

# PATHWAYFIT® DIET, NUTRITION & EXERCISE PERSONAL GENETIC REPORT

# SAMPLE REPORT

**Protected Health Information** 



PAGE 2

Hello John,

Pathway Genomics is pleased to present you with your personalized metabolism, diet, nutrition and exercise report. This report is based on a DNA test and the lifestyle information that you recently submitted. It is well-documented that genetics accounts for 40 to 70% of a person's predisposition to obesity<sup>1</sup>. If you are thinking about starting a weight loss program or just maintaining a healthy diet, the goal of this test is to give you information about yourself that may help you modify your behavior.

Pathway Genomics' onsite laboratory is accredited by the College of American Pathologists (CAP), accredited in accordance with the U.S. Health and Human Services' Clinical Laboratory Improvement Amendments (CLIA), and licensed by the California Department of Public Health. We tested over 75 genetic markers to provide you with the latest, most comprehensive, and scientifically-advanced recommendations on diet, nutrition, exercise, addictive behaviors and weight-related health conditions. Pathway Fit® was developed with input from medical and scientific experts from leading institutions, such as UC Berkeley, Harvard Medical School, Scripps Clinic, Pennington Biomedical Research Center, Salk Institute for Biological Studies, and the University of Copenhagen. This report provides personalized information, based on your genetics and lifestyle, to help you meet the following goals:

- Understand your metabolism and behavior traits
- Reach and maintain a healthy weight
- Get the most benefit from physical activity and exercise
- Optimize the nutritional balance of your diet

A licensed physician has already reviewed your report; however, it is important that you discuss any modifications to your diet, exercise and nutritional supplementation with your physician before making any changes.

Pathway is here to help. If you have guestions or concerns regarding any aspect of this report, please contact our staff of genetic, medical or nutritional counselors by logging in to your Pathway Genomics account at www.pathway.com, or call us at (877) 505-7374. We are delighted we can help you on your path to optimum wellness.

Michael P. Nova, M.D. Chief Medical Officer

NOTE: This report has not been evaluated by the FDA. This product is not intended to diagnose, treat, cure, or prevent any disease.

SEX: MALE ACC #: C4215580

DATE:



IMPORTANT INFORMATION

PAGE 3



**Personal Details** 

Name: John DEPLYAYAYA

DOB: Jan 1, 1966 Gender: Male

Ethnicity: Caucasian

**Indication:** Population Screening

**Report Date:** 

Received Date: Jun 22, 2012

**Test Performed / Method** 

Genotyping by array-based evaluation of

multiple molecular probes

**Ordering Healthcare Professional** 

Carlos Doctor1Deploy M.D. 95646 Sorrento Valley Blvd Atlanta GA, 30375 US **License:** 123456

Lab Director: James R. Nickel, M.D.

James White Mo.

**Laboratory Info** 

**Accession #:** C4215580

Activation Code: DEPLY-AYAYA Specimen Source: Saliva Collected Date: Jun 20, 2012

Test Result Reviewed & Approved by Dir. Clin. Genetics: Linda Wasserman.

M.D., Ph.D.

Link Wessen







#### SCIENTIFIC STRENGTH RATING SYSTEM

The genetic markers and studies selected for this report represent the best and most recent genetic research in diet, nutrition, exercise and weight-related health conditions. Some research can be described as stronger than others based on the size of the population studied and whether the outcome has been replicated. Due to the current state of scientific research on the genetics of diet, exercise and nutrition, most of the studies referenced in your report are based on individuals of Caucasian ethnicity. While we all have the same genes, there are genetic and non-genetic factors in different ethnicities that might yield different outcomes for non-Caucasian populations. Your report includes a star system, described below, to rate the strength of the research evidence for the genetic marker and the associated result.

****	Results derived from a large study of approximately 2,000 or more people, with at least one additional study showing the same results (replication study).
***	Results derived from a moderately-sized study of at least 400 people, with or without a replication study.
****	Small study of less than 400 people in some cases, with other small replicated studies. Results in this category are preliminary, but pass our criteria for statistical significance.
****	Results in this category should be considered extremely preliminary.

#### Disclaimer

This report is intended as educational information, and is not intended to be used solely by the patient in medical decision-making without the consultation of a licensed health care professional. This test was developed and its performance characteristics determined by Pathway Genomics. It has not been cleared or approved by the U.S. Food and Drug Administration. If you have any questions about this report or wish to speak with one of Pathway Genomics' genetic counselors, please call (877) 505-7374.



