

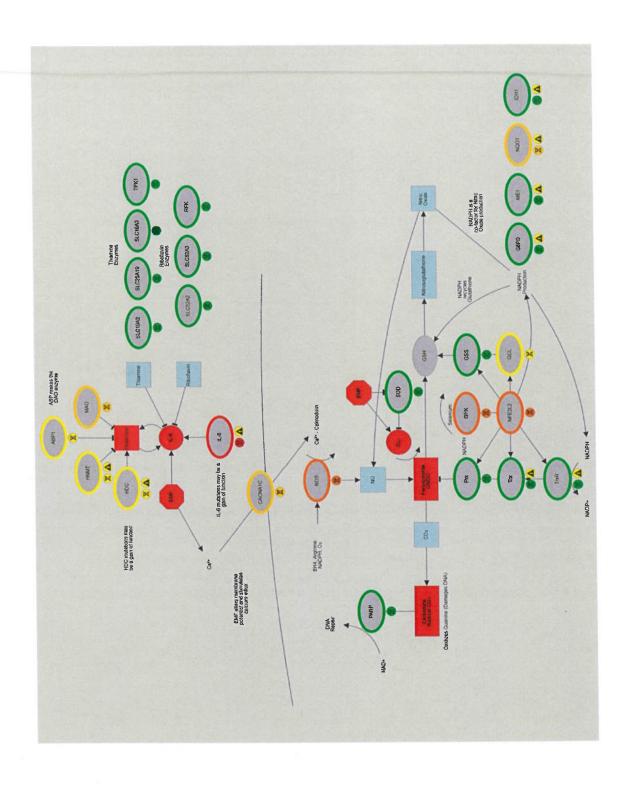
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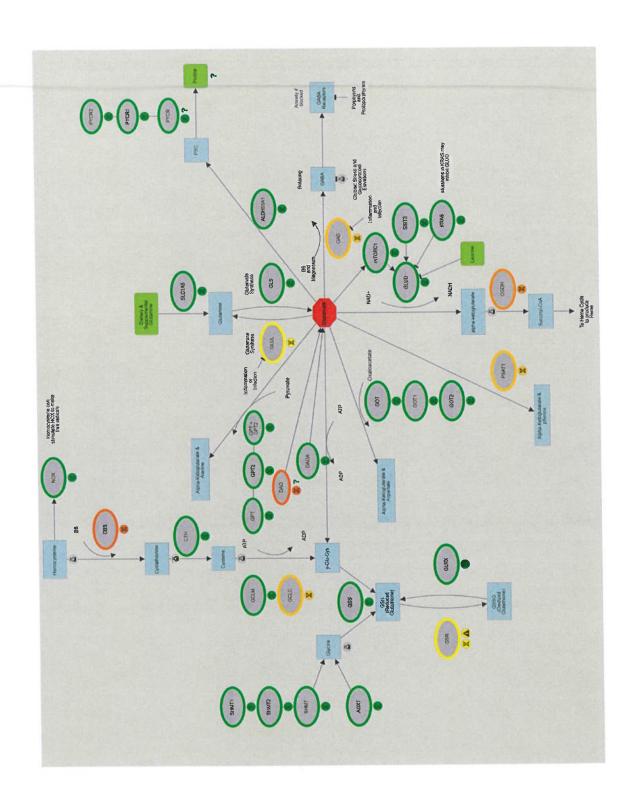


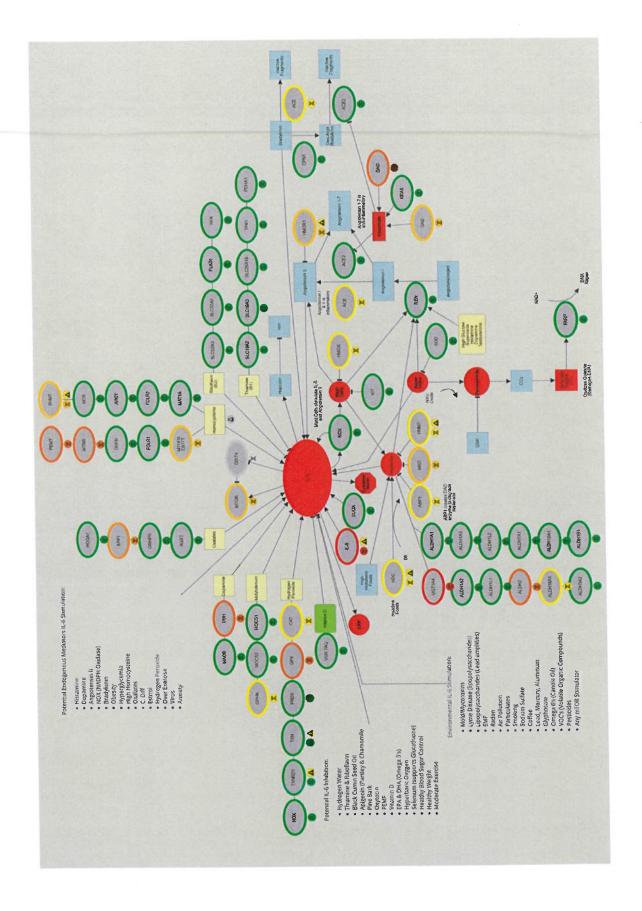
〒212-0013 神奈川県川崎市幸区堀川町 580 ソリッドスクエア東館 1F 医療法人社団あえん会 スクエアクリニック

