Epigenetics: Your DNA Is NOT Your Destiny!*

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<u>Epigenetics</u>: study of biological mechanisms that turn genes on and off and hence, affect gene EXPRESSION

• <u>Genotype</u> describes the inherited genes. <u>Phenotype</u> describes gene expression.

Resources: Genetics Home Reference: <u>http://ghr.nlm.nih.gov/gene</u>/

National Center for Biotechnology Information: <u>http://www.ncbi.nlm.nih.gov/gene/</u>

Epigenetics: the power to affect expression of our genes through our choices!

 Food intake, Environmental exposures (where you live, where you work, toxins, drug use, medications), Lifestyle (exercise, stress), Sleep, Social/Spiritual, Age

Genetic variations are called **<u>SNPs</u>** (Single Nucleotide Polymorphisms)

- Normal alleles from both parents = Wild-type genotype (Normal function/expression)
- A variation (SNP) in one allele = Heterozygous genotype
- A variation (SNP) in both alleles = Homozygous genotype (More severe variation)

Methylation Genes

• MTHFR (A1298C, C677T): activates folate to 5-MTHF

Functions of Methylation

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Controls genetic expression	Stress and relaxation response
Conversion of food to energy	Detoxification
Cellular protection	Immune response
Brain and muscle health	 Inflammation & gut integrity
Production and balance of	DNA repair and reduction of errors
neurotransmitters	

Health Risks with Methylation SNPs

Abnormal blood clotting	Behavioral health problems
Heart Disease	Cognitive problems
Stroke	Miscarriages
• High blood pressure (HTN, preeclampsia)	Cleft lip and palate
Eye problems, including glaucoma	Spina bifida

Supporting Poor Methylators

Higher intake of folate sources, such as	Avoidance of food additives,
leafy greens	preservatives and pesticides
Higher intake of B12 sources (i.e. eggs)	Avoidance of high-intensity exercise
• Supplements: B6, B12 and/or Folate	Adequate sleep
	Beneficial gut bacteria

Vitamin & Mineral Genes

- **<u>BCO1 (aka BCMO1)</u>**: converts dietary beta-carotene to active vitamin A
- **<u>GC</u>**: binding protein that transports Vitamin D to target tissues
- **<u>SLC23A1</u>**: encodes protein that transports vitamin C to organs

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• **<u>SLC30AB</u>**: encodes protein that transports zinc intracellularly

<u>Supporting Vitamin/Mineral SNPs</u> with specific vitamin/mineral that is affected (i.e. Vit D, Vit C, zinc)

Detoxification Genes

- Start w/ <u>CYP</u>: involved in the formation (synthesis) and breakdown (metabolism) of various molecules/chemicals
- Start w/ <u>GPX</u>: reduce hydrogen peroxide and hydroperoxides to protect cells against oxidative damage
- Start w/ GST: conjugate certain toxins with glutathione
- **<u>COMT</u>**: produces enzymes that breaks down brain neurotransmitters (dopamine, epinephrine, norepinephrine) and estrogen
 - SNPs assoc. with alcohol use disorder, fibromyalgia, opioid addiction, schizophrenia
- **<u>SOD</u>**: makes superoxide dismutase to break down toxic superoxide radicals
 - SOD1 SNP assoc. with amyotrophic lateral sclerosis (ALS)
 - SOD2 SNP assoc. with premature aging, motor neuron disease, cancer

Supporting Detoxification SNPs

Organic, whole for	oods that are less	•	Supplement cofactors for the enzymes
 processed Foods rich in ant Supplementation glutathione, SAM 	ioxidants n of vitamin C, NAC, 1e rbs that support detox	•	affected Limit exposure to pesticides, food additives, household chemicals, VOCs Sleep! Stress management

Weight Management Genes

- ADIPOQ: SNP associated with poor glucose and lipid metabolism
- **<u>FTO</u>**: SNP associated with increased appetite and decreased satiety
- MC4R: SNP assoc. with increased appetite
- **<u>PPARG</u>**: SNP assoc. with poor lipid management

Supporting Weight Management SNPs

• Best supported through diet high in unprocessed, whole plant foods, and high-quality protein and fats; Probiotics, fermented foods; Regular exercise/activity

Cardiovascular Health Genes

- APOA2: SNP may result in hypercholesterolemia
- Factor II & Factor V: rare SNP's increasing risk for bleeding
- Factor V Leiden (R506Q): SNP increased risk for venous thrombosis, hence blood clotting
- **NOS3**: nitric oxide gene involved in vascular smooth muscle relaxation
 - SNP assoc. with coronary spasm, hypertension, stroke, Alzheimer's disease
- AGT: angiotensinogen gene regulates blood pressure and balance of fluids and salt in the body
 - heterozygous SNP increases risk for hypertension
 - homozygous SNP causes abnormal kidney development at birth & hypotension