PATHWAY SENOMICS

PAGE 4



#### DIET





Eat a balanced diet of protein, fat and carbohydrates, rather than a diet that is targeted towards being specifically low in fat or carbohydrates.



You may indulge more than average on tempting foods, as you have a genetic marker associated with eating disinhibition. Reduce your exposure to foods that tempt you.



As someone who has enhanced bitter taste perception, you may not like the taste of certain healthy vegetables, such as broccoli or leafy greens. Try recipes that mask the bitter flavors without adding too many calories.



You are less likely to be lactose intolerant, which means you may consume dairy products and not have gastrointestinal side effects. Choose dairy products that are lower in calories, fat and added sugar.



As a slow caffeine metabolizer, you may want to limit your caffeine intake to one or two cups of coffee per day, or an equivalent amount of caffeine. This may limit the likelihood of some of the common side effects.



#### **NUTRITIONAL-NEEDS**

#### 9 Genetic Markers Tested



You have a genetic variant associated with lower vitamin B-6 levels. Be sure your diet includes foods rich in vitamin B-6, such as dark green leafy vegetables, whole grains, legumes, poultry, fish and eggs.



Review the recommended daily allowance (RDA) for vitamin and nutrient levels for your age, gender and health status. Do not exceed recommended levels. Sometimes too much can be harmful.



Talk to your physician, a registered dietitian, or a certified nutritionist about monitoring your nutrient levels and assessing your nutritional needs.



PATHWAY GENOMICS

PAGE 5



#### EXERCISE





Your genetics are associated with enhanced health benefits from endurance exercises, such as mid-long distance walking, jogging and bicycling. Weight resistance exercises may be less beneficial.



You have a genetic variant associated with being overweight. You can lower your chances by leading a physically active lifestyle.



You have a genetic variant associated with elevated blood pressure if you are not physically active. Exercise may help you manage your blood pressure.



Continue a vigorous exercise regimen after losing weight. You have genes that are associated with an increased chance of gaining weight back.



#### **METABOLIC-HEALTH**

#### 53 Genetic Markers Tested



You have a higher than average genetic risk for obesity. If you are having difficulties controlling your weight, you should take action as soon as possible to reduce your risks. Regardless of your current weight, your Pathway Fit report provides you with guidance that can help you work actively to achieve and maintain a healthy weight.



To ensure that your diet and exercise program are helping you maintain optimal health, work with your physician to track your blood sugar and cholesterol levels.



You have an "Above Average" genetic likelihood for some metabolic health factors, but you do not have a "High" likelihood in any category. Discuss these results with your physician.



**SUMMARY** 



PAGE 6

YOUR MATCHING DIET	T1
Matching Diet Type p. 8	BALANCED DIET
Response To Monounsaturated Fats p. 11	NEUTRAL

Response To Polyunsaturated	
Fats p. 11	INCREASED BENEFIT

Omega-6 And Omega-3 Levels	
p. 12	TYPICAL

		_
FATING	<b>BEHAVIOR</b>	TRAITS
LAIIIVO	DLIIAVION	IIIAII

ATING BEHAVIOR TRAITS	
	T) (D)

Snacking p. 14	TYPICAL
Hunger p. 14	TYPICAL
Satiety - Feeling Full p. 14	TYPICAL
Eating Disinhibition p. 15	MORE LIKELY
Food Desire p. 15	TYPICAL
Sweet Tooth p. 16	TYPICAL

#### **FOOD REACTIONS**



Caffeine Metabolism p. 18	SLOW METABOLIZER
Bitter Taste p. 18	TASTER
Sweet Taste p. 19	TYPICAL
Lactose Intolerance p. 19	LESS LIKELY
Alcohol Flush p. 20	LESS LIKELY

#### **NUTRITIONAL NEEDS**



Vitamin B2 p. 22	STAY BALANCED
Vitamin B6 p. 23	OPTIMIZE INTAKE
Vitamin B12 p. 23	STAY BALANCED
Folate - Folic Acid p. 24	STAY BALANCED
Vitamin A p. 25	INCONCLUSIVE
Vitamin C p. 26	STAY BALANCED
Vitamin D p. 26	STAY BALANCED
Vitamin E p. 27	OPTIMIZE INTAKE

