP1B1*6 L432V- rs1056836		CG	
CYP2C9*3 A1075C Variants in CYP2C9 rs1057910 may alter the metabolism of THC, the psychoactive compound found in cannabis.	CYP2C9*3 A1075C- rs1057910		AC	
CYP2D6 T100C CYP2D6 metabolizes approximately 50% of drugs in clinical use.	CYP2D6 T100C- rs1065852	GG		
CYP2E1 Research has identified CYP2E1 as the primary P450 isozyme responsible for benzene metabolism at low concentrations, acrylamide to glycidamide, alcohol, Tylenol, and nitrosamines.	CYP2E1-rs2031920	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
CYP3A4*1B The CYP3A4 enzyme is involved in the metabolism of approximately 50% of drugs that are used today, cholesterol homeostasis, and the oxidative deactivation of testosterone.	CYP3A4*1B- rs2740574	ΤT		
CYP2C19*17 Genetic variability impacts expression and activity of CYP2C19 and therefore can influence drug metabolism and catabolism of estrogens.	CYP2C19*17- rs12248560	СС		
NAT2 The NAT2 gene encodes an enzyme that functions to activate and deactivate arylamine, hydrazine drugs, and carcinogens.	NAT2-rs1495741			AA
VKORC1*2 Variants in VOKRC1*2 may increase the need for vitamin K2 and a sensitivity to dosing of the drug Warfarin.	VKORC1*2- rs9923231		СТ	
COQ2 The COQ2 gene encodes an enzyme that functions in the final steps in the biosynthesis of CoQ10 and homozygous variants may increase the risk of statin induced myopathy.	COQ2-rs4693596			СС
CYP17A2	CYP17A2-rs743572	AA		

MY CLINICAL RESEARCH SUMMARY: IMMUNE SUPPORT,

ANTIOXIDANTS AND INFLAMMATION

Cell Protection-SOD2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
SOD2 rs4880	Heterozygous AG

Recap



Improves SOD2 Gene Function: Manganese, boron, vitamin A, C, E, omega-3 fatty acids, CoQ10, lutein, lycopene, milk thistle, cordyceps, holy basil, reishi and cryotherapy.



Decreases SOD2 Gene Function: Glyphosate, fluoridated water, chronic stress, poor sleep, shallow breathing, high iron levels and food dyes.

CELL PROTECTION-SOD2

Research: SOD2 is superoxide dismutase, which protects against the inflammatory superoxide inside the cell for the mitochondria (power house of the cell). SOD2 is manganese dependent, and adequate intake is important. Manganese is crucial for heart health, blood sugar, male fertility, bone health and protecting the brain against glutamate toxicity.

Exercise also helps improve SOD2 activity. Studies show exercise intensity can reduce cardiac arrhythmias and myocardial infarction due to improved SOD2 function.

Glutathione level and activity of antioxidant enzymes (catalase, superoxide dismutase, glutathione peroxidase and glutathione reductase) have been found to be increased in yoga practitioners. One year of Tai Chi training has been reported to promote superoxide dismutase activity and lessen lipid peroxidation.

One study found that young men exposed to cryotherapy for 3 minutes at -202°F (-130°C) everyday for 20 days doubled the activity of one the antioxidant enzyme glutathione reductase, and increased superoxide dismutase by 43%.

Chronic stress, poor sleep, shallow breathing and food dye consumption are examples of ways intracellular inflammation can occur. Food dyes have been found to inhibit mitochondrial respiration; the ability of the powerhouse of your cells to convert nutrients to energy and food dyes are often used ironically in sports drinks and multivitamins.

Fluoride decreases SOD2 activity in studies, and 75% of the water in the U.S. is fluoridated compared to 3% of western Europe. Reverse osmosis systems remove fluoride from water.

Variants in SOD2 increase the need for manganese to protect the mitochondria and lactobacillus in the gut. Colitis has been linked to impaired SOD2 genes.

Vitamin, A, C, E, omega-3 fatty acids, cordyceps and reishi help protect mitochondria against intracellular superoxide in red blood cells.

Glutathione-GSTM1

GENE	GENOTYPE
GSTM1 rs366631	Wild Type AA



GLUTATHIONE-GSTM1

Research: GSTM1 rs366631 is a pseudo-SNP that can be used as a GSTM1 deletion marker. The deletion is also known as the null genotype and confers the absence of the GSTM1 protein. The frequency of the null genotype varies from 20% to 80%, depending on the ethnic group studied.

For example, the null genotype is less frequent in western and southern African populations, less frequent in South American populations, intermediate in the Japanese, but is higher in Egyptian, European, American, and Asian populations.

High frequencies of the GSTM1 null genotype have been found in patients with lung cancer (East Asians), breast cancer (over 50 age group and in Asians), bladder cancer (with NAT2 slow acetylator), colorectal cancer, skin cancer, gastric cancer (among Asians with H. Pylori), chronic bronchitis, kidney disease progression, acute myeloid leukemia, acute lymphoblastic leukaemia, head and neck cancer (combined with CYP1A1 variant), endometriosis, type 2 diabetes retinopathy, and recurrent pregnancy loss. All have been regarded as environmentally induced and the risk may change with ethnicity.

