



Gene Comprehensive Nutrigenomic Report

Accession Number: #####

Specimen Collected: ##/##/####

Specimen Received: ##/##/####

Report Generated: November 30, 2022

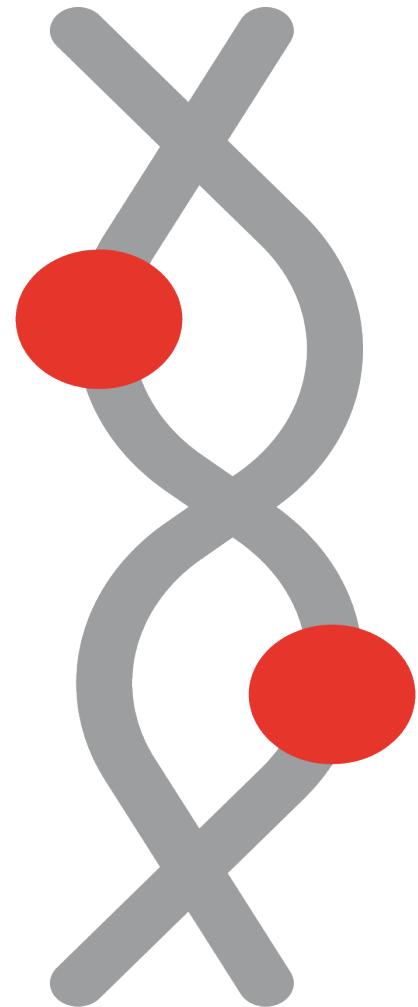
Specimen Type: Buccal Swab

Provider: #####

Patient Name: #####

Patient DOB: ##/##/####

Patient Gender: Female



Do not make any decisions about your health solely based on the information contained in this report.
Always consult with a licensed and experienced health practitioner when you receive this report.

- 28 – Female

(-/-) No clinical abnormality (+/-) Heterozygous result (+/+) Homozygous result

rsID	Gene	Genetic Result	Therapeutics Associated With Positive Result	Highly Recommended Therapeutics	Provider Discretion: As Needed Formula Recommendations	Lifestyle Recommendations	Laboratory Recommendations
Neurotransmitters							
rs4680	COMT V158M	+/-	Taurine, Choline, Trimethylglycine (TMG), Dimethylglycine (DMG), Methionine, SAMe, Inositol		May Benefit from Full Focus+™ if Anxiety or Depression Present	Higher Risk of Depression / Anxiety with Stressful Events	Consider Neurotransmitter Metabolite Testing
rs769407	GAD1	-/-	Prescription Amantadine, Glycine, N-Acetyl-Cysteine (NAC), Zinc, Magnesium, Elderberry, L-Theanine, Melatonin	May Benefit from Prescription Amantadine	May Benefit from Pro GAD Enhancer™ if Anxiety is Present May Benefit from Neuro Night Essentials™ if Sleep Initiation is Problematic	Be cautious with MSG (Monosodium Glutamate) Be cautious with Glutamine Supplementation	Consider Neurotransmitter Metabolite Testing
rs3828275	GAD1	+/-	B2 (Riboflavin), Methyl Donors (Taurine, Choline, Trimethylglycine (TMG), Dimethylglycine (DMG), Inositol, Methionine				
rs6323	MAO-A	+/-	Methyl Donors (Taurine, Choline, Trimethylglycine (TMG), Dimethylglycine (DMG), Inositol, Methionine		May Benefit from Full Focus+™ if Anxiety or Depression Present		
rs1799836	MAO-B	-/-					

- 28 – Female

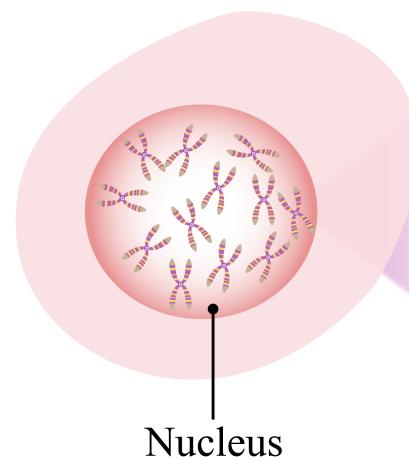
(-/-) No clinical abnormality (+/-) Heterozygous result (+/+) Homozygous result

rsID	Gene	Genetic Result	Therapeutics Associated With Positive Result	Highly Recommended Therapeutics	Provider Discretion: As Needed Formula Recommendations	Lifestyle Recommendations	Laboratory Recommendations
rs6313	HTR2	+/-	5-HTP (Hydroxytryptophan)		May Benefit from Mood Plus™ if Anxiety or Depression Present	May Have Less Than Expected Efficacy To SSRI Medications	Consider PGx Testing
rs1042173	SLC6A4	+/+					
rs4570625	TPH2	+/-	Niacinamide, 5-HTP				
rs1108580	DBH	+/-	Droxidopa Phenylpropanolamine Pseudoephedrine Fludrocortisone Strattera				Consider Neurotransmitter Metabolite Testing
Methylation for Neurotransmitters							
rs1801131	MTHFR 1298	-/-	Methyltetrahydrofolate (5-MTHF)	Methyl Folate Plus twice daily			
rs1801133	MTHFR 677	+/-					

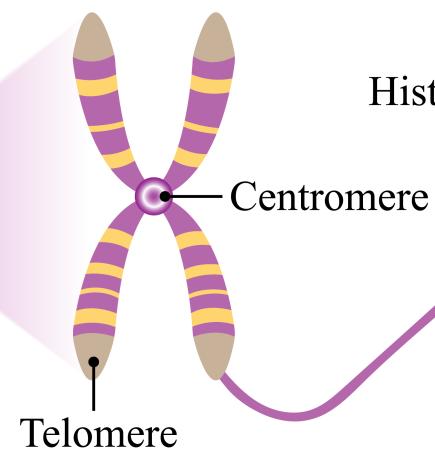
Summary for Neurotransmitters / Mood

Highly Recommended Therapeutics	Provider Discretion: As Needed Formula Recommendations	Lifestyle Recommendations	Laboratory Recommendations
<ul style="list-style-type: none">• May Benefit from Prescription Amantadine• Methyl Folate Plus twice daily	<ul style="list-style-type: none">• May Benefit from Full Focus+™ if Anxiety or Depression Present• May Benefit from Pro GAD Enhancer™ if Anxiety is Present• May Benefit from Neuro Night Essentials™ if Sleep Initiation is Problematic• May Benefit from Full Focus+™ if Anxiety or Depression Present• May Benefit from Mood Plus™ if Anxiety or Depression Present	<ul style="list-style-type: none">• Higher Risk of Depression / Anxiety with Stressful Events• Be cautious with MSG (Monosodium Glutamate)• Be cautious with Glutamine Supplementation• May Have Less Than Expected Efficacy To SSRI Medications	<ul style="list-style-type: none">• Consider Neurotransmitter Metabolite Testing• Consider PGx Testing