#### **EXERCISE**



Endurance Training p. 30	ENHANCED BENEFIT
Strength Training p. 31	LESS BENEFICIAL
Aerobic Capacity (VO2max) p. 31	DECREASED
Muscle Power p. 32	LESS MUSCLE POWER
Achilles Tendinopathy p. 32	TYPICAL
Weight Loss Response To Exercise p. 33	EXERCISE STRONGLY RECOMMENDED
Blood Pressure Response To Exercise p. 33	EXERCISE STRONGLY RECOMMENDED
HDL (Good) Cholesterol Response To Exercise p. 34	NORMAL BENEFIT
Loss Of Body Fat Response To Exercise p. 34	NORMAL BENEFIT
Insulin Sensitivity Response To Exercise p. 34	ENHANCED BENEFIT

#### YOUR BODY AND WEIGHT



Obesity p. 37	ABOVE AVERAGE
Weight Loss-regain p. 37	MORE LIKELY TO GAIN WEIGHT BACK
Metabolism p. 38	NORMAL
Adiponectin Levels p. 38	TYPICAL

#### **METABOLIC HEALTH FACTORS**



Elevated LDL Cholesterol p. 40	AVERAGE
Decreased HDL Cholesterol p. 41	AVERAGE
Elevated Triglycerides p. 42	AVERAGE
Elevated Blood Sugar p. 43	AVERAGE

SEX: MALE ACC #: C4215580 DATE:



DIET

PAGE 7



# PERSONALIZE YOUR DIET WITH GENETICS

The way we eat, how our bodies process foods, and our overall health are impacted by our genetics. Scientific studies have shown that genetics can also be important for diet effectiveness. Your results have been calculated to determine the best diet likely to help you optimize your metabolism, lose weight and improve your health.

#### ▶ YOUR RESULTS ◀

MATCHING DIET TYPE

page:8

**BALANCED DIET** 

RESPONSE TO MONOUNSATURATED FATS

page:11

**NEUTRAL** 

W1

RESPONSE TO POLYUNSATURATED FATS

page:11

**INCREASED BENEFIT** 



OMEGA-6 AND OMEGA-3 LEVELS

page:12

**TYPICAL** 





DIET



PAGE 8



### MATCHING DIET TYPE

Your diet has been selected by looking at many genetic variants associated with how people respond to the different macronutrients (proteins, fats and carbohydrates) in their food<sup>2,3,4,5,6,7</sup>. Your genetic risk profiles discussed in the Metabolic Health Factors section of this report were also evaluated to determine your recommended diet<sup>8,9</sup>. Together, your genetic results suggest which one of the following diets may be best for you: "Low Fat," "Low Carb," "Mediterranean" or a "Balanced Diet." It is highly recommended to discuss any change in your diet plan with your health care provider.

#### YOUR DIET RECOMMENDATIONS

- ✓ Eat a balanced diet of protein, fat and carbohydrates, rather than a diet that is targeted towards being specifically low in fat or carbohydrates.
- ✓ You may indulge more than average on tempting foods, as you have a genetic marker associated with eating disinhibition. Reduce your exposure to foods that tempt you.
- ✓ As someone who has enhanced bitter taste perception, you may not like the taste of certain healthy vegetables, such as broccoli or leafy greens. Try recipes that mask the bitter flavors without adding too many calories.
- ✓ You are less likely to be lactose intolerant, which means you may consume dairy products and not have gastrointestinal side effects. Choose dairy products that are lower in calories, fat and added sugar.
- ✓ As a slow caffeine metabolizer, you may want to limit your caffeine intake to one or two cups of coffee per day, or an equivalent amount of caffeine. This may limit the likelihood of some of the common side effects.

#### YOUR RESULT

#### **BALANCED DIET**

Your genotype is associated with weight loss or other health benefits from a diet balanced in healthy fats, low glycemic index carbohydrates, protein and overall low calories.