Of the major glutathione enzymes, GSTM1 appears to be the most effective at neutralizing cytotoxic and genotoxic reactive compounds. However, the research shows that the null genotype of GSTM1 on its own may not be able to determine carcinogen exposure cancer risk. Instead, a combination of genotypes in the other glutathione and antioxidant genes like GSTP1 and NFE2L2, detoxification genes like CYP1A1 and NAT2, and/or compounding epigenetic habits that appear to modify the effect.

GSTM1 catalyzes the detoxification of alkyl and polycyclic aromatic hydrocarbons, intermediate forms of many carcinogens, specifically metabolically generated epoxide intermediates of benzo(a)pyrene. Benzo(a)pyrene is part of a class of chemicals called polycyclic aromatic hydrocarbons. Sources of benzo(a)pyrene include the burning of wood or trash, tobacco smoke, asphalt, coal, diesel exhaust, and grilled or charred meat. There is evidence that it causes skin, lung, and bladder cancer in humans and in animals. Research has also shown that early markers of cardiovascular disease are associated with occupational exposure to polycyclic aromatic hydrocarbons.

A study also found sensitivity to gas cooking and the GSTM1 null genotype, increasing the sensitivity of the lungs to nitrogen dioxide. Nitrogen dioxide is also found in diesel exhaust. Exposure of human blood plasma to nitrogen dioxide caused rapid losses of ascorbic acid, uric acid, protein thiol groups, lipid peroxidation, and depletions of alpha-tocopherol, bilirubin, and ubiquinol leading to high levels of oxidative stress.

Animal studies and in vitro studies have shown that vitamin C, vitamin E, vitamin A, resveratrol, curcumin, green tea, and white tea can inhibit the carcinogenic effect of benzo(a)pyrene and nitrogen dioxide. In the Norwegian Mother and Child Cohort Study 50,651 women, a higher prenatal exposure to dietary benzo(a)pyrene was found to reduce birth weight. However, increasing dietary vitamin C intake during pregnancy helped reduce any adverse effects of benzo(a)pyrene on birth weight.

Isothiocyanates from cruciferous vegetables are known for their anti-cancer activity. They are stored as glucosinolates in cruciferous vegetables and are hydrolyzed by myrosinase (an enzyme found in plants and intestinal microflora) to form isothiocyanates. Isothiocyanates from cruciferous vegetables are substrates and inducers of GSTM1.

GSTM1 variants may alter isothiocyanates clearance, with the null genotype retaining higher levels of isothiocyanates and therefore the benefits. In numerous studies, the GSTM1 null genotype was the most responsive to cruciferous vegetables for anti-cancer effects against lung cancer, colon cancer, breast cancer, and kidney disease.

The isothiocyanate levels in cruciferous vegetables will range based on growing conditions including sulfur and nitrogen levels, time after harvest and storage (cold transportation and storage of broccoli also cause a loss of glucosinolates up to 70-80%), plant genetics, and cooking preparation. Broccoli sprouts will yield the highest isothiocyanate levels.

Glutathione-GSTP1

GENE	GENOTYPE
GSTP1 I105V rs1695	Homozygous GG



Improves GSTP1 Gene Function: Glycine, cysteine, selenium, vitamin C, B1, B6, zinc, magnesium, optimal iron levels, magnesium, alpha lipoic acid, milk thistle, holy basil and vitamin E supplementation for the homozygous GG genotype only.



Decreases GSTP1 Gene Function: Mercury, arsenic, cadmium, pesticides, and air

GLUTATHIONE-GSTP1

Research: Glutathione is the master antioxidant system involved in oxidative stress, detoxification and immunity. It requires the amino acids glycine, cysteine and glutamate. Selenium activates the glutathione system and works in concert with vitamin E as a potent antioxidant against plasma and LDL lipid peroxidation.

The functional polymorphism of the GSTP1 Ile105Val gene, which reduces enzymatic activity, involves an A-G substitution. Carriers of these mutations are less able to detoxify carcinogens, and epidemiological studies have suggested that individuals differing in the expression of allelic variants of GSTP1 gene differ in susceptibility to various chemical carcinogens.

A meta-analysis of 10,067 cancer cases and 12,276 controls in 41 independent case—control studies from 19 articles found a significant increase in risk in breast cancer in Caucasions with variants in GSTP1 rs1695. A second meta-analysis found the same results with Asians that had the GG genotype. A 2020 study found that the rs1695 homozygous GG genotype was associated with an increased risk of breast cancer, but not the AG genotype. Other research has shown the risk to be higher in premenopausal women vs. post-menopausal women.

An analysis of that included 3,035 breast cancer cases and 3,037 population controls in a Chinese population found that cruciferous vegetable intake helped offset the risk of the GG genotype, with a lower risk associated with a higher cruciferous vegetable intake.

A meta-analysis of 11,762 cases and 15,150 controls from 51 studies showed a statistically significant association between GSTP1 rs1695 polymorphism with prostate cancer risk and urinary system cancer among Asians.