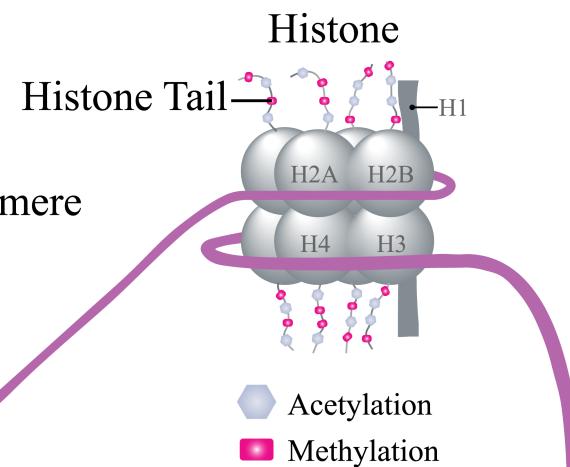
Cell



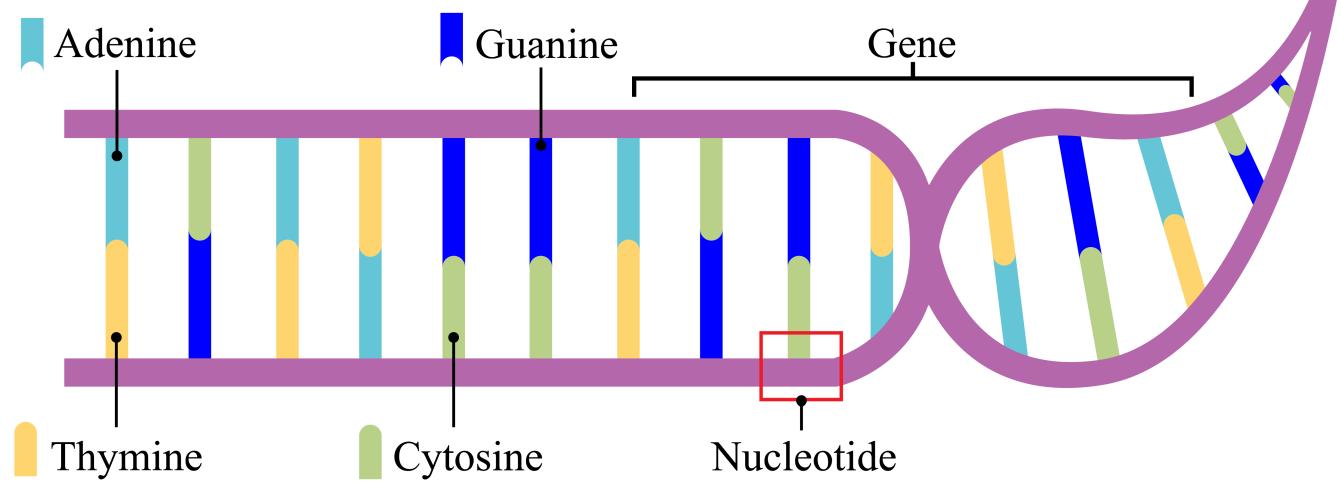
Chromosome



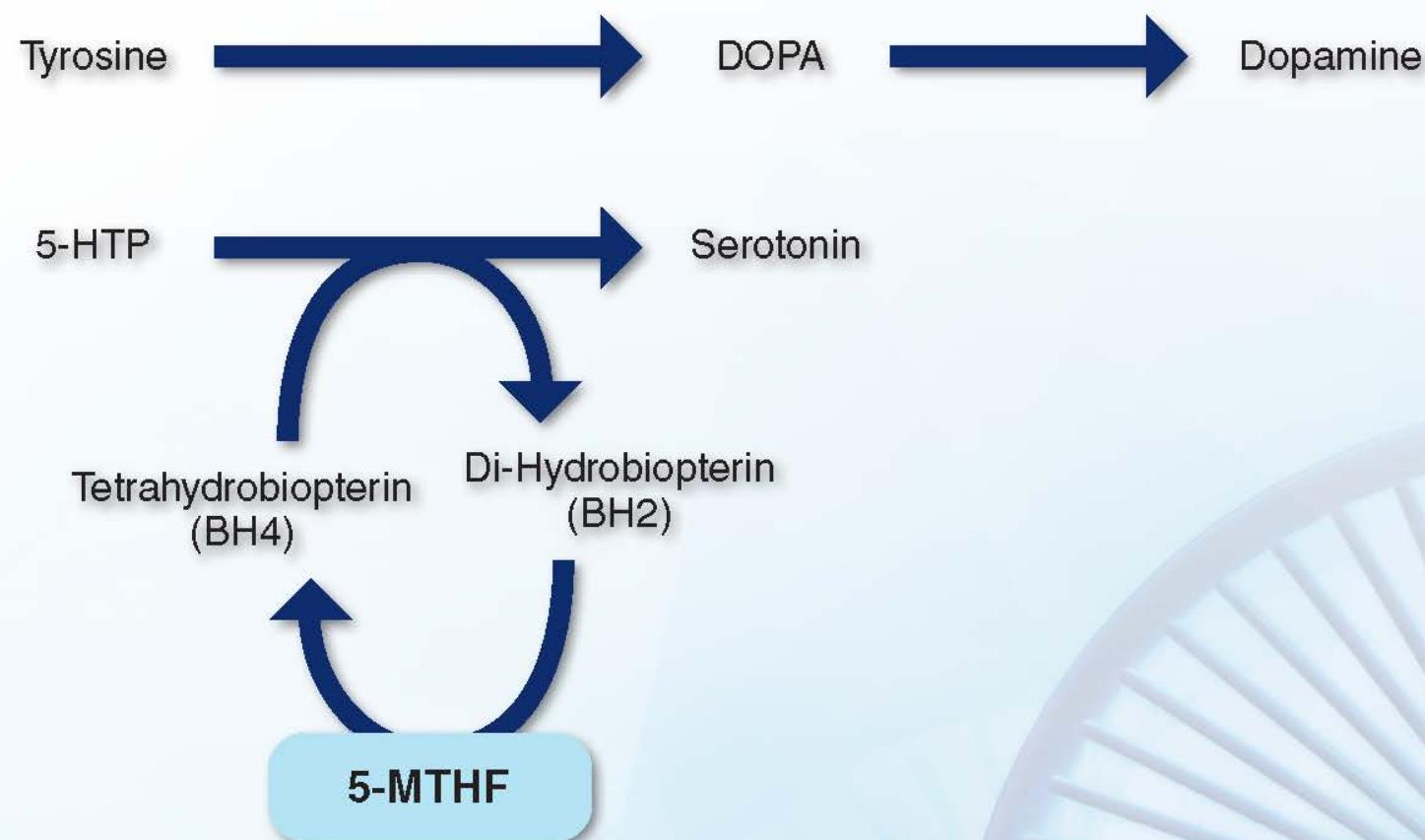
Nucleosome



DNA

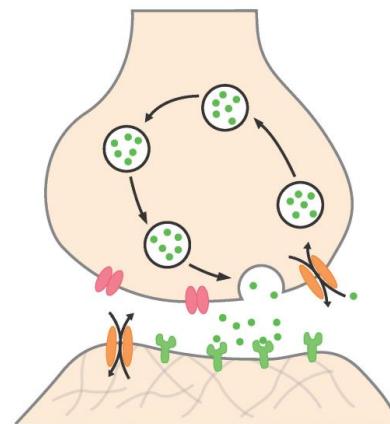


5-MTHF & Neurotransmitter Production



NEUROTRANSMITTERS & PATHWAY

TRANSMIT INFORMATION FOR ESSENTIAL PROCESSES SUCH AS DIGESTION, BREATHING, HEARTBEAT, MOVEMENT, PAIN REGULATION ETC.



RELEVANT GENES

- **HTR2, TPH2, SLC6A4, MAO-A** genes are important in the synthesis, breakdown, transport and/or functioning of serotonin
 - **COMT, MAO-A, MAO-B** genes are important for the breakdown of serotonin, norepinephrine and/or dopamine
 - The **DBH** gene is important for norepinephrine synthesis
- The **GAD1** gene is important for GABA synthesis
 - Variants in **COMT, MAO-A, MAO-B** and **GAD1** genes have been associated with mood, anxiety and focus issues

WAYS TO INCREASE LEVELS



Aerobic Exercise



Dietary Factors



Mediation/Yoga



Increase Sun Exposure

Gene Information Key

rsID	Gene	"-" variant	"+" variant
rs4680	COMT V158M	G	A
rs1108580	DBH	A	G
rs3828275	GAD1	C	T
rs769407	GAD1	G	C
rs6313	HTR2	G	A
rs6323	MAO-A	T	G
rs1799836	MAO-B	T	C
rs1801131	MTHFR 1298	T	G
rs1801133	MTHFR 677	G	A
rs1042173	SLC6A4	A	C
rs4570625	TPH2	G	T