YOUR RELATED GENES			
Gene Tested Your Scientific Genotype Strength			
FTO-rs9939609	T/A	****	
MMAB-rs2241201	G/C	****	
PPARG-rs1801282	C/C	****	
LIPC-rs1800588	C/T	****	
ADIPOQ- rs17300539	G/G	****	
KCTD10- rs10850219	G/G	****	
APOA2-rs5082	T/T	****	
AND MORE			

#### LOW CARB

vegetables, highquality proteins and healthy fats

#### LOW FAT

rich vegetables, grains and fruits, and healthy fats



Balance of healthy fats, carbohydrates and proteins

#### **MEDITERRANEAN**

Fish, monounsaturated fats, and low glycemic, high fiber vegetables, fruits, grains and legumes

SEX: MALE ACC #: C4215580

DATE:







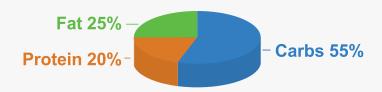
PAGE 9

PATHWAY GENOMICS

DIET

#### **BALANCED DIET**

To achieve a diet that is balanced, you should eat an array of food from each of the food groups. Focus on the most nutrient-dense foods, including strongly colored fruits and vegetables with bold flavors, whole grains, such as oatmeal, whole wheat bread, quinoa, brown rice and corn tortillas. For your protein intake, incorporate legumes, fish (and other seafood), and limit your red meat consumption to about 3 ounces or less, 2 to 3 times per week. Regarding fats and oils, it's best to choose vegetable fats and to minimize your intake of animal fats. Processed and highly refined foods, trans fats, as well as added sugars, should be avoided.



Balanced Diet: Key Aspects

#### Fruits and Vegetables

- > Bright colors, bold flavor.
- > Consume a variety of colors.
- Try to eat 9 servings of fruits and vegetables per day.
- ➤ Leafy green veggies are optimal.
- Limit store-bought juice to 1/2 cup per day (no sugar or sweetener).

#### Grains and Starchy Vegetables

> Avoid processed and refined grains.

#### Protein Foods

- ➤ At least 1 to 2 servings of protein should come from legumes.
- Limit red meats.
- ➤ Eat fish or other seafood at least 2 to 3 times per week.
- > Remove all visible fat and skin from meat, fish and poultry.
- Prepare meat by baking, broiling, steaming or poaching.
- > Avoid frying meat.

#### Milk Products

- > Plain Greek-style yogurt is optimal.
- > Avoid milk products with added sugar.
- > Limit cheese.

#### Fats and Oils

- > Avoid hydrogenated and trans fats.
- ➤ Limit saturated fats.

#### General

Minimize or avoid added sugars and foods with added sugar. This is especially important if you are trying to lose weight or control your blood sugar levels, or if your triglyceride levels are elevated.

SEX: MALE ACC #: C4215580

DATE:



DIET

PAGE 10

DIFT

#### TYPES OF FAT IN YOUR DIET

Acting as an important part of any diet and a source of energy, fat provides flavor to your diet, but more importantly, it is a vital element in the absorption of fat-soluble vitamins such as vitamins A, D, E and K. The two major types of fat include saturated and unsaturated (polyunsaturated and monounsaturated) fats. In order for your body to function normally, you need to maintain a consistent and balanced supply of saturated and unsaturated fats. A third type of fats consists of hydrogenated fats, which are processed fats that are not found naturally, such as in margarine and fried fast foods. Hydrogenated fats may also contain trans fatty acids and are generally unhealthy and should be avoided.

#### SATURATED FAT ➤ Veal

- ➤ Beef
- ➤ Lamb
- ➤ Lard

meat)

- ➤ Milk
- ➤ Cream
- ➤ Poultry (dark
- ➤ Pork
- ➤ Butter
- ➤ Cheeses
- ➤ Coconut oil
- salmon,
- herring, halibut.
  - ➤ Walnuts

  - > Flaxseed
  - ➤ Chia seed
  - > Pumpkin Seed
  - primrose oil

  - seeds
  - eggs

#### **UNSATURATED FAT**

Monounsaturated

➤ Avocados

➤ Extra virgin

olive oil

➤ Nuts

➤ Olives

#### Polyunsaturated

- ➤ Cold water fish (e.g., sardines, mackerel)
- ➤ Almonds

- ➤ Evening
- ➤ Borage seed
- ➤ Nuts and
- ➤ Poultry and

#### HYDROGENATED FAT

- ➤ Margarine (stick)
- ➤ Most fast foods
- ➤ Fried foods
- ➤ Highly processed foods
- ➤ Shortening
- ➤ Foods
- containing trans fats



DIET



PAGE 11



# DIET RESPONSE TO MONOUNSATURATED FATS

Fat is an important part of any diet, and not all fats are bad. Monounsaturated fat is considered a healthy dietary fat found in avocados, olives, and some nuts, as well as oils, such as olive oil. The two possible outcomes for this test are "Increased Benefit" or "Neutral." Having an "Increased Benefit" from monounsaturated fat suggests you could benefit from eating foods containing monounsaturated fats. In general, it is best to avoid trans fats and limit saturated fat intake.