GSTP1 rs1695 variants were reported to be associated with the risk of esophageal cancer and malignant melanoma in the Caucasian population, but not childhood acute lymphoblastic leukemia or bladder cancer.

Glutathione-related polymorphisms, such as GSTM1 and GSTP1 have also been found to increase the elevation and toxicity of mercury. Selenium blocks mercury uptake, folate decreases mercury levels and magnesium and holy basil protect against mercury toxicity.

One benefit of the GSTP1 AG and GG genotype appears to be in athletic training. GSTP1 rs1695 AG and GG may be high responders to endurance training due to an impaired ability to remove excess reactive oxygen species. The hypothesis is that better activation of cell signaling pathways results in positive muscle adaptations. Women with at least one copy of the G allele showed a significantly greater increase in V?O2max in response to applied training.

In healthy control subjects, the effect of a-tocopherol supplementation on the production of inflammatory cytokines appears to be dependent on an individual's GSTP1 rs1695 genotype. These genotype-specific differences may help explain some of the discordant results in studies that used vitamin E. Persons having the alleles AA or AG in GSTP1 rs1695 had an increase in inflammatory interleukin-6 (IL-6) upon supplementing alpha-tocopherol (the most common form of Vitamin E in a North American diet) while those with GG saw a decrease.

Glutathione-GPX1

GENE	GENOTYPE
GPX1 rs1050450	Heterozygous AG



Improves GPX1 Gene Function: Selenium, optimal testosterone and estradiol levels, melatonin, vitamin C, vitamin E, black cumin seed oil, flavonoids, milk thistle, ginger, cumin, anise, fennel, caraway, cardamom and cryotherapy.



Decreases GPX1 Gene Function: Selenium deficiency, statin drugs, iron deficiency or elevated iron, and lead.

GLUTATHIONE-GPX1

Research: Superoxide dismutase (SOD) transforms the inflammatory superoxide to hydrogen peroxide (H2O2), and the next step is for glutathione peroxidase (GPX1) to transform it to water (H2O). When GPX1 function is modulated by polymorphisms and other factors affecting its function, a hydroxyl radical may be more likely to form which attacks DNA and causes strand breaks.

Research has shown that there is reason to believe that individual requirements for selenium will differ because of polymorphisms in seleno-protein genes. In a study looking at a New Zealand population, homozygous minor allele carriers of GPX1 rs1050450 had lower GPX1 activity than other genotypes with the same selenium status.

Elevated lead levels may have more toxic effects with GPX1 polymorphisms. A study looking at 362 patients and 494 controls found that lead exposure and GPX1 polymorphisms were significantly associated with glioblastoma and meningioma. Vitamin C decreases blood lead levels, and calcium reduces lead uptake.

GPX1 activity is considered to be the most important antioxidant enzyme defense mechanism in the skin. In a study from the Journal of Dermatological Science, the homozygous genotype for GPX1 rs1050450 was associated with a two-fold increased risk of melanoma.

Statins inhibit the biosynthesis of selenium-containing proteins, one of which is glutathione peroxidase serving to suppress peroxidative stress. An impairment of selenoprotein biosynthesis may be a factor in congestive heart failure, reminiscent of the dilated cardiomyopathies seen with selenium deficiency. A meta-analysis found that East Asian populations may be prone to cardiovascular issues with GPX1 polymorphisms.

Oxidative stress and inflammation play a pivotal role in the pathogenesis of Hashimoto's disease, an autoimmune disorder. A study looking at patients in Northwest Iran found that antioxidant capacity in Hashimoto's patients was lower than healthy controls. There was also a significant association with variants in GPX1 rs1050450, elevated anti-TPO levels, and Hashimoto's risk. The thyroid is the organ with the highest amount of selenium per gram of tissue. Research has suggested that selenium supplementation of patients with Hashimoto's disease is associated with a reduction in anti-TPO levels, improved thyroid ultrasound features, and improved quality of life.

In an experiment investigating the effect of heat and cold stress on glutathione metabolism in human erythrocytes, men were immersed at three different water temperatures for 10 min. At 39 degrees C (102 F), glutathione peroxidase decreased from 35.90 (1.83) to 34.33 (1.66) IU.g. The researchers concluded that "these changes indicate that heat stress causes oxidative stress in the human body; however, cold stress is thought to augment the activity of the antioxidative defense system. It is suggested that body exposure to hot environmental conditions should not be recommended for patients suffering from a damaged antioxidative defense system."

One study found that elite kayakers that engaged in whole body cryotherapy (-248 to 284°F or -120 to 140°C) for 3 minutes a day for 10 days increased the activity of superoxide dismutase by 36% and glutathione peroxidase by 68%.

Glutathione-CTH

GENE	GENOTYPE
CTH rs1021737	Homozygous TT





GLUTATHIONE-CTH

Research: Cysteine is an important amino acid for glutathione. Variants in CTH may increase the need for dietary cysteine. Homozygotes for this gene also had significantly higher mean plasma homocysteine concentration than subjects with other genotypes, and the effect sizes of CTH and MTHFR genotypes were similar.