Definitions

METHYLATION	Methylation is a primary biochemical process in the body that involves the addition of a "methyl" chemical group to a vitamin or neurotransmitter. The addition of the "methyl" group allows for very specific biochemical interactions. Poor "methylation" function alters the effectiveness, delivery and function of many vitamins and important chemicals in the cell.
MTHFR A1298C	Methylenetetrahydrofolate reductase (MTHFR) catalyzes the conversion of 5,10-methylenetetrahydrofolate to 5-methyltetrahydrofolate, the bioactive form of folic acid. Two significant polymorphism variants exist in this gene, the A1298C and the C677T. The 1298 confers a conversion weakness of 10% for one copy and approximately 20% for two copies. In contrast, the 677 variant is much more severe and conveys a 40% conversion weakness for one copy and 70% for two copies. A reduced level of MTHFolate produces significant biochemical effects including poor production of dopamine and serotonin, pregnancy complications, poor healing of the nervous system, weak mitochondrial function, reduced production of glutathione, poor cell turnover and poor function of T cell lymphocytes.
MTHFR C677T	Methylene tetrahydrofolate reductase (MTHFR) catalyzes the conversion of 5,10-methylenetetrahydrofolate to 5-methyltetrahydrofolate, the bioactive form of folic acid. Two significant polymorphism variants exist in this gene, the A1298C and the C677T. The 1298 confers a conversion weakness of 10% for one copy and approximately 20% for two copies. In contrast, the 677 variant is much more severe and conveys a 40% conversion weakness for one copy and 70% for two copies. A reduced level of MTHFolate produces significant biochemical effects including poor production of dopamine and serotonin, pregnancy complications, poor healing of the nervous system, weak mitochondrial function, reduced production of glutathione, poor cell turnover and poor function of T cell lymphocytes.
NEUROTRANSMITTER	Neurotransmitters are chemicals that are used to produce specific effects in the nervous system. These specific neurotransmitter genomics assess a person's risk for anxiety, depression and dysphoria.
COMT V158M	Catechol-O-methyltransferase (COMT) is one of several enzymes that degrade catecholamine neurotransmitters such as dopamine, epinephrine, and norepinephrine. COMT's main function is to inactivate neurotransmitters (dopamine, epinephrine, and norepinephrine) by the addition of a methyl group to the catecholamine. Normal COMT function allows people to rapidly reverse feelings of anxiety or depression. COMT (+/-) patients have sluggish ability to alter anxiety or depression episodes. COMT (+/+) patients are more prone to prolonged episodes of anxiety, depression and OCD.
DBH	DBH (Dopamine Beta Hydroxylase) is an oxidoreductase belonging to the copper type II, ascorbate-dependent monooxygenase family. The encoded protein, expressed in neurosecretory vesicles catalyzes the conversion of dopamine to norepinephrine, which functions as both a hormone and sympathetic nervous system function. Polymorphisms in this gene lower the production of norepinephrine which causes poor autonomic and cardiovascular function, including hypotension and ptosis. Polymorphisms in this gene have also been linked to Autism, ADD, bipolar disorder and major depression.
GAD1 rs3828275	Glutamic Acid Decarboxylase (GAD 1) is the enzyme responsible for conversion of glutamic acid (a stimulant neurotransmitter) to GABA (a calming neurotransmitter). Deficiency of GABA from polymorphisms in this enzyme are associated with sleep disorders, "half glass empty" syndrome, dysphoria, and spasticity.
HTR2A	5-hydroxytryptamine receptor 2 (HTR2) is one of the neuronal receptors for the neurotransmitter serotonin. Mutations in the HTR2 gene are associated with individual response to antidepressants, appetite, and mood.
MAOA	Monoamine oxidase A (MAOA) is one of the classic neurotransmitter degradation enzymes. By degrading serotonin, dopamine, epinephrine, and norepinephrine, MAO-A ends neuronal signaling induced by those neurotransmitters. Mutations in the MAO-A gene leads to decreased degradation of these neurotransmitters and can be associated with increased aggression, mood disorders and drug addiction.
MAOB	Monoamine Oxidase B (MAO B) catalyzes the neuroactive amines, such as dopamine, epinephrine, norepinephrine, and plays a role in the stability of mood in the central nervous system.. MAO B's primary purpose is to degrade dopamine. Patients who possess polymorphisms of MAO B have a higher risk of clinical depression and mood disorders.
SLC6A4	The SLC6A4 gene encodes the serotonin transporter, also known as SERT. The serotonin transporter is responsible for clearing the serotonin neurotransmitter from the synaptic space. SERT is the target of many therapeutic drugs. Polymorphisms in the SLC6A4 gene are associated with increased risk of anxiety and depression and less effective response to SSRI medications.
TPH2	TPH2 (Tryptophan Hydroxylase 2) gene catalyzes the first and rate limiting step in the biosynthesis of serotonin (5HT), an important hormone and neurotransmitter. Mutations in this gene have been shown to be associated with psychiatric diseases such as bipolar affective disorder, anxiety and major depression. Polymorphisms in this gene are also correlated to an increased response rate to SSRI medications.

Disclaimers

TESTING:

Testing Performed By: AMH

METHODOLOGY AND LIMITATIONS:

Testing for genetic variation/mutation on listed genes was performed using ProFlex PCR and Real-Time PCR with TaqMan® allele-specific probes on the QuantStudio 12K Flex. All genetic testing is performed by GX Sciences, 4150 Freidrich Lane, Ste H, Austin, TX. 78744. This test will not detect all the known alleles that result in altered or inactive tested genes. This test does not account for all individual variations in the individual tested. Test results do not rule out the possibility that this individual could be a carrier of other mutations/variations not detected by this gene mutation/variation panel. Rare mutations surrounding these alleles may also affect our detection of genetic variations. Thus, the interpretation is given as a probability. Therefore, this genetic information shall be interpreted in conjunction with other clinical findings and familial history for the administration of specific nutrients. Patients should receive appropriate genetic counseling to explain the implications of these test results. Details of assay performance and algorithms leading to clinical recommendations are available upon request. The analytical and performance characteristics of this laboratory developed test (LDT) were determined by GX Sciences' laboratory pursuant to Clinical Laboratory Improvement Amendments (CLIA) requirements.

CLIA #: 45D2144988 Laboratory Director: James Jacobson, PhD

DISCLAIMER:

This test was developed and its performance characteristics determined by GX Sciences. It has not been cleared or approved by the FDA. The laboratory is regulated under CLIA and qualified to perform high-complexity testing. This test is used for clinical purposes. It should not be regarded as investigational or for research. rsIDs for the alleles being tested were obtained from the dbSNP database (Build 142).

DISCLAIMER:

UND Result: If you have received the result Variant undetermined (UND) this indicates that we were not able to determine your carrier status based on your raw data. Please refer to the GX Sciences genetic knowledge database for more information: https://www.gxsciences.com/kb_results.asp

DISCLAIMER:

Report contents and report recommendations are created and approved by GX Sciences. Sole responsibility for the proper use of the information on the GX Sciences report rests with the user, or those professionals with whom the user may consult. Nutrigenomic Testing and Dietary Supplements are not "Designated Health Services" covered by Medicare or Medicaid and may not be reimbursed under any state or Federal health care program.

DISCLAIMER:

These products are not approved by the Food and Drug Administration and are not intended to diagnose, treat, cure or prevent disease. These recommendations are for report purposes only and an individual is not required to use such products. These are recommendations only and do not replace the advisement of your own healthcare practitioner.