Genetic variants in two genes, ADIPOQ and PPARG, have been associated with a lower body weight in individuals when more than 13% of their calories come from monounsaturated fats<sup>6,7</sup>. This would be equivalent to a person on an 1,800-calorie diet consuming about 1 to 2 tablespoons of olive oil and a quarter cup of nuts each day as part of their total caloric intake. While the ADIPOQ study was done in a population of both men and women, the PPARG study was done only in women. There is not enough scientific evidence to support if the PPARG association is also true in men.



#### **NEUTRAL**

For people with your genotype, the amount of dietary monounsaturated fat you eat is not likely to affect your body weight. However, avoiding trans fats and substituting some saturated fats with monounsaturated fats is still recommended, as it has several health benefits.

YOUR RELATED GENES			
Gene Tested Your Scientific Genotype Strength			
PPARG-rs1801282	C/C	****	
ADIPOQ- rs17300539	G/G	****	



## DIET RESPONSE TO POLYUNSATURATED FATS

Polyunsaturated fat is considered a healthy fat and is important for heart and brain function, as well as growth and development. Two types of polyunsaturated fats are omega-3 and omega-6 fats. Good sources of omega-6 fats include evening primrose and borage oils, as well as olives, nuts and poultry. Additionally, good sources of omega-3 fats include fish and seafood, as well as flaxseed, walnuts, hemp seeds, and dark green leafy vegetables.

The two possible outcomes in this report are "Increased Benefit" or "Neutral." Having an "Increased Benefit" from polyunsaturated fat means you should try to eat foods containing polyunsaturated fats. In general, it is best to avoid trans fats and minimize saturated fats. One study in women has shown that those with a certain genetic variant in the PPARG gene tend to have a lower body weight when they consume more polyunsaturated fats than saturated fats<sup>7</sup>. This association has not been studied in men.



#### **INCREASED BENEFIT**

People with your genotype who have a diet that includes more polyunsaturated fats, rather than saturated fats, tend to have a lower body weight, compared to those who do not.

ntific ngth				
PPARG-rs1801282				
1				

SEX: MALE ACC #: C4215580 DATE:



DIET

PAGE 12

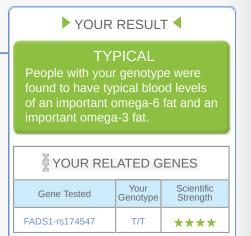


**"1** 

# DIET OMEGA-6 AND OMEGA-3 LEVELS

Polyunsaturated fats (PUFAs) in our diet are composed of omega-3 and omega-6 fatty acids, both of which are recommended by the American Heart Association (AHA) for good heart health. Long-chain PUFAs are provided by our diet, but can also be synthesized in our bodies starting from the precursor essential fatty acids, linoleic acid (LA, omega-6) and alpha-linolenic acid (ALA, omega-3). Both omega-3 and omega-6 fats are processed in the body by the same enzyme complex<sup>10</sup>. The major dietary sources of omega-3 fatty acids include foods, such as flaxseed and walnuts, as well as fish oils and fish such as salmon. Processed foods often contain high levels of omega-6, while healthy sources of omega-6 include evening primrose and borage oils, as well as olives, nuts and poultry. Historically, the ratio of omega-6 to omega-3 fats in the diet was maintained close to a healthy 1:1, while in the current Western diet it is estimated to be about 15:1<sup>11</sup>.

In recent genome-wide association studies that included over 10,000 people, it was found that those with the C/C or C/T genotypes at a variant in the FADS1 gene, which codes for one of the enzymes involved in processing omega-3 and omega-6 fats, had "Decreased" blood levels of arachidonic acid (AA), a long-chain omega-6 fat, as well as eicosapentaenoic acid (EPA), a long-chain omega-3 fat. On the other hand, those with a T/T genotype had "Typical" levels of these two omega-fats 12,13. Since both AA and EPA are precursors of biologically important metabolites, those with a "Decreased" outcome should increase their dietary intake of both omega-3 and omega-6 fatty acids. However, considering the current skewed ratio of omega-6:omega-3 fats, it is recommended that people monitor the intake of omega-6 fats from processed foods, while increasing their intake of omega-3 fats.