You have the homozygous TT genotype for CTH. Oxidative stress has been suggested as a mechanism of autism. A three-SNP joint effect was observed for genotype combinations of SNPs in glutaredoxin, glutaredoxin 3 (GLRX3), and cystathionine gamma lyase (CTH). These results suggest that variation in genes involved in counterbalancing oxidative stress may contribute to autism, though replication is necessary.

The homozygous CTH genotype may require more cysteine-rich foods and a need to focus on other genes in lowering homocysteine and boosting glutathione.

Eye Health-CFH

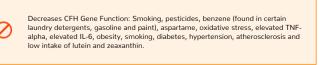
Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CFH rs1061170	Heterozygous CT

Recap

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EYE HEALTH-CFH

Research: Age related macular degeneration (AMD) is the leading cause of blindness in Western societies, but its etiology remains largely unknown.

Variants in CFH confers a 2-fold higher risk of late AMD per copy in individuals of European descent. Research indicates that CFH (rs1061170) polymorphism impacts significantly on retinal function in early AMD patients, and supports the hypothesis that a dysfunctional CFH might result in early retinal function loss due to a reduction in the immune antioxidant defense mechanism. A study from 2005 found that variants in CFH likely explains approximately 43% of AMD in older adults.

Malondialdehyde (MDA) is a common lipid peroxidation product that accumulates in many pathophysiological processes, including AMD. In vivo studies in mice found CFH as a major MDA-binding protein that blocks MDA-modified proteins by macrophages and MDA-induced pro-inflammatory effects. The CFH polymorphism markedly reduces the ability of CFH to bind MDA, indicating a causal link to a cause of age related macular degeneration.

A recent meta-analysis found that the rates of myopia (nearsightedness) will increase 140% by 2050 due to our increased time

in front of a screen. Myopia can increase the risk of numerous eye disorders. Blue light is a high-energy or short-wavelength visible light from your phone and computer that induces inflammation and retinal diseases such as age-related macular degeneration and retinitis pigmentosa. Research has found that bilberry and lingonberry exert protective effects against blue LED light-induced retinal photoreceptor cell damage due to their polyphenol content.

Lutein and zeaxanthin can inhibit oxidation of cell membranes and may be protective against UV-induced eye damage. Studies have demonstrated that people in the highest quintile of intake of dietary carotenoids, especially lutein and zeaxanthin concentrations have significantly lower risk of macular degeneration. Blue-eyed adults have far less lutein and zeaxanthin in their retinas.

One study compared diets of 356 patients with macular degeneration with 520 patients with other eye diseases. The data revealed that beta carotene was not especially effective, but that lutein and zeaxanthin were. Another study found that the risk of macular degeneration was reduced 65 percent with high amounts of lutein and zeaxanthin.

Research has found that MDA levels are significantly increased in groups of subjects with deficient levels of vitamin C and vitamin E. Deficiency in these two antioxidants leads to insufficient defense against free radicals and increased MDA levels. Those with polymorphisms in CTH should increase vitamin C and vitamin E intake. In another study, the risk for macular degeneration was found to be 77% lower when vitamin C supplements and a low-glycemic diet was used.

One study followed 3,600 people ages 55-80 years old for six years and found that those that took antioxidants plus zinc were less likely than those who took only antioxidants or only zinc to lose their vision.

Studies show that people who consume more fish, which is rich in DHA-fish fat, are less likely to develop macular degeneration. Eating fish one to three times a week has been associated with a 40 to 75 percent reduction in macular degeneration.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
SOD2 Superoxide dismutase (SOD2) is manganese dependent and protects against superoxide for the mitochondria of the cell. Variants here increase the need for intracellular antioxidant protection.	SOD2-rs4880		AG	
SOD3 Superoxide dismutase (SOD3) is zinc/copper dependent and protects against superoxide for the cell membrane. Variants here increase the need for intracellular and extracellular antioxidant protection.	SOD3-rs1799895	CC		
CAT C-262T CAT makes an enzyme called catalase, which helps reduce oxidative stress.	CAT C-262T- rs1001179	СС		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
GSTM1 GSTM1 catalyzes the detoxification of alkyl and polycyclic aromatic hydrocarbons (PAHs), intermediate forms of many carcinogens, specifically metabolically generated epoxide intermediates of benzo(a)pyrene.	GSTM1-rs366631	AA		
GSTP1 I105V Glutathione S-Transferase (GSTP1) is linked to the metabolism of mutagens, carcinogens, and other poisonous chemicals. It plays a crucial role in the detoxification process, thereby protecting cells from these compounds. GSTP1 rs1695 is connected to breast, prostate, urinary, esophagus, and skin health.	GSTP1 I105V- rs1695			GG
GSTP1 C341T Glutathione S-Transferase (GSTP1) is linked to the metabolism of mutagens, carcinogens, and other poisonous chemicals. It plays a crucial role in the detoxification process, thereby protecting cells from these compounds. GSTP1 rs1138272 is connected to the colon, prostate, lung, throat, and fertility.	GSTP1 C341T- rs1138272	CC		
GPX1 The GPX1 (Glutathione peroxidase 1) gene encodes a protein responsible for the modulation and detoxification of hydroperoxides and hydrogen peroxide to protect the mitochondria and cytoplasm of cells against oxidative damage.	GPX1-rs1050450		AG	