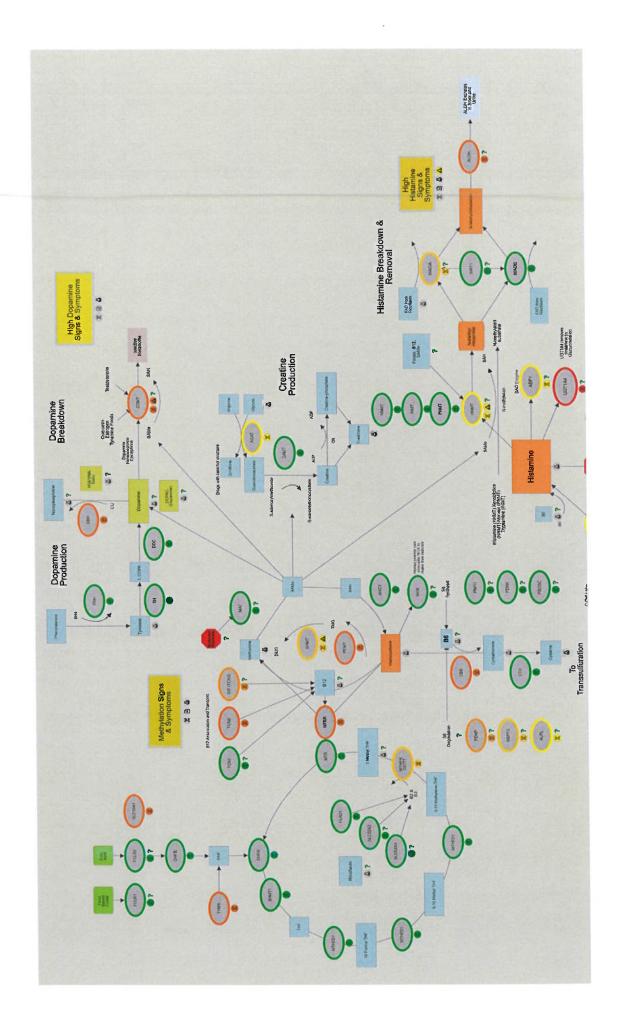
NOX, Mast Cells, and EMF 13 Sensitivity 13 FOXO Phase II Methylation 13 (Detox) M Heme Pathway & Sulfites 13 SHBG Glutathione Conjugation 13 Phase II 13 Sirtuins 13 PON1 **(1)** Circulation, Muscle, Phase II Acetylation 13 and Collagen M 2 Food Gut Histamine SOD and Catalase MTHFR, Folate, and Methylation (building) 13 Oxalates 13 Glucuronidati 13 Phase II **Nutrient Metabolism** M Mitochondria Phase I - CYP 13 13 Function **a** M M 13 Phase II Sulfation Glutathione 13 2 13 Glutamate M Neurotransmitters (3) (11) and BH4 M 13 **Urea Cycle** 2 M **MTOR** M NAD+ NADPH 13 Proteins, Vit A & Vit NOS Uncoupling 13 Autophagy 13 13 Fats, Carbs, 2 M M M 13 Phase III Detox 13 Nrf2 Keap1 M Reaction/Ferroptosis M 13 Fenton M

Functional Genomic Analysis Report Makoto Terashita様









# Functional Genomic Analysis Report

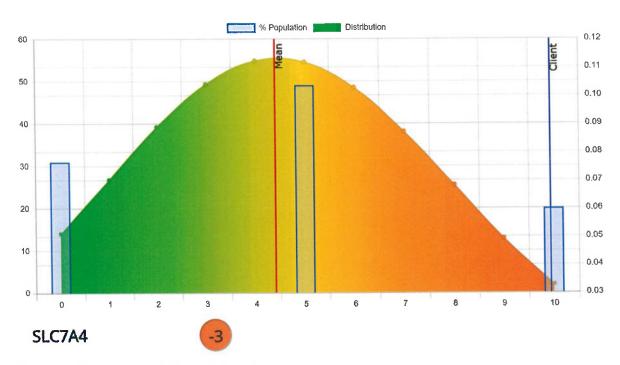
# Makoto Terashita 様

GPX4



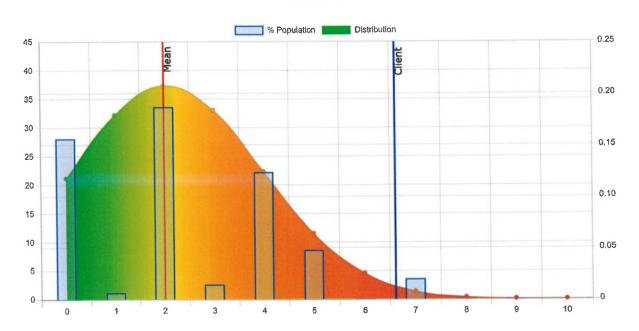
The proteins encoded by GPX genes belong to the glutathione peroxidase family. Glutathione peroxidase functions in the detoxification of hydrogen peroxide, and is one of the most important antioxidant enzymes in humans.