GX Sciences SNP References

METHYLATION SNP References

MTHFR

- Mischoulon, D. & Raab, M. F. The role of folate in depression and dementia. *Journal of Clinical Psychiatry* (2007).
- Goyette, P. et al. Human methylenetetrahydrofolate reductase: Isolation of cDNA, mapping and mutation identification. *Nat. Genet.* (1994). doi:10.1038/ng0694-195 • Hua, Y., Zhao, H., Kong, Y. & Lu, X. Association between Alzheimer's disease and the NOS3 gene Glu298Asp polymorphism. *Int. J. Neurosci.* (2014). doi:10.3109/00207454.2013.834336 • Tran, P. et al. Multiple transcription start sites and alternative splicing in the methylenetetrahydrofolate reductase gene result in two enzyme isoforms. *Mamm. Genome* (2002). doi:10.1007/s00335-002-2167-6 • Földinger, M., Hörl, W. H. & Sunder-Plassmann, G. Molecular biology of 5,10-methylenetetrahydrofolate reductase. *Journal of Nephrology* (2000). doi:10.5860/CHOICE.39-4838 • Trimmer, E. Methylenetetrahydrofolate Reductase: Biochemical Characterization and Medical Significance. *Curr. Pharm. Sci.* (2013). doi:10.2174/1381612811319140008 • Wu, X. et al. Association Between the MTHFR C677T Polymorphism and Recurrent Pregnancy Loss: A Meta-Analysis. *Genet. Test. Mol. Biomarkers* (2012). doi:10.1089/gtmb.2011.0318 • Voutilainen, S., Rissanen, T. H., Virtanen, J., Lakka, T. A. & Salonen, J. T. Low dietary folate intake is associated with an excess incidence of acute coronary events: The Kuopio Ischemic Heart Disease Risk Factor Study. *Circulation* (2001). doi:10.1161/01.CIR.103.22.2674 • Ducker, G. S. & Rabinowitz, J. D. One-Carbon Metabolism in Health and Disease. *Cell Metabolism* (2017). doi:10.1016/j.cmet.2016.08.009 • Papakostas, G. I. et al. Effect of adjunctive L-methylfolate 15 mg among inadequate responders to SSRIs in depressed patients who were stratified by Biomarker levels and genotype: Results from a randomized clinical trial. *Journal of Clinical Psychiatry* (2014). doi:10.4088/JCP.13m08947 • Schneider, J. A., Rees, D. C., Liu, Y.-T. & Clegg, J. B. Worldwide Distribution of a Common Methylenetetrahydrofolate Reductase Mutation. *The American Journal of Human Genetics* 62, 1258–1260 (1998).
- Moore, L. D., Le, T. & Fan, G. DNA methylation and its basic function. *Neuropharmacology* (2013). doi:10.1038/npp.2012.112 • Scaglione, F. & Panzavolta, G. Folate, folic acid and 5-methyltetrahydrofolate are not the same thing. *Xenobiotica* (2014). doi:10.3109/00498254.2013.845705 • 5-Methyl-tetrahydrofolate. National Center for Biotechnology Information. PubChem Compound Database (2004). Available at: <https://pubchem.ncbi.nlm.nih.gov/compound/135483998>.
- Office of Dietary Supplements - Folate. NIH Office of Dietary Supplements (2020). Available at: <https://ods.od.nih.gov/factsheets/Folate-HealthProfessional/> • Sibani, S. et al. Characterization of six novel mutations in the methylenetetrahydrofolate reductase (MTHFR) gene in patients with homocystinuria. *Hum. Mutat.* (2000). doi:10.1002/(SICI)1098-1004(200003)15:33.O;CO;2-1 • Goyette, P. et al. Human methylenetetrahydrofolate reductase: Isolation of cDNA, mapping and mutation identification. *Nat. Genet.* (1994). doi:10.1038/ng0694-195 • Abu Seman, N., Wan Mohamad, W. N., Östenson, C. G., Brisman, K. & Gu, H. F. Increased dna methylation of the slc30a8 gene promoter is associated with type 2 diabetes in a malay population. *Clin. Epigenetics* (2015). doi:10.1168/13148-15-0049-5 • Link, R. 15 Healthy Foods That Are High in Folate (Folic Acid). *Healthline* (2020). Available at: <https://www.healthline.com/nutrition/foods-high-in-folate-folic-acid#6-Citrus-fruits> • Matthews, R. G. & Daubner, S. C. Modulation of methylenetetrahydrofolate reductase activity by Sadenosylmethionine and by dihydrofolate and its polyglutamate analogues. *Adv. Enzyme Regul.* (1982). doi:10.1016/0065-2571(82)90012-7 • Yamada, K., Strahler, J. R., Andrews, P. C. & Matthews, R. G. Regulation of human methylenetetrahydrofolate reductase by phosphorylation. *Proc. Natl. Acad. Sci.* (2005). doi:10.1073/pnas.0504786102 • Tanaka, T. et al. Genome-wide Association Study of Vitamin B6, Vitamin B12, Folate, and Homocysteine Blood Concentrations. *Am. J. Hum. Genet.* (2009). doi:10.1016/j.ajhg.2009.02.011 • Schwahn, B. & Rozen, R. Polymorphisms in the methylenetetrahydrofolate reductase gene: Clinical consequences. *American journal of pharmacogenomics: genomics-related research in drug development and clinical practice* (2001). doi:10.2165/00129785-200101030-00004 • Yamada, K., Chen, Z., Rozen, R. & Matthews, R. G. Effects of common polymorphisms on the properties of recombinant human methylenetetrahydrofolate reductase. *Proc. Natl. Acad. Sci.* (2001). doi:10.1073/pnas.261469998 • Voutilainen, S., Rissanen, T. H., Virtanen, J., Lakka, T. A. & Salonen, J. T. Low dietary folate intake is associated with an excess incidence of acute coronary events: The Kuopio Ischemic Heart Disease Risk Factor Study. *Circulation* (2001). doi:10.1161/01.CIR.103.22.2674 • Bailey, L. B. Folate, Methyl-Related Nutrients, Alcohol, and the MTHFR 677C>T Polymorphism Affect Cancer Risk: Intake Recommendations. *J. Nutr.* (2003). doi:10.1093/jn/133.11.3748S • Nishiyama, M., Kato, Y., Hashimoto, M., Yukawa, S. & Omori, K. Apolipoprotein E, Methylenetetrahydrofolate Reductase(MTHFR) Mutation and the Risk of Senile Dementia: An Epidemiological Study Using the Polymerase Chain Reaction(PCR) Method. *J. Epidemiol.* (2000). doi:10.2188/jea.10.163 • Reilly, R., McNulty, H., Penteiva, K., Strain, J. J. & Ward, M. MTHFR 677TT genotype and disease risk: Is there a modulating role for B-vitamins'. *Proc. Nutr. Soc.* (2014). doi:10.1017/S0029665113003613