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
CTH The CTH (Cystathionine Gamma-Lyase) gene encodes an enzyme in the trans- sulfuration pathway that converts cystathionine derived from methionine into cysteine. Glutathione synthesis in the liver is dependent upon the availability of cysteine.	CTH-rs1021737			TT
NOS1 NOS1 (nNOS) codes for brain neural transmission, memory, learning, psychological stress, the peripheral nervous system and potentially the lymph nodes.	NOS1-rs3782218	СС		
NOS2 NOS2 (iNOS) encodes for wound, tissue damage, infection and hypoxia (low oxygen).	NOS2-rs2248814		AG	
CFH CFH (complement factor H) polymorphism is associated with increased risk of age related macular degeneration.	CFH-rs1061170		СТ	
ARMS2 ARMS2 polymorphism is associated with increased risk of age related macular degeneration (AMD).	ARMS2- rs10490924	GG		
LZTFL1 The LZTFL1 gene influences the transition of specialized lung cells to less specialized lung cells during infection and inflammation.	LZTFL1- rs17713054	GG		
IL-10	IL-10-rs1800872			тт
IL-10	IL-10-rs1800871			AA

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
IL-10	IL-10-rs1800896			ТТ
NQ01	NQ01-rs1800566	GG		

MY CLINICAL RESEARCH SUMMARY: DNA PROTECTION, DAMAGE & REPAIR

Longevity-SIRT1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
SIRT1 rs7895833	Wild Type AA

Recap



Improves SIRT1 Gene Function: Exercise, fasting, 7-8 hours of sleep per night, sauna, polyphenols, vitamin D, omega-3 fatty acids, resveratrol, magnesium, and melatonin.



LONGEVITY-SIRT1

Research: SIRT1 regulates numerous genes that accelerate the aging process, modulate DNA repair mechanisms and transcription factors like p53 (tumor suppressor gene), FOXOs (key regulators of lipid metabolism, stress resistance, and apoptosis) and inhibits NF-kb, a pathway connected to viral inflammation.

SIRT1 activity goes down as we age, and DNA damage accumulates, and its activity is especially harmed by a sedentary lifestyle, poor diet, and obesity. Activation of sirtuins induces the growth of blood vessels, insulin sensitivity and better glucose control, and other health benefits in a wide range of age-related cardiovascular and metabolic disease models. Experimental models have shown that increasing the activity of the sirtuins is associated with the delay of age-related diseases and potentially increasing longevity.

Researchers have observed a significant increase in SIRT1 levels in longevity populations and found a significant positive correlation between SIRT1 levels and age in a Turkish population. The oldest people carrying AG genotypes for rs7895833 had the highest SIRT1 level compared to the AA genotype, suggesting an association between rs7895833 SNP and lifespan longevity.

The average age of older people carrying AG genotype (76.0 \pm 1.5 years) was significantly higher than the average age of older people carrying AA genotype (71.3 \pm 1.4 years).

Your APOE genotype may also affect SIRT1 activity for longevity. Research from the Buck Institute group found that APOE-e4 reduced expression of SIRT1. The reduced expression of SIRT1 was thought to impair beta-amyloid clearance observed in Alzheimer's. If you have the APOE-e4 allele, the AA SIRT1 genotype may require more SIRT1 activation.

Polyphenols are activators of SIRT1 and contain anti-inflammatory and apoptosis properties. These include piceatannol (a metabolite of resveratrol), olive oil, fisetin (strawberries, apples, grapes), quercetin (wine, peppers, berries, apples) and resveratrol (wine, blackberries, blueberries, pistachios and dark chocolate).

Other activators of SIRT1 that also benefit the APOE-e4 carriers include magnesium, melatonin, vitamin D, and omega-3 fatty acids. One study found that centenarians (those living over 100) have higher total body magnesium and lower calcium levels than most elderly people.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
ATM D1853N ATM coordinates DNA repair by activating enzymes that fix double stranded DNA breaks.	ATM D1853N- rs1801516	GG		
ESR2 ESR2 acts as a tumor suppressor gene that codes for estrogen receptor beta (ER-beta), one of two main types of estrogen receptor activated by estrogen. ESR2 is strongly expressed in the breast.	ESR2-rs2987983	AA		
TP53 TP53 is a tumor suppressor gene responsible for DNA repair.	TP53-rs1042522		CG	
MDM2 Variants in the MDM2 gene encode a protein that reduces cellular levels of the p53 tumor suppressor protein.	MDM2-rs2279744	тт		
MLH1 MLH1 codes for a DNA repair enzyme linked to colon health.	MLH1-rs1800734		AG	
GATA3 GATA3 factors are involved in cellular maturation with proliferation arrest and cell survival.	GATA3-rs4143094	GG		
SIRT1 SIRT1 senses changes in intracellular NAD+ levels and plays a role in DNA damage and repair.	SIRT1-rs7895833	АА		
XRCC3 XRCC3 participates in DNA double-strand break/recombination repair.	XRCC3-rs861539	GG		

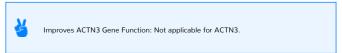
MY CLINICAL RESEARCH SUMMARY: CARDIOVASCULAR HEALTH AND ATHLETIC PERFORMANCE

Power and Recovery-ACTN3

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
ACTN3 rs1815739	Heterozygous CT

Recap





POWER AND RECOVERY-ACTN3

The T (R) allele is associated with enhanced strength and training adaptation, improved protection from eccentric training-induced muscle damage, lower risk of sports injury, and reduced frailty in the elderly. Testosterone levels were also higher in male and female athletes with at least one R allele compared to the XX genotypes. When stratified by race and gender for power athletes in a 2019 meta-analysis, Asian and male athletes benefited the most from the RX (TC) genotype.