GPX4



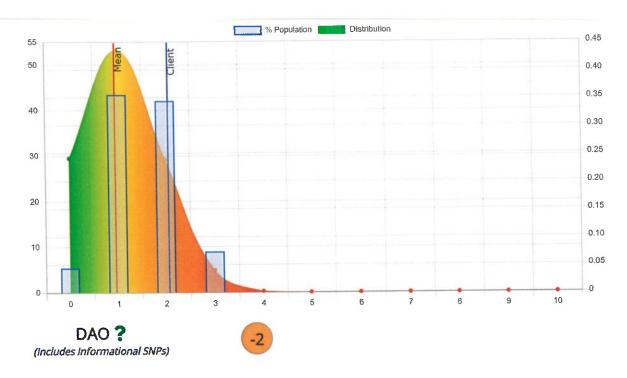
Supports the transportation of Arginine.

SLC7A4



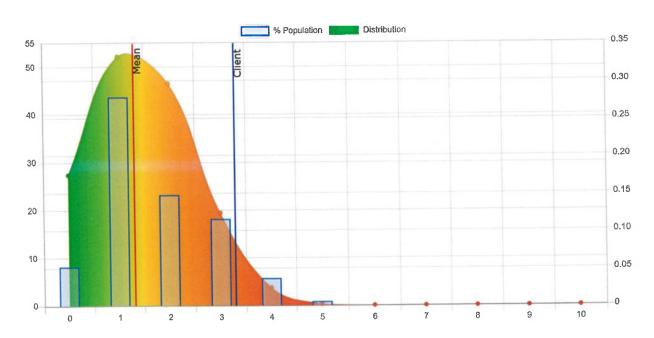
The GAD enzyme converts glutamate to GABA. When someone has high glutamate and a lot of variants in GAD, it creates conditions that may have high glutamate and low GABA that could increase stress and conditions related to high glutamate. It has been observed, that Homozygous variants in GAD have more impact that many Heterozygous. SER-GAB Assist and GABA Assist may be helpful if there is low GABA.

#### GAD2



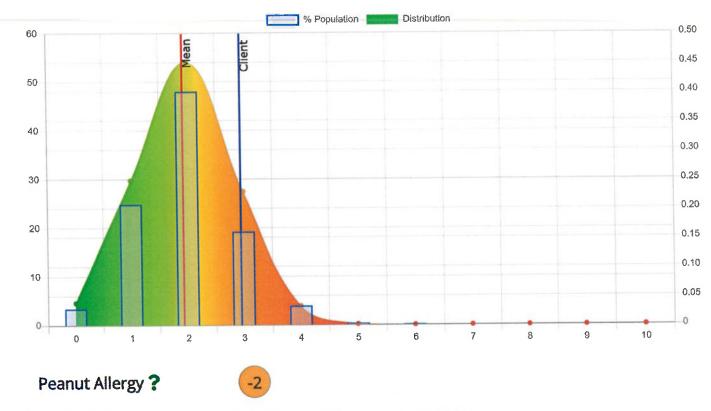
The name of this gene is D-amino-acid oxidase and DAO is the gene's official symbol. Health conditions observed with this variant are: Schizophrenia, Bipolar Disorder, Primary Hyperoxaluria, ALS (Type 18), Autism and Crohn's Disease. Studies have found that the A allele at rs2391191 in DAOA is a possible genetic feature of certain health conditions such as Schizophrenia, and Bipolar Disorder.

#### DAO



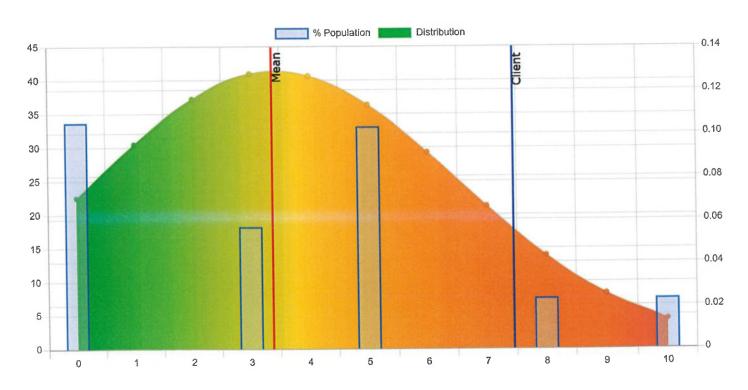
Variants in these genes may increase the chances of Celiac Disease or gluten intolerance. Other factors that may impact gut health are FUT variants that impact probiotics, HNMT and ABP1 variants that lessen histamine degradation and consequently cause zonulin production, low folate or high peroxynitrite.

## HLA



Variations in this gene may cause the potential for peanut sensitivity.