NEUROTRANSMITTER SNP References

COMT

- Lotta, T. et al. Kinetics of Human Soluble and Membrane-Bound Catechol O-Methyltransferase: A Revised Mechanism and Description of the Thermolabile Variant of the Enzyme. *Biochemistry* (1995). doi:10.1021/bi00013a008 • Stein, M. B., Fallon, M. D., Schork, N. J. & Gelernter, J. COMT polymorphisms and anxiety-related personality traits. *Neuropharmacology* (2005). doi:10.1038/sj.npp.0507877 • Lee, L. O. & Prescott, C. A. Association of the catechol-O-methyltransferase val158met polymorphism and anxiety-related traits: A meta-analysis. *Psychiatr. Genet.* (2014). doi:10.1097/YPG.0000000000000018 • Ulmanen, I. et al. Expression and intracellular localization of catechol O-methyltransferase in transfected mammalian cells. *Eur. J. Biochem.* (1997). doi:10.1111/j.1423-1033.1997.0452a.x • Axelrod, J. O-methylation of epinephrine and other catechols in vitro and in vivo. *Science* (80-). (1957). doi:10.1126/science.126.3270.400 • Tai, C. H. & Wu, R. M. Catechol-O-methyltransferase and Parkinson's disease. *Acta Medica Okayama* (2002).
- Grossman, M. H., Emanuel, B. S. & Budarf, M. L. Chromosomal mapping of the human catechol-O-methyltransferase gene to 22q11.1q11.2. *Genomics* (1992). doi:10.1016/0888-7543(92)90316-K • Golani, D. E., Armstrong, E. J. & Armstrong, A. W. Principles of pharmacology: the pathophysiology basis of drug therapy. (Wolters Kluwer Health, 2017).
- Bonifácio, M. J., Palma, P. N., Almeida, L. & Soares-Da-Silva, P. Catechol-O-methyltransferase and its inhibitors in Parkinson's disease. *CNS Drug Reviews* (2007). doi:10.1111/j.1527-3458.2007.00020.x • Wicher, M. et al. The catechol-O-methyl transferase Val158Met polymorphism and experience of reward in the flow of daily life. *Neuropharmacology* (2008). doi:10.1038/sj.npp.1301520 • Diamond, A., Briand, L., Fossella, J. & Gehlbach, L. Genetic and Neurochemical Modulation of Prefrontal Cognitive Functions in Children. *Am. J. Psychiatry* (2004). doi:10.1176/appi.ajp.161.1.125 • Robinson, S., Goddard, L., Dritschel, B., Wisley, M. & Howlin, P. Executive functions in children with Autism Spectrum Disorders. *Brain Cogn.* (2009). doi:10.1016/j.bandc.2009.06.007 • Bruder, G. E. et al. Catechol-O-methyltransferase (COMT) genotypes and working memory: Associations with differing cognitive operations. *Biol. Psychiatry* (2005). doi:10.1016/j.biopsych.2005.05.010 • Chen, J. et al. Functional analysis of genetic variation in catechol-O-methyltransferase (COMT): Effects on mRNA, protein, and enzyme activity in postmortem human brain. *Am. J. Hum. Genet.* (2004). doi:10.1086/425589

DBH

- Rahman, M. K., Rahman, F., Rahman, T. & Kato, T. Dopamine-β-hydroxylase (DBH), its cofactors and other biochemical parameters in the serum of neurological patients in Bangladesh. *Int. J. Biomed. Sci.* (2009). doi:10.1016/j.jicard.2009.09.092 • Barrie, E. S., Weinshenker, D., Verma, A., Pendergrass, S. A., Lange, L. A., Ritchie, M. D. ... Sadee, W. (2014). Regulatory polymorphisms in human DBH affect peripheral gene expression and sympathetic activity. *Circulation Research*. <https://doi.org/10.1161/CIRCRESAHA.116.304398> • Das, M., Bhownik, A. Das, Bhaduri, N., Sarkar, K., Ghosh, P., Sinha, S. ... Mukhopadhyay, K. (2011). Role of gene/gene-environment interaction in the etiology of eastern Indian ADHD probands. *Progress in Neuro-Psychopharmacology and Biological Psychiatry*. <https://doi.org/10.1016/j.pnpbp.2010.12.027> • Das Bhownik, A., Sarkar, K., Ghosh, P., Das, M., Bhaduri, N., Sarkar, K. ... Mukhopadhyay, K. (2017). Significance of Dopaminergic Gene Variants in the Male Biasness of ADHD. *Journal of Attention Disorders*. <https://doi.org/10.1177/1087054713494004> • Fang, Y., Ji, N., Cao, Q., Su, Y., Chen, M., Wang, Y., ... Yang, L. (2015). Variants of Dopamine Beta Hydroxylase Gene Moderate Atomoxetine Response in Children with Attention-Deficit/Hyperactivity Disorder. *Journal of Child and Adolescent Psychopharmacology*. <https://doi.org/10.1089/cap.2014.0178> • Parasuraman, R., de Visser, E., Lin, M. K., & Greenwood, P. M. (2012). Dopamine beta hydroxylase genotype identifies individuals less susceptible to bias in computer-assisted decision making. *PLOS ONE*. <https://doi.org/10.1371/journal.pone.0039675> • Sezer, S., Kurt, S., & Ates, O. (2016). Analysis of dopamine beta hydroxylase gene polymorphisms in migraine. *Clinical Neurology and Neurosurgery*. <https://doi.org/10.1016/j.clineuro.2016.02.002> • Sun, Z., Ma, Y., Li, W., He, J., Li, J., Yang, X., ... Tang, Y. L. (2018). Associations between the DBH gene, plasma dopamine β-hydroxylase activity and cognitive measures in Han Chinese patients with schizophrenia. *Schizophrenia Research*. <https://doi.org/10.1016/j.schres.2017.06.028> • Sundararajan, R. (2013). Effect of DBH, DRD2 and ADRA2A gene variants on human working memory. *Dissertation Abstracts International: Section B: The Sciences and Engineering*.