The ACTN3 RR and ACTN3 RX groups have not been significantly different, indicating that the presence of one or two R alleles does not have a dose-dependent effect on 200-meter sprint speed in elite athletes. However, there was some evidence for a dose-effect of the ACTN3 R allele and 200-meter sprint speed in elite male African athletes. The ACTN3 RR individuals had (on average) a faster best personal sprint time than ACTN3 RX individuals.

Having the RX genotype may represent the best of both worlds for cold adaptation, longevity, strength training, and exercise recovery.

VO2 Max-PPARGC1A

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
PPARGC1A rs8192678	Heterozygous CT

Recap





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VO2 MAX-PPARGC1A

Research: Peroxisome proliferator-activated receptor gamma coactivator 1-alpha (PPARGC1A) is a master regulator of mitochondrial biogenesis, mitochondrial respiration, skeletal muscle fiber transformation (from fast to slow twitch), glucose and fatty acid metabolism, and the anti-oxidation machinery. PPARGC1A is expressed in cell types with high oxidative function

(heart, skeletal muscle slow twitch fibers, liver, and pancreas) and in brown adipose tissue.

Several studies have shown that SNPs in PPARGC1A are associated with a significant lower level in aerobic power (i.e., VO2 max) in insulin resistant and untrained individuals as well as in athletes. Healthy untrained adults display a large individual variation in VO2 max that ranges from -20% to more than 50%.

Research indicates that the exercise-induced variation in VO2 max is 47% explained by genetics. If you have heterozygous or homozygous variants in PPARGC1A, you may have a naturally lower VO2 max for aerobic exercise and increased CRP (C-reactive protein) levels.

To increase VO2 max, consider cold exposure. Since mitochondria are what give us the ability to use oxygen in order to produce cellular energy, the more we have the more the aerobic potential.

Cold exposure activates the PPARGC1A gene and PGC1® protein, which makes more mitochondria in the muscle. One study found that 15 minute exposure to cold water (50°F or 10°C) following high intensity running, increases PGC1® in muscle tissue. Another study found that men that were immersed in cold water at 50°F (10°C) for 15 minutes, 3 times a week for four weeks after running were able to increase mitochondrial biogenesis occurring in their muscle tissue.

Adaptogens are another way to increase your VO2 max. One study found that ashwagandha increased velocity, power, VO2 max, lower limb muscular strength and neuromuscular coordination. A second study used elite Indian cyclists for 8 weeks. One group received 500mg of the root extract 2x a day, while the other group received a placebo. There was significant improvement in the experimental group in all parameters, namely, VO2 max and time for exhaustion on treadmill.

A study using eleuthero root found that using 800mg for 8 weeks increased VO2 max of by 12%, endurance time improved 23%, the highest heart rate increased 4%, and metabolism was altered which spared glycogen storage. The study concluded that "this was the first well-conducted study that shows that 8-week ES supplementation enhances endurance capacity, elevates cardiovascular functions and alters the metabolism for sparing glycogen in recreationally trained males."

Muscle Recovery-IL6

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
IL6 rs1800795	Heterozygous CG

Recap



Improves IL6 Gene Function: Whey protein, cold water immersion, American ginseng, curcumin, allicin, optimal testosterone levels, vitamin C, and collagen protein.



Decreases Gene Function: Low testosterone (men), depression, obesity, bacterial

MUSCLE RECOVERY-IL6

Research: Exercise increases IL6 cytokines even when muscle damage hasn't occurred. It is produced in large amounts during heavy weight lifting and endurance races. The CG genotype is more common in sprint and power athletes compared to endurance and non-athletes.

C-allele carriers of the IL6 SNP have been found to have higher creatine kinase values (a marker of muscle damage) following exercise compared with GG homozygotes.

The highest post-exercise creatine kinase levels are found after prolonged exercise such as ultra distance marathon running, weight lifting and downhill running.

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To accelerate recovery, whey protein, cold water immersion, American ginseng, curcumin, optimal testosterone levels, vitamin C and collagen protein have all been found to attenuate creatine kinase levels.

Research has also found that purple sweet potatoes, cranberries, blueberries and beet root juice have verified health,

performance-enhancing, and exercise recovery benefits.