SEX: MALE ACC #: C4215580 DATE:



EATING BEHAVIOR TRAITS

PAGE 13



# EATING BEHAVIORS CAN BE INFLUENCED BY GENES. There are certain genes that have the potential to impact how we perceive and desire particular foods, and influence our eating behaviors, such as excessive snacking and difficulty feeling full. For example, variants in the ANKK1 and DRD2 genes, which result in a reduced density of dopamine receptors in your brain, have been associated with eating and addictive behaviors 14,15,16.

This type of information, generated from testing genetic markers in a number of genes, is included in your report, and can be used to understand how to modify your lifestyle and behaviors for optimum wellness.

#### **▶ YOUR RESULTS** ◀

SNACKING	page:14	TYPICAL

HUNGER	page:14	TYPICAL

<b>(3)</b>	SATIETY - FEELING FULL	page:14	TYPICAL

<b>3</b>	EATING DISINHIBITION	page:15	MORE LIKELY

FOOD DESIRE	page:15	TYPICAL

(E)	SWEET TOOTH	page:16	TYPICAL

SEX: MALE ACC #: C4215580

DATE:



EATING BEHAVIOR TRAITS

PAGE 14

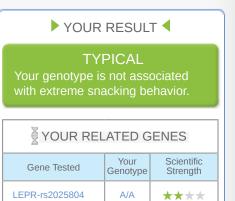




# EATING BEHAVIOR TRAITS SNACKING

Snacking can be a healthy or unhealthy behavior. Snacking on balanced foods, containing healthy fats, lean protein, fiber and low glycemic index carbohydrates, in small portions, throughout the day can help control hunger cravings and reduce total caloric intake, while snacking on junk food can have negative health effects. Genetic markers associated with snacking behavior include variants in the receptor for leptin, an essential hormone for the regulation of food intake. The possible results in this report are "Typical" and "Increased." If you receive the "Increased" result, you may want to curtail the negative effects of snacking by choosing healthy snacks, eating slowly and reducing the size or calories of snacks. People with the G/G genotype in a leptin receptor (LEPR) genetic marker were more likely to show

"Increased" snacking behavior <sup>17</sup>. "Typical" genotypes were not associated with "Increased" snacking behavior in the same study. This association has not been studied in men.





# EATING BEHAVIOR TRAITS HUNGER



While most of us know the feeling of hunger, some people feel hunger more intensely and more often than others. Susceptibility to hunger can now be partially explained by genetics. A variation in the NMB gene has been associated with increased feelings of hunger 18. When asked about their own eating behaviors on a questionnaire, people with a T/T genotype were more likely to report an "Increased" susceptibility to hunger, while others were likely to have a "Typical" hunger response. This preliminary information is based on a study rated with one star of scientific strength.



#### **TYPICAL**

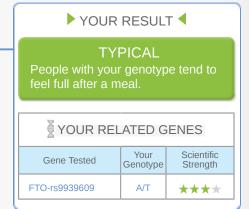
People with your genotype are more likely to exhibit typical levels of susceptibility to hunger.

YOUR RELATED GENES			
Gene Tested Your Scientific Genotype Strength			
NMB-rs1051168	G/T	****	



# EATING BEHAVIOR TRAITS SATIETY - FEELING FULL

Satiety can be described as the feeling of fullness after you eat. The FTO (fat mass and obesity-associated) gene is known to be an important factor that predisposes a person to a healthy or unhealthy level of body weight<sup>19</sup>. The two possible outcomes in this report are "Difficulty in Feeling Full" and "Typical." People who experience "Difficulty in Feeling Full" tend to eat more without feeling satisfied. To help manage this outcome, you could increase the amount of fiber in your diet and balance meals and snacks throughout the day. Examples of foods high in fiber include whole wheat bread, oatmeal, barley, lentils, black beans, artichokes, raspberries, and peas. In a 2008 study, the A/A genotype at rs9939609 in the FTO gene was associated with "Difficulty in Feeling Full"<sup>20</sup>. Although this study was done in children, there is preliminary data to support that the association also holds true in adults<sup>21</sup>.