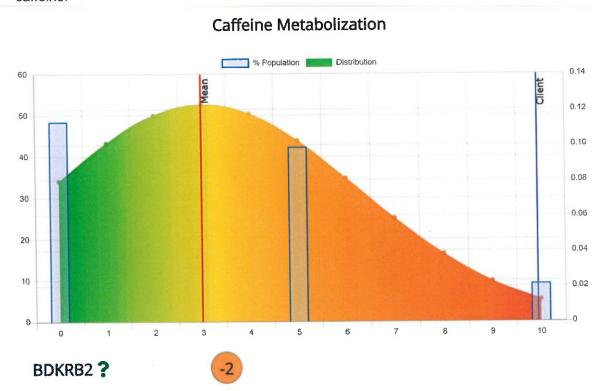
# **Peanut Allergy**



#### Caffeine Metabolization

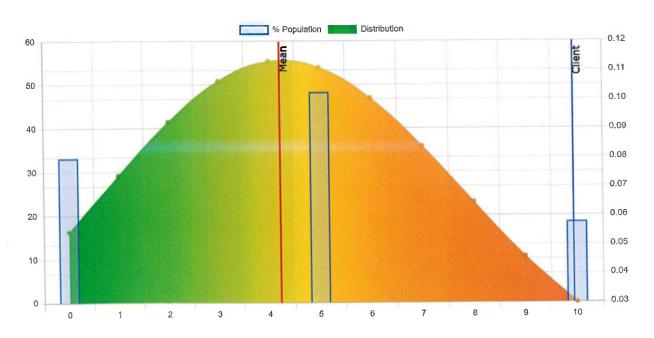


The CYP1A2 gene encodes a member of the cytochrome p450 family of proteins. These proteins metabolize nutrients and drugs. One well known substrate of CYP1A2 is caffeine. Caffeine is a bitter substance that can be found in coffee, tea, soft drinks, chocolate, kola nuts, and certain medicines. It has many effects on the body's metabolism, including stimulating the central nervous system. Studies have shown that individuals with the A allele in this gene are faster metabolizers of caffeine and therefore will feel less of a stimulating effect from caffeine.



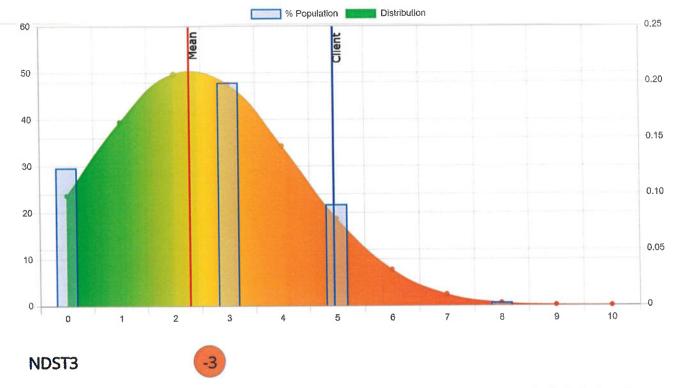
Encoded by BDKRB2, bradykinin receptor B2 (B2R) is a G-protein coupled receptor that appears to be involved in the inflammatory responses. Research suggests B2R forms an inhibitory complex with NOS, which is released following receptor activation. Angiotensin 1-7 appears to potentiate bradykinin-induced B2R responses.

#### **BDKRB2**



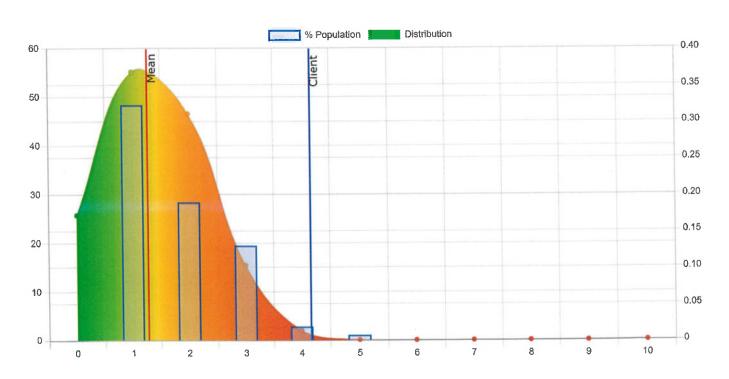
Under normal conditions, Nrf2 is repressed by a negative regulator KEAP1. When cells are exposed to oxidative stress, or electrophiles, Nrf2 escapes KEAP1 and activates the ARE to maintain cellular redox homeostasis.

## KEAP1



Heparan sulfate biosynthetic enzymes are key components in generating a myriad of distinct heparan sulfate fine structures that carry out multiple biologic activities.

# NDST3

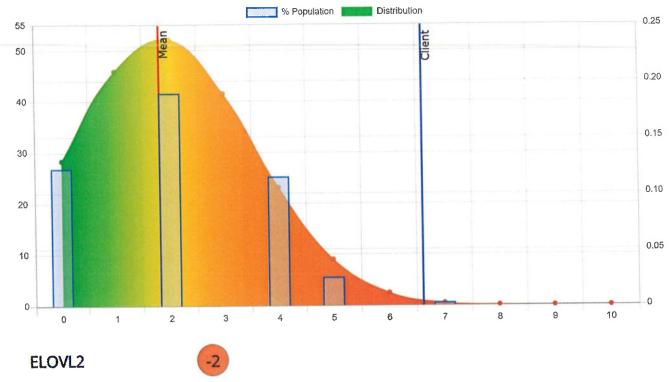


#### **UGT1A4?**

-3

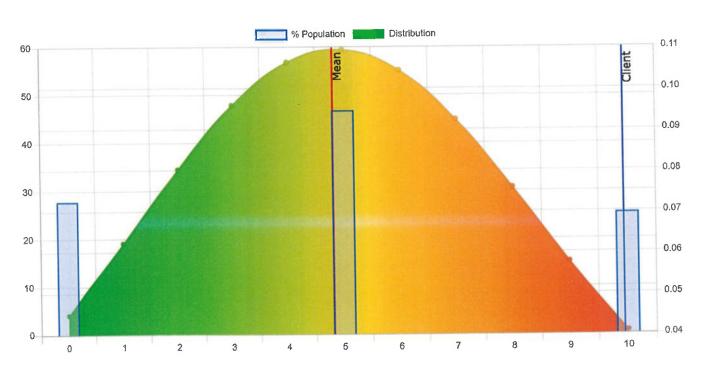
UGT1A4 has some glucuronidase activity towards bilirubin, although it is more active on amines, steroids, and sapogenins.