- <u>Apo E</u>
 - Apo E3/3: normal genotype in >50% of population
 - Apo E2/2: rare SNP assoc. with type III hyperlipidemia and 2-3 fold increased risk for CVD
 - Apo E4: most common SNP associated with increased risk for Alzheimer's disease and dementia, and hyperlipidemia with 1.5-fold increase for CVD
 - Heterozygous (E4/3) increased risk for late-onset Alzheimer's disease
 - Homozygous (E4/4) increased risk for earlier-onset Alzheimer's disease

Supporting Cardiovascular SNPs

•	Whole plant foods, that are less processed &	•	Regular exercise
	low sugar	•	Stress Management
•	High-quality protein that is low in saturated fats	•	Supplementation: B-vitamins, Vit. C, Vit. D, Vit.
•	Emphasis of omega-3 fats, unsaturated fats		K, Omega-3, Resveratrol
•	Probiotics, fermented foods	•	Herbs: Ashwagandha, Rhodiola

Glucose Homeostasis Genes

- **<u>SLC30A8</u>**: SNP reduces zinc transport to increase risk for type 2 diabetes
- **TCF7L2**: SNP implicated in blood glucose homeostasis to increase risk for type 2 diabetes

<u>Support Glucose SNPs</u> through diet high in unprocessed, whole plant foods, and high-quality protein and fats; also regular exercise; supp. zinc if applicable

Cognition & Neurological Genes

- Includes Apo E, COMT, MTHFR (discussed)
- MAOA: involved in breakdown of serotonin, epinephrine, norepinephrine, dopamine and tyramine
 - SNP causing low MAOA activity primarily in males, causing mild intellectual disability and behavioral problems, including aggressive and violent outburst → helps to reduce high tyramine foods
 - SNP causing high MAOA activity in females, causing panic disorders, severe anxiety and overwhelming fear
- **DRD2**: encodes subtype of dopamine receptor
 - SNPs associated with schizophrenia, opioid addiction and alcohol use disorder, due to dysregulation of dopamine
- **<u>TPH2</u>**: encodes enzyme in first step of serotonin production
 - SNPs assoc. with attention deficit-hyperactivity disorder (ADHD) and major depressive disorder

Supporting Cognition & Neurological SNPs (CAUTION: supplementation depends on individual needs!)

- Support methylation and detoxification SNPs
- Add neurotransmitter pre-cursors/amino acids: 5-HTP, I-Tyrosine, Taurine, L-theanine
- Add co-enzymes in neurotransmitter production: B6, Folate, Mg, Zinc, etc.
- Add herbs and food extracts
 - Bacopa, Vinpocetine, Ginkgo biloba, Rhodiola, etc.
 - Blueberry/Grape polyphenols, Curcumin, Green tea catechins, Quercetin flavonoid, etc.
- Add neurotransmitters: such as GABA

Immunity Genes

- IL Genes: SNPs affect various functions in inflammation to increase risks for the following
 - <u>IL-1B</u>: alopecia areata (AA) and gastric cancer
 - <u>IL-4</u>: AA and ischemic stroke
 - IL-6: arthritis, type 1 diabetes, IBS
 - **<u>IL-10</u>**: shingles, Crohn disease, rheumatoid arthritis
 - **<u>IL-13</u>**: psoriatic arthritis, asthma, allergic rhinitis
- <u>TNF</u>: tumor necrosis factor superfamily involved in cell proliferation, differentiation, apoptosis (cell death), lipid metabolism, coagulation
 - SNPs implicated in autoimmune diseases (arthritis), insulin resistance, cancer

Supporting Immunity SNPs

•	Gut support: over 70% of immune support!	•	Herbs/Extracts: Aloe vera, mushrooms,
•	Vitamins: Vit C, Vit D,		quercitin, citrus bioflavonoids, elderberry,
•	Minerals: Zinc		lemon balm, hesperidin
		•	Support immune function with healthy diet,
			sleep and stress management

Genes Specific to Medication Tolerance

- **<u>SLCO1B1</u>**: involved in removal of drug compounds such as statins
- **NAT2**: involved in activation and deactivation of arylamine and hydrazine drugs and carcinogens

Testing Options

- Genetic testing can get very expensive! Ranging \$60-\$700+
- Specific profiles: cardiovascular, detoxification, estrogen, immunity, neurological, pharmacogenic...
- A la carte options: MTHFR, EpoE, COMT, SOD2, IL, etc.
- Other: interpret raw data from 23andME, Ancestry.com (least expensive and gives variety of SNPs)

Can assist gene function no matter what!*

Organic, whole foods that are less processed	• Limit exposure to pesticides, food additives,
Foods rich in antioxidants	household chemicals, VOCs
Support a healthy microbiome	Sleep!
• Supplementation based on symptoms, blood	Stress management
tests or other evaluations	Choose exercise that invigorates vs.
	depletes/drains

*Gene testing will guide more specific support. Supplementation depends on individual needs/tolerance; caution with over-supplementation.