SEX: MALE ACC #: C4215580 DATE:



EATING BEHAVIOR TRAITS

PAGE 15

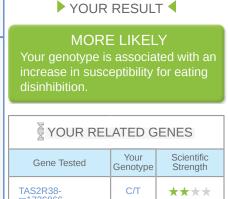




# EATING BEHAVIOR TRAITS EATING DISINHIBITION

Eating disinhibition describes the tendency to eat more than normal in response to a stimulus, such as a tasty food or in situations that trigger overeating (e.g., emotional stress or specific social situations). In a 2010 study, the T allele of rs1726866 was "More Likely" to be associated with eating disinhibition in women<sup>22</sup>. The C/C genotype at the same marker was "Less Likely" to be associated with eating disinhibition. There is not enough scientific evidence yet to;determine if this association also holds for men.







## EATING BEHAVIOR TRAITS FOOD DESIRE

Although there is no objective method to quantify someone's feeling of hunger or liking for a particular type of food, behavioral scientists have devised techniques to measure an individual's motivation to consume food and compare it with that of others. This measurement, called the reinforcing value of food<sup>23</sup>, describes how much effort an individual is willing to put forth to get access to food. The reinforcing value can be determined through a series of tests in a laboratory setting. In each of those tests, the individual being tested is asked to complete a task in exchange for a small portion of his or her favorite foods. The task of the initial test is easy, so the food is not difficult to win. As the tests continue, the task gets more and more difficult until, at some point, the participant feels that the food is no longer worth the effort and decides to quit. This experiment tells us that early quitters, when compared with late quitters, are low in food reinforcement. Using this technique, a 2007 study<sup>14</sup> identified a genetic component in food reinforcement. Among people who were considered obese, those who had a specific variant (T allele) of the genetic marker rs1800497 had an "Increased" likelihood to make more effort to obtain their favorite foods and eat more of them. In contrast, the C/C genotype was associated with "Typical" levels of food reinforcement.



#### TYPICAL

Your genotype is not associated with an increased desire or willingness to put forth additional effort to obtain your favorite foods.

§ YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
ANKK1/DRD2- rs1800497	C/C	****

SEX: MALE ACC #: C4215580 DATE:



EATING BEHAVIOR TRAITS

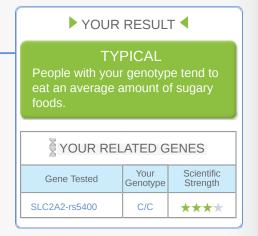
PAGE 16





# EATING BEHAVIOR TRAITS SWEET TOOTH

Craving sweet foods is sometimes described as having a "sweet tooth." The possible outcomes in this report are "Increased" or "Typical." If your genotype shows an "Increased" likelihood to eat lots of sweets, try choosing fruit as a healthy sweet alternative to sugary foods or soda. Be sure to follow your diet as some diet plans, such as the low carbohydrate diets, significantly limit the amount of sugar you can eat. Sweet foods can include healthy foods, such as fruits, or unhealthy foods like candy and sweetened beverages. People with the C/T and T/T genotypes showed an "Increased" likelihood to eat more sweets and sugary foods, while people with the C/C genotype were more likely to have a "Typical" intake of sugary foods<sup>24</sup>.



SEX: MALE ACC #: C4215580 DATE:



**FOOD REACTIONS** 

PATHWAY GENOMICS

PAGE 1

# YOUR GENETICS MAY IMPACT HOW YOU RESPOND TO SOME FOODS.

Genetic studies have been reported on some types of food reactions. Our tests draw on the current genetic data for responses to caffeine, bitter foods, sweet foods, milk products and alcohol. A summary of your results is provided below.