# UGT1A4



ELOVL2 encodes one of the main enzymes in the in vivo synthesis of long-chain polyunsaturated fatty acids. The G allele of rs953413 has been associated with increased expression of ELOVL2 which may result in an increase in the amount of long-chain polyunsaturated fatty acids synthesized

# **ELOVL2**



APOE

TC

CC

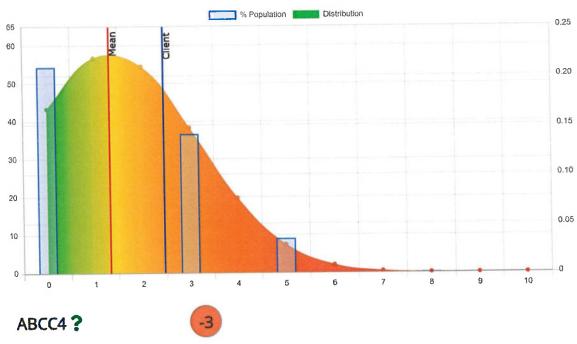


Gene Name	Variants	Metrics
APOE (rs429358)	1 🕒	TC 26.3%
APOE (rs7412)	6	CC 85%

E3/E4

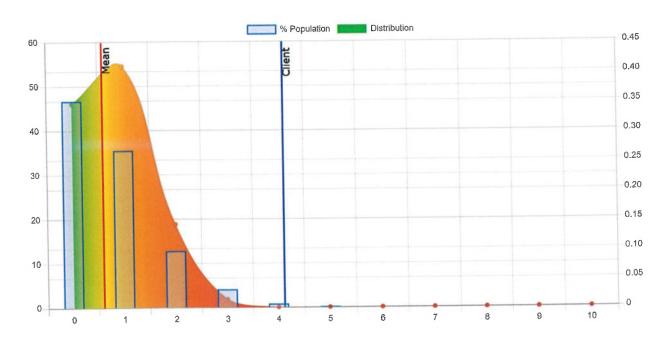
Studies have shown that of all six genotypes, E4/E4 is associated with the greatest risk for high cholesterol. The APO E3/E3 genotype is considered normal. E2 is associated with lower levels of cholesterol. E4 is associated with higher levels of cholesterol.

#### **APOE**



MRP4 is an inducible cotransporter of taurine and glycine conjugates of cholic acid with glutathione, and sulfated bile salts out of cells.

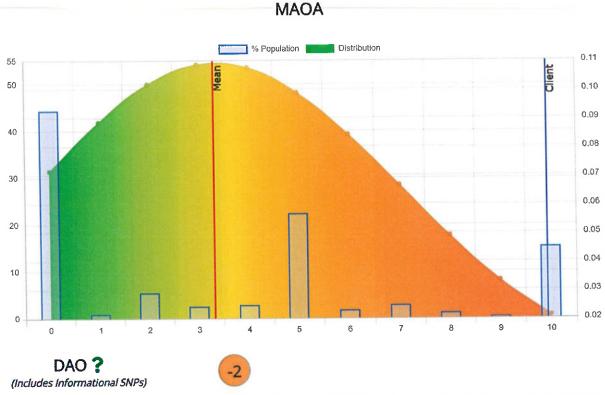
# ABCC4



#### MAOA?

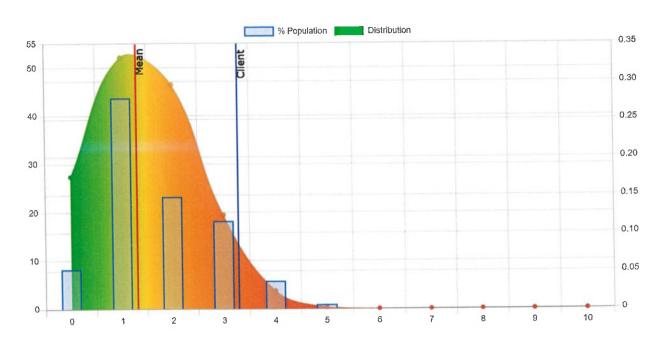


MAOA is one of two neighboring gene family members that encode mitochondrial enzymes which catalyze the oxidative deamination of amines, such as dopamine, norepinephrine, and serotonin. Variants in this gene will actually preserve serotonin. This can be helpful when there is low BH4 and poor availability of amino acids that are the precursors to neurotransmitters. Variations in this gene have also been associated with antisocial behavior



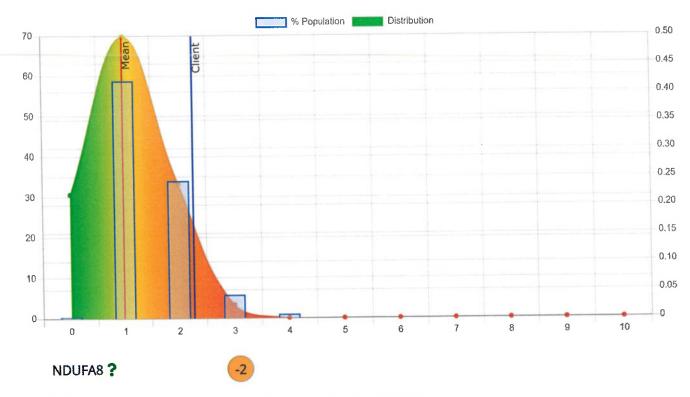
The name of this gene is D-amino-acid oxidase and DAO is the gene's official symbol. Health conditions observed with this variant are: Schizophrenia, Bipolar Disorder, Primary Hyperoxaluria, ALS (Type 18), Autism and Crohn's Disease. Studies have found that the A allele at rs2391191 in DAOA is a possible genetic feature of certain health conditions such as Schizophrenia, and Bipolar Disorder.