GAD1

- GAD1 glutamate decarboxylase 1 [Homo sapiens (human)] - Gene - NCBI. National Center for Biotechnology Information (2020). Available at: <https://www.ncbi.nlm.nih.gov/gene/2571>.
- KELLY, C. D. et al. Nucleotide sequence and chromosomal assignment of a cDNA encoding the large isoform of human glutamate decarboxylase. *Ann. Hum. Genet.* (1992). doi:10.1111/j.1469-1809.1992.tb01150.x • Giorda, R., Peakman, M., Tan, K. C., Vergani, D. & Trucco, M. Glutamic acid decarboxylase expression in islets and brain. *The Lancet* (1991). doi:10.1016/0140-6736(91)92781-V • Dirksen, R. et al. Targeting of the 67-kDa isoform of glutamic acid decarboxylase to intracellular organelles is mediated by its interaction with the NH2-terminal region of the 65-kDa isoform of glutamic acid decarboxylase. *J. Biol. Chem.* (1995). doi:10.1074/jbc.270.5.2241 • Bu, D. F. & Tobin, A. J. The exon-intron organization of the genes (gad1 and gad2) encoding two human glutamate decarboxylases (gad67 and gad65) suggests that they derive from a common ancestral gad. *Genomics* (1994). doi:10.1006/geno.1994.1246 • Demakova, E. V., Korobov, V. P. & Lemkina, L. M. Determination of gamma-aminobutyric acid concentration and activity of glutamate decarboxylase in blood serum of patients with multiple sclerosis. *Klin. Lab. Diagn.* (2003).
- Asada, H. et al. Mice lacking the 65 kDa isoform of glutamic acid decarboxylase (GAD65) maintain normal levels of GAD67 and GABA in their brains but are susceptible to seizures. *Biochem. Biophys. Res. Commun.* (1996). doi:10.1006/bbrc.1996.1898 • McHale, D. P. et al. A Gene for Autosomal Recessive Symmetrical Spastic Cerebral Palsy Maps to Chromosome 2q24-25. *Am. J. Hum. Genet.* (1999). doi:10.1086/302237

HTR2

- Anton, R. F. et al. Pharmacogenomics. *Nat. Genet.* (2008). doi:10.1016/j.ejca.2015.06.122 • Unschuld, P. G. et al. Polymorphisms in the serotonin receptor gene HTR2A are associated with quantitative traits in panic disorder. *Am. J. Med. Genet. Part B Neuropsychiatr. Genet.* (2007). doi:10.1002/ajmg.b.30412 • Kling, a et al. Genetic variations in the serotonin 5-HT2A receptor gene (HTR2A) are associated with rheumatoid arthritis. *Ann. Rheum. Dis.* (2008). doi:10.1136/ard.2007.074948 • HTR2A gene - Genetics Home Reference - NIH. U.S. National Library of Medicine (2020). Available at: <https://ghr.nlm.nih.gov/gene/HTR2A>.

MAO-A

- Kim, S. K. et al. Association study between monoamine oxidase A (MAOA) gene polymorphisms and schizophrenia: Lack of association with schizophrenia and possible association with affective disturbances of schizophrenia. *Mol. Biol. Rep.* (2014). doi:10.1007/s11033-014-3207-5 • Anton, R. F. et al. Pharmacogenomics. *Nat. Genet.* 16, 268–278 (2008).
- Karmakar, A. et al. Pilot study indicate role of preferentially transmitted monoamine oxidase gene variants in behavioral problems of male ADHD probands. *BMC Med. Genet.* (2017). doi:10.1186/s12881-017-0469-5 • Bortolato, M. & Shih, J. C. Behavioral outcomes of monoamine oxidase deficiency: Preclinical and clinical evidence. *In International Review of Neurobiology* (2011). doi:10.1016/B978-0-12-386467-3.00002-9

MAO-B

• Ukraintseva, S. V., Arbeev, K. G., Michalsky, A. I. & Yashin, A. I. Antiaging treatments have been legally prescribed for approximately thirty years. in *Annals of the New York Academy of Sciences* (2004). doi:10.1196/annals.1297.014 • Shih, J. C. & Chen, K. MAO-A and -B gene knock-out mice exhibit distinctly different behavior. *Neurobiology* (Budapest, Hungary). (1999). Available at: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC10591056/>. • Riederer, P. & Laux, G. MAO-Inhibitors in Parkinson's Disease. *Exp. Neuropiol.* (2011). doi:10.5607/en.2011.20.1.1 • Saura, J. et al. Increased monoamine oxidase b activity in plaque-associated astrocytes of Alzheimer brains revealed by quantitative enzyme radioautography. *Neuroscience* (1994). doi:10.1016/0306-4522(94)90311-5 • Mallajosyula, J. K., Chinta, S. J., Rajagopalan, S., Nicholls, D. G. & Andersen, J. K. Metabolic control analysis in a cellular model of elevated MAO-B: Relevance to parkinson's disease. *Neurotox. Res.* (2009). doi:10.1007/s12640-009-9032-2 • Nagatsu, T. & Sawada, M. Molecular mechanism of the relation of monoamine oxidase B and its inhibitor to Parkinson's disease: possible implications of glial cells. *J. Neural Transm. Suppl.* (2006). doi:10.1007/978-3-211-33328-0_7 • Kumar, M. J. & Andersen, J. K. Perspectives on MAO-B in Aging and Neurological Disease: Where Do We Go From Here? *Mol. Neurobiol.* (2004). doi:10.1385/MN:30:1:077 • Shih, J. C., Chen, K. & Riddle, M. J. MONOAMINE OXIDASE: From Genes to Behavior. *Annu. Rev. Neurosci.* (1999). doi:10.1146/annurev.neuro.22.1.197 • MAOB monoamine oxidase B [*Homo sapiens* (human)] - Gene - NCBI. National Center for Biotechnology Information Available at: <https://www.ncbi.nlm.nih.gov/gene/4129>. • Bortolato, M. & Shih, J. C. Behavioral outcomes of monoamine oxidase deficiency: Preclinical and clinical evidence. in *International Review of Neurobiology* (2011). doi:10.1016/B978-0-12-386467-3.00002-9 • Bortolato, M., Godar, S. C., Davarian, S., Chen, K. & Shih, J. C. Behavioral disinhibition and reduced anxiety-like behaviors in monoamine oxidase b-deficient mice. *Neuropsychopharmacology* (2009). doi:10.1038/npp.2009.118 • Miller, G. M. The emerging role of trace amine-associated receptor 1 in the functional regulation of monoamine transporters and dopaminergic activity. *Journal of Neurochemistry* (2011). doi:10.1111/j.1471-4159.2010.07109.x • Edmondson, D. E., Bindu, C. & Mattevi, A. Structural insights into the mechanism of amine oxidation by monoamine oxidases A and B. *Archives of Biochemistry and Biophysics* (2007). doi:10.1016/j.abb.2007.05.006 • Nolen, W. A., Hoencamp, E., Bouvy, P. F. & Haffmans, P. M. Reversible Monoamine Oxidase-A Inhibitors In Resistant Major Depression. *Clinical Neuropharmacology* 15, (1992).