Perhaps the most promising results have come from two separate studies showing decreased muscle soreness and increased recovery from cherry juice and dehydrated cherry supplements. One of these studies had subjects perform ten sets of ten repetitions at 70% of a 1-RM back squat. The researchers found that Montmorency powdered tart cherry supplementation used daily and 48 hours post-workout significantly lowered muscle soreness strength decrement during recovery, and markers of muscle catabolism throughout the 48 hour post-lifting recovery period compared to placebo.

Muscle Injury-COL1A1

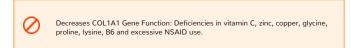
Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
COL1A1 rs1800012	Wild Type CC

Recap



Improves COL1A1 Gene Function: Vitamin C, zinc, copper, glycine, proline, lysine and B6 (all precursors to collagen production) and cryotherapy.



MUSCLE INJURY-COL1A1

Research: According to one study, the gene encoding for the alpha1 chain of type I collagen (COL1A1) has been shown to be associated with cruciate ligament ruptures and shoulder dislocations.

You have the CC genotype for COL1A1, which lowers the production of Type 1 collagen. Approximately 90% of collagen in the body is Type I. Type I collagen is found in the skin, tendons, corneas, lungs and in 95% of bone.

ACL ruptures are considered the most severe injury sustained in sports. The A variant produces more COL1A1. Two AA's reduced risk of ACL rupture by ten times, while only 5% of the population have two AA's.

Cryotherapy has been shown to inhibit harmful collagenase (activity on collagen enzyme that breaks down collagen) and also decreased the production of inflammatory E2 series prostaglandins. For athletes, cryotherapy post-training could be a useful tool to help prevent injuries.

Caffeine-CYP1A2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
CYP1A2 C164A rs762551	Homozygous AA

Recap



Increases CYP1A2 Gene Function: A higher cruciferous vegetable intake may help increase caffeine metabolism for those with the CC slow metabolizer genotype, along with exercise.



CAFFEINE-CYP1A2

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metabolized from your body and the effects lasting a shorter period of time. It is important to review your COMT gene function to better understand a sensitivity to coffee intake.

For the AA genotype, caffeine decreased 40-km time in cyclists by an average of 3.8 minutes in the AA homozygotes as compared to 1.3 minutes in the C allele carriers.

Triglycerides-FADS1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
FADS1 rs174546	Heterozygous CT

Recap





TRIGLYCERIDES-FADS1

Research: Variants in the FADS1 SNP (rs174546) are associated with elevated triglyceride levels, which appears to be due to a higher need for EPA and DHA from animal foods. Studies have found that plasma triglyceride levels were lower in wild-type CC genotype when compared to carriers of the minor T allele.

Population average triglyceride levels have increased since 1976 in parallel with the constant growing epidemic of obesity, insulin resistance and Type-2 diabetes. A meta-analysis of 17 population-based prospective trials including 46,413 men and 10,864 women identified plasma triglycerides levels as an independent risk factor for cardiovascular disease.

Triglycerides are essentially fat in the blood that are driven by excess sugar and carbohydrate consumption. They are the driving force behind lipoprotein particles that are potent causes of heart disease, such as small LDL and very low-density lipoprotein (VLDL).

Numerous studies have found that omega-3 fatty acids administered as fish oil supplements lowers plasma triglyceride levels by 25% to 34%. While fish oil is known to lower triglycerides, there doesn't appear to be a difference in the FADS1 genotype response to supplementation.

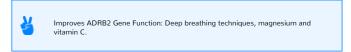
A meta-analysis of 13 randomized controlled trials found that 500mg of vitamin C resulted in a significant decrease in serum LDL cholesterol and triglyceride concentrations.

Stress-ADRB2

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
ADRB2 rs1042713	Homozygous AA

17 Recap





Research: ADRB2 activation regulates various biological functions, including heart rate, blood pressure or respiration, and it may modulate the vasodilatation in normal coronary arteries.

In a meta-analysis of seven case-control studies with a total of 6,843 subjects, a higher frequency of polymorphisms in rs1042713 was found with heart attacks or coronary artery disease compared to healthy controls. A similar result was also obtained with polymorphisms in rs1042714. Ethnicity-stratified subgroup analysis suggested that the rs1042714 variants correlated with an increased risk of the two diseases in both Asians and Caucasians, while rs1042713 only contributed to the risk of two diseases in Asians.

If cardiovascular disease runs in your family, you may be more prone to anxiety, high blood pressure and arterial damage from stress. You may also be more sensitive to beta blockers, leading to high triglycerides.

Since ADRB2 is connected to the catecholamine epinephrine, it is also important to look at your COMT gene function. Magnesium, vitamin C and adaptogens should be considered while making lifestyle changes and strategies if chronic stress is present.

Recently, the ADRB2 rs104271 was linked to fibromyalgia and temporomandibular joint disorder, two health issues that are frequently encountered in GI disorder patients. The AA genotype in the was more prevalent in the articular TMD group than in the muscular TMD group.

Relaxation techniques that involve slow, deep breathing have been found to be an effective therapeutic intervention that counteracts the adverse clinical effects of stress in disorders including hypertension, anxiety, insomnia and aging.