#### DAO



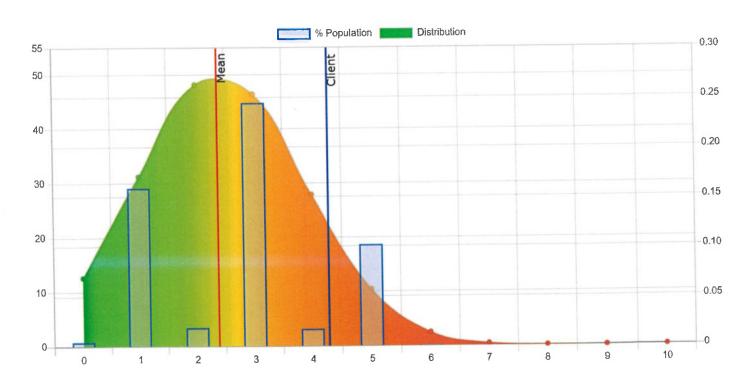
Variants in OGDH may lead to hypotonia, metabolic acidosis, hyperlactatemi, and Alpha-ketoglutarate dehydrogenase deficiency.

# **OGDH**



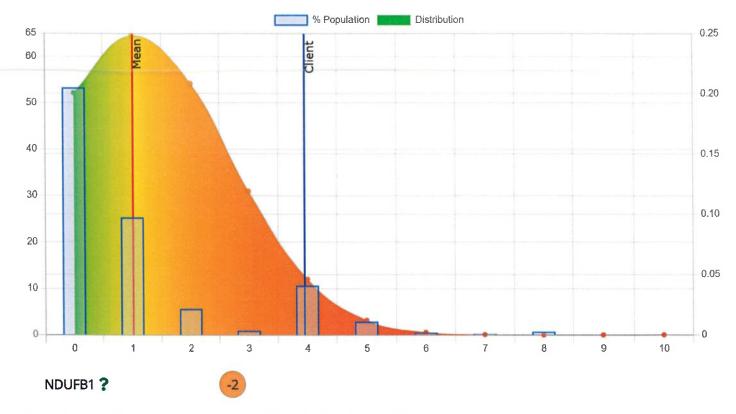
Variations in these genes may cause Complex I to be deficient.

# NDUFA8



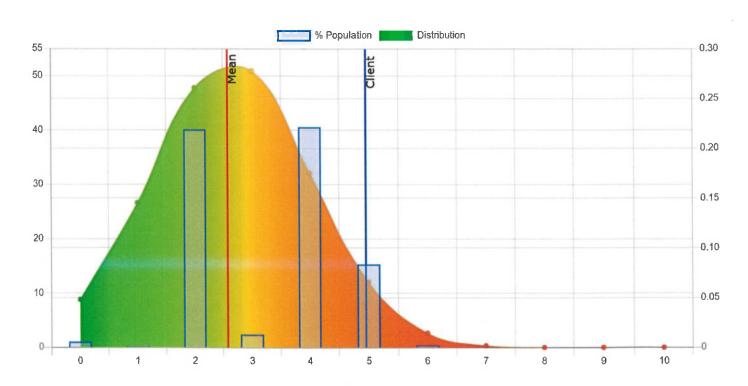
Variations in these genes may cause Complex I to be deficient.

# NDUFAB1



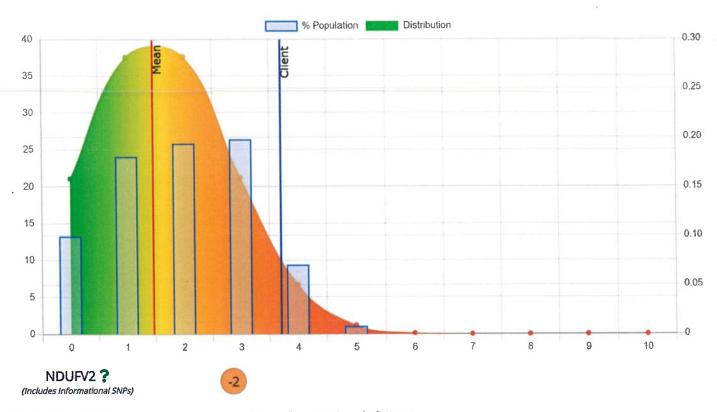
Variations in these genes may cause Complex I to be deficient.

# NDUFB1



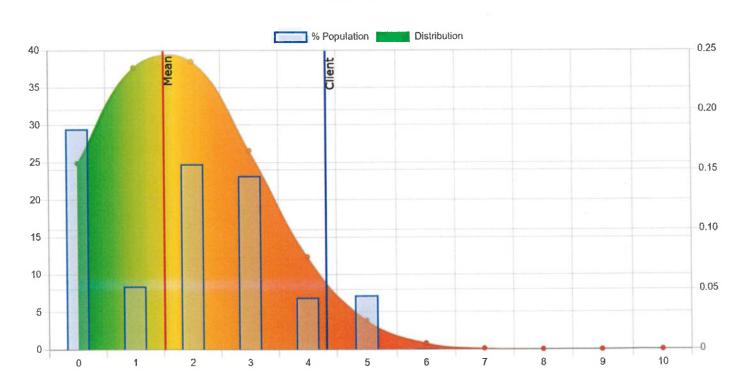
Variations in these genes may cause Complex I to be deficient.