SLC6A4

• Johnson, B. A. et al. Pharmacogenetic approach at the serotonin transporter gene as a method of reducing the severity of alcohol drinking. *Am. J. Psychiatry* 168, 265–275 (2011). • SLC6A4 gene - Genetics Home Reference - NIH. U.S. National Library of Medicine (2020). Available at: <https://ghr.nlm.nih.gov/gene/SLC6A4>. • Anton, R. F. et al. Pharmacogenomics. *Nat. Genet.* 16, 268–278 (2008). • Ait-Daoud, N. et al. Preliminary Evidence for cue-induced Alcohol Craving Modulated by Serotonin Transporter Gene Polymorphism rs1042173. *Front. Psychiatry* 3, 6 (2012). • Landgren, S. et al. Genetic Variation of the Ghrelin Signaling System in Females With Severe Alcohol Dependence. *Alcohol. Clin. Exp. Res.* 34, 1519–1524 (2010).

TPH2

• Bragatti, J. A., Bandeira, I. C., de Carvalho, A. M., Abujamra, A. L., Leistner-Segal, S., & Bianchin, M. M. (2014). Tryptophan hydroxylase 2 (TPH2) gene polymorphisms and psychiatric comorbidities in temporal lobe epilepsy. *Epilepsy and Behavior*. <https://doi.org/10.1016/j.yebeh.2014.01.007> • Zhang, X., Beaulieu, J. M., Gainetdinov, R. R. & Caron, M. G. Functional polymorphisms of the brain serotonin synthesizing enzyme tryptophan hydroxylase-2. *Cellular and Molecular Life Sciences* (2006). doi:10.1007/s00018-005-5417-4 • Plemenitaš, A., Kores Plesničar, B., Kastelic, M., Porcelli, S., Serretti, A., & Dolžan, V. (2015). Genetic variability in tryptophan hydroxylase 2 gene in alcohol dependence and alcohol-related psychopathological symptoms. *Neuroscience Letters*. <https://doi.org/10.1016/j.neulet.2015.07.037> • Natarajan, R., Einarsdóttir, E., Riutta, A., Hagman, S., Raunio, M., Mononen, N., ... Elovaara, I. (2012). Melatonin pathway genes are associated with progressive subtypes and disability status in multiple sclerosis among Finnish patients. *Journal of Neuroimmunology*. <https://doi.org/10.1016/j.jneuroim.2012.05.014> • Kim, Y. K., Lee, H. J., Yang, J. C., Hwang, J. A., & Yoon, H. K. (2009). A tryptophan hydroxylase 2 gene polymorphism is associated with panic disorder. *Behavior Genetics*. <https://doi.org/10.1007/s10519-008-9254-8> • Gao, J., Pan, Z., Jiao, Z., Li, F., Zhao, G., Wei, Q., ... Evangelou, E. (2012). TPH2 gene polymorphisms and major depression - a meta-analysis. *PLoS ONE*. <https://doi.org/10.1371/journal.pone.0036721> • Baehne, C. G., Ehliis, A. C., Plichta, M. M., Conzelmann, A., Pauli, P., Jacob, C., ... Fallgatter, A. J. (2009). Tph2 gene variants modulate response control processes in adult ADHD patients and healthy individuals. *Molecular Psychiatry*. <https://doi.org/10.1038/mp.2008.39> • Pae, C. U., Chiesa, A., Porcelli, S., Han, C., Patkar, A. A., Lee, S. J., ... De Ronchi, D. (2012). Influence of BDNF variants on diagnosis and response to treatment in patients with major depression, bipolar disorder and schizophrenia. *Neuropsychobiology*. <https://doi.org/10.1159/000327605> • Walitza, S., Renner, T. J., Demple, A., Konrad, K., Wewetzer, C., Halbach, A., ... Lesch, K. P. (2005). Transmission disequilibrium of polymorphic variants in the tryptophan hydroxylase-2 gene in attention-deficit/hyperactivity disorder. *Molecular Psychiatry*. <https://doi.org/10.1038/sj.mp.4001734> • Singh, A. S., Chandra, R., Guhathakurta, S., Sinha, S., Chatterjee, A., Ahmed, S., ... Rajanma, U. (2013). Genetic association and gene-gene interaction analyses suggest likely involvement of ITGB3 and TPH2 with autism spectrum disorder (ASD) in the Indian population. *Progress in Neuropsychopharmacology and Biological Psychiatry*. <https://doi.org/10.1016/j.pnpbp.2013.04.015> • Reuter, M., Kuepper, Y., & Hennig, J. (2007). Association between a polymorphism in the promoter region of the TPH2 gene and the personality trait of harm avoidance. *International Journal of Neuropsychopharmacology*. <https://doi.org/10.1017/S1461145706007073> • Su, Y. A., Li, J. T., Dai, W. J., Liao, X. M., Dong, L. C., Lu, T. L., ... Si, T. M. (2016). Genetic variation in the tryptophan hydroxylase 2 gene moderates depressive symptom trajectories and remission over 8 weeks of escitalopram treatment. *International Clinical Psychopharmacology*. <https://doi.org/10.1093/ICP.YIC.0000000000000115> • Serretti, A., Liappas, I., Mandelli, L., Albani, D., Forloni, G., Malitatis, P., ... Kalofoutis, A. (2009). TPH2 gene variants and anxiety during alcohol detoxification outcome. *Psychiatry Research*. <https://doi.org/10.1016/j.psychres.2007.12.006>