Potassium and Magnesium-ADD1

Below is a summary of your most significant variant genotypes:

GENE	GENOTYPE
ADD1 rs4961	Heterozygous GT

Recap



Improves ADD1 Gene Function: Lower sodium intake, magnesium, potassium, calcium, garlic, vitamin D and omega-3's.



Decreases Gene Function: High sodium intake, excess weight, high sugar intake, sedentary lifestyle, smoking and stress.

POTASSIUM AND MAGNESIUM-ADD1

Research: A meta-analysis of 33 studies with 40,432 participants found that variants in rs4961 was significantly associated with hypertension in Asians. Other research found that carriers of the risk (T) allele responded better to diuretics and sodium-restricted diets, in that they tended to lower their blood pressure by ~10 mmHg points compared to rs4961(GG) homozygotes similarly treated.

Excess weight, high sugar intake, sedentary lifestyle, smoking, stress and high sodium intake all raise blood pressure. People living at higher latitudes throughout the world are at higher risk of hypertension, and patients with cardiovascular disease are often found to be deficient in vitamin D. Magnesium, potassium, calcium, vitamin D, garlic and omega-3's all lower blood pressure.

One study found that increasing potassium-rich foods to 4.7 grams was equivalent to cutting out 4 grams of sodium in terms of reducing blood pressure.

In another study, aged garlic extract given at a dose of 600-1500mg was just as effective as the drug atenolol in reducing blood pressure over a 24-week period.

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
ACTN3 ACTN3 encodes for the alpha- actin-3 protein found exclusively within type-II fast- twitch muscle fibers.	ACTN3-rs1815739		СТ	
PPARGC1A It has been demonstrated that variants in the PPARGC1A gene affect the exercise-induced change in maximal oxygen uptake (VO2).	PPARGC1A- rs8192678		СТ	
TNFA Tumor necrosis factor (TNF-a) is a pro-inflammatory cytokine. Variants may increase the risk of asthma in Asian populations.	TNFA-rs1800629	GG		
IL6 IL6 is an interleukin that acts as both a pro-inflammatory cytokine and an anti-inflammatory myokine.	IL6-rs1800795		CG	
SOD2 Superoxide dismutase (SOD2) is manganese dependent and protects against superoxide for the mitochondria of the cell. The homozygous genotype increases the need for antioxidant support in high-intensity athletes.	SOD2-rs4880		AG	
COL1A1 COL1A1 produces alpha 1 chain of type I collagen, a major protein in tendons and ligaments.	COL1A1- rs1800012	СС		
PON1 PON1 (Paraoxonase) plays a large role in removing pesticides. It is also involved with supporting HDL function and LDL oxidation.	PON1-rs662	тт		

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
LPA Lp(a)is a sticky form of LDL that appears to affect plaque growth, LDL particle size and increase the risk of plaque rupture and blood clotting.	LPA-rs3798220	тт		
CYP1A2 C164A Variants in CYP1A2 determine caffeine metabolism and effects on bone density and cardiovascular health.	CYP1A2 C164A- rs762551			AA
9p21 9p21 is considered an important genetic marker for cardiovascular health.	9p21-rs4977574			GG
FADS1 FADS1 is involved in fatty acid metabolism, and variants in this gene are associated with elevated triglyceride levels.	FADS1-rs174546		СТ	
ESR2 ESR2 codes for estrogen receptor beta (ER-I), one of two main types of estrogen receptor activated by estrogen and is linked to fibrinogen levels in post-menopausal women.	ESR2-rs4986938	CC		
F5 Variants in F5 increase the risk of deep vein thrombosis, especially if using oral contraceptives.	F5-rs6025	СС		
ADRB2 Beta-2 adrenergic receptor (ADRB2) is abundantly expressed in cardiac cells, and bronchial smooth muscle cells and is connected to stress levels and heart health.	ADRB2-rs1042713			AA

Gene & Gene Function	Gene Rsid	Wild Type	Heterozygous	Homozygous
ACE1 G2350A ACE1 is part of the reninangiotensin system responsible for the conversion of angiotensin I to angiotensin II, constricting blood vessels and elevating blood pressure.	ACE1 G2350A- rs4343		AG	
ADD1 Variants in ADD1 are associated with hypertension in Asians.	ADD1-rs4961		GT	
AGTR1 Angiotensin-II receptor type 1 (AGTR1) is a major component of the renin-angiotensin system for regulating blood pressure and is highly expressed in adipose tissue, liver, leukocytes and the intestine. The homozygous genotype may increase the risk of high blood pressure from excess dietary fat and carbohydrate intake.	AGTR1-rs5186		AC	
ACE2 A8790G ACE2 is part of the reninangiotensin system, responsible for degrading angiotensin II and providing balance to ACE1 by dilating blood vessels and lowering blood pressure.	ACE2 A8790G- rs2106809	AA		
TMPRSS2 Transmembrane Serine Protease 2 is highly expressed in the prostate and lungs, and the expression is associated with viral susceptibility and prostate cancer.	TMPRSS2- rs2070788		AG	

Sources

Please click the link below if you'd like to browse peer-reviewed studies referenced by this analysis:

https://www.mydna.life/wp-content/uploads/myDNA-Comprehensive-Health-Report-references.pdf