# NDUFS6



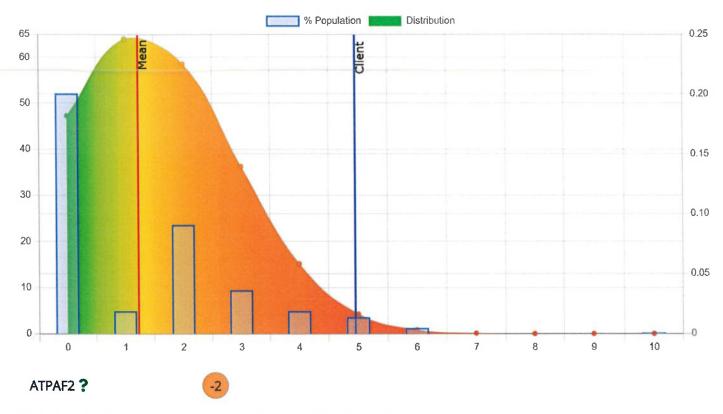
Variations in these genes may cause Complex I to be deficient.

# NDUFV2



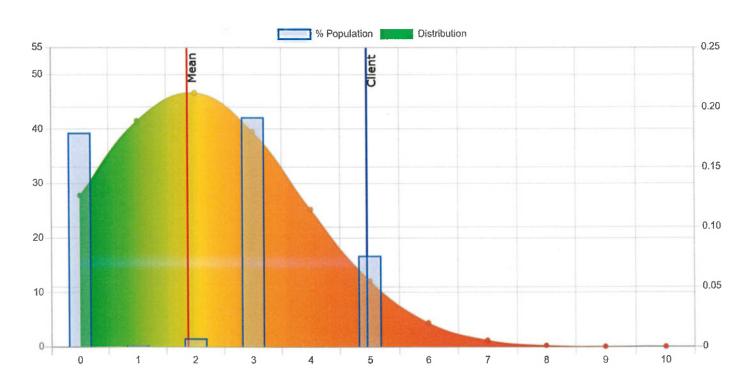
Variations in these genes may cause Complex IV to be deficient.

# **COX411**



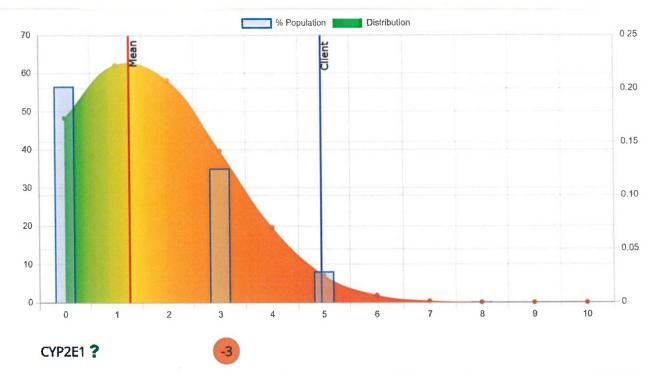
Variations in these genes may cause Complex V to be deficient.

# ATPAF2



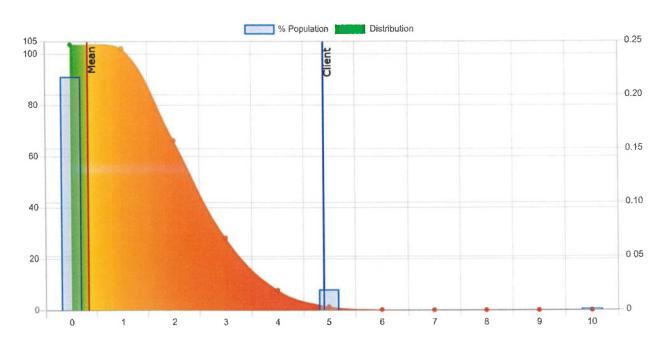
The CYP1A2 gene encodes a member of the cytochrome P450 superfamily of enzymes. CYP1A2 metabolizes polyunsaturated fatty acids into signaling molecules that have physiological as well as pathological activities. Vegetables such as cabbages, cauliflower and broccoli are known to increase levels of CYP1A2.

#### CYP1A2



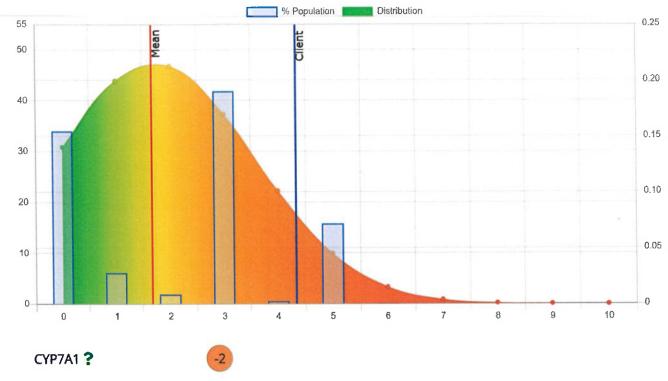
The CYP2E1 gene encodes a member of the cytochrome P450 superfamily of enzymes. CYP2E1 metabolizes both endogenous substrates, such as ethanol, acetone, and acetal, as well as exogenous substrates including benzene, carbon tetrachloride, ethylene glycol, and nitrosamines which are premutagens found in cigarette smoke.

## CYP2E1



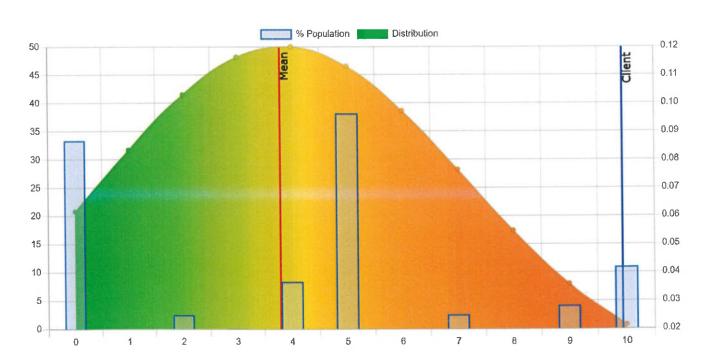
The CYP4X1 gene encodes a member of the cytochrome P450 superfamily of enzymes. The expression pattern of a similar rat protein suggests that CYP4X1 may be involved in neurovascular function in the brain.

## CYP4X1



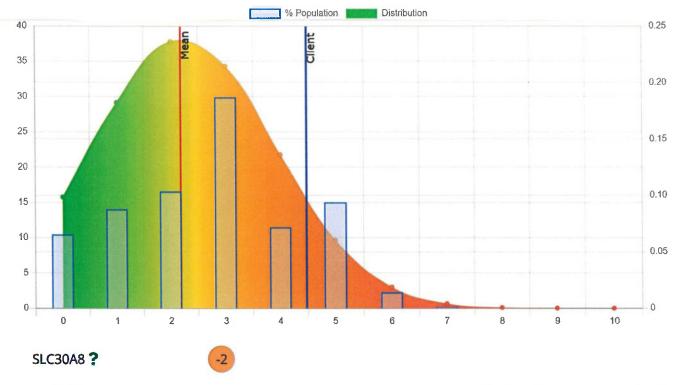
Cholesterol 7 alpha-hydroxylase, encoded by the CYP7A1 gene, catalyzes the first and rate-limiting step of bile acid synthesis from cholesterol. Cholesterol 7 alpha-hydroxylase converts cholesterol to  $7\alpha$ -hydroxycholesterol, a precursor to bile acids.

#### CYP7A1



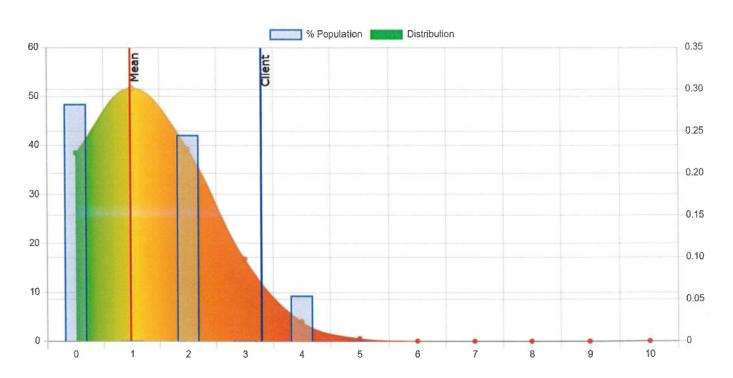
SLC39A14 is involved in the cellular uptake of manganese (Mn) in various cell types, yet has been further demonstrated to have a role in the transportation of Mn from the blood to the liver, for excretion via bile: thus, this enzyme may work synergistically with SLC30A10 for the elimination of manganese.

## **SLC39A14**



Encoded by SLC30A8, zinc transporter 8 (ZNT8) is a zinc transporter involved in the storage of zinc from the cytoplasm into intracellular vesicles. Research suggests ZNT8 may be related to insulin resistance.

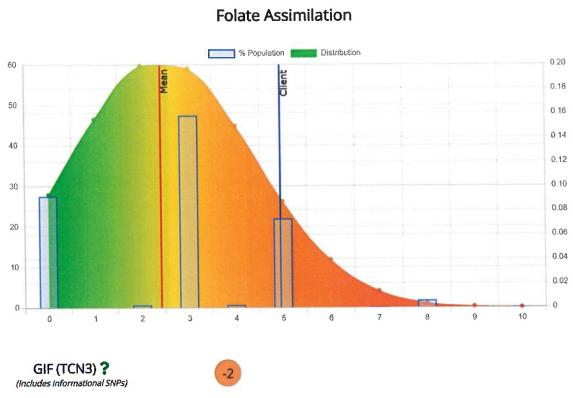
# SLC30A8



#### Folate Assimilation ?



The FOLR1 gene provides instructions for making a protein called folate receptor alpha. This protein helps regulate transport of the B-vitamin folate into cells. Folate (also called vitamin B9) is needed for many processes, including the production and repair of DNA, regulation of gene activity (expression), and protein production. Folate from food is absorbed in the intestines and then released in a form called 5-methyltetrahydrofolate (5-MTHF) into the bloodstream, where it can be taken in by cells in various tissues.



GIF (gastric intrinsic factor) reduces the absorption of B12, while TCN1 and TCN2 limit the transport of B12. Any combination of higher demand, less absorption and transport will impair the functions dependent upon B12.

# GIF (TCN3)