#### ▶ YOUR RESULTS ◀

CAFFEINE METABOLISM

page:18

**SLOW METABOLIZER** 

**BITTER TASTE** 

page:18

**TASTER** 

SWEET TASTE

page:19

**TYPICAL** 

LACTOSE INTOLERANCE

page:19

**LESS LIKELY** 

ALCOHOL FLUSH

page:2

**LESS LIKELY** 





FOOD REACTIONS



PAGE 18



#### FOOD REACTIONS CAFFEINE METABOLISM



Caffeine is one of the most widely consumed stimulants in the world, and it is found in the leaves and seeds of many plants. It is also produced artificially and added to some foods. Caffeine is found in tea, coffee, chocolate, many soft drinks and energy drinks, as well as in some pain relievers and other over-the-counter medications. Caffeine is metabolized by a liver enzyme, which is encoded by the CYP1A2 gene. Variation at a marker in the CYP1A2 gene results in different levels of enzyme activity, and thus, different metabolism rates for caffeine<sup>25,26,27</sup>. Therefore, the two possible genetic results in this report are "Fast Metabolizer" and "Slow Metabolizer." If you are a "Slow Metabolizer," then caffeine may have longer lasting stimulant effects for you. In addition to genetics, your body's ability to metabolize caffeine also depends on other lifestyle factors<sup>28,29</sup>. For example, how much coffee you drink, whether you smoke or whether you take hormonal birth control, may also affect your ability to metabolize caffeine. Because these and other lifestyle factors may both increase or decrease your caffeine metabolism, the most sensible advice is to make lifestyle choices that have the maximum benefit for your overall health.

#### YOUR RESULT

#### **SLOW METABOLIZER**

You are likely to slowly metabolize caffeine. It is suggested that you keep your total caffeine intake to less than 200mg per day, which is about one to two cups of coffee per

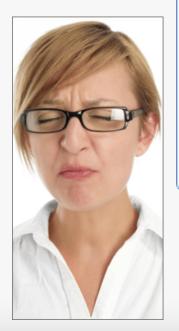
YOUR RELATED GENES			
Gene Tested	Your Genotype	Scientific Strength	
CYP1A2-rs762551	A/C	++++	

\*\*\*



#### **FOOD REACTIONS BITTER TASTE**

People taste things differently. Variations in the TAS2R38 gene are associated with different levels of sensitivity to a chemical called PTC 30,31. which produces a strong bitter taste. The possible results for bitter taste are "Taster," "Non-Taster," or "Inconclusive." A person described as a "Taster" may be more sensitive to bitter flavors found in foods, such as grapefruit, coffee, dark chocolate and cruciferous vegetables, such as Brussels sprouts, cabbage and kale. Being a "Taster" does not mean you do not enjoy these foods, but you may sense a stronger bitter taste compared to a "Non-Taster." In addition, tasters may need to watch their salt intake. because they may have an increased preference for salty foods, which mask the bitterness<sup>32</sup>. A genetic result of "Inconclusive" means that there is not enough scientific evidence for how your genotype is associated with bitter taste sensitivity.



#### YOUR RESULT **TASTER** You are likely to have a high sensitivity to bitter taste. YOUR RELATED GENES Your Genotype Scientific Strength Gene Tested TAS2R38-rs713598 \*\*\* TAS2R38-C/T \*\*\*

rs1726866



**FOOD REACTIONS** 

PATHWAY

PAGE 19



# FOOD REACTIONS SWEET TASTE

Sweet is one of the most basic tastes we can experience, and is usually found in sugar and sugary foods. The sensation of sweet taste is triggered to the brain from the taste buds. There are receptors on your tongue that are programmed by your genes to determine how you taste sweetness. A 2009 study showed that genetic variants found in the sweet taste receptors can result in "Typical" or "Decreased" sensitivity to the sweet taste of sugar<sup>33</sup>. People with "Decreased" sensitivity may prefer foods with more sugar since they are less likely to taste sweetness in foods that are low sugar.



#### YOUR RESULT

#### **TYPICAL**

You are likely to have typical sensitivity to the sweet taste of sugar.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
TAS1R3- rs35744813	G/G	****



# FOOD REACTIONS LACTOSE INTOLERANCE



Lactose intolerance is the inability to digest lactose, the sugar found in milk and milk products. This condition is caused by the lack of an enzyme called lactase. The rs4988235 variant lies close to the lactase (LCT) gene, in the MCM6 gene, and has been shown to regulate lactase levels 34,35,36. If you are lactose intolerant you should make sure that you are getting enough calcium from non-dairy or lactose-free sources. On the other hand, if you are not lactose intolerant, be aware that dairy products can be high in calories, fat, or both. You need to watch your intake accordingly or select low fat dairy products. People with a C/C genotype at rs4988235 are "More Likely" to be lactose intolerant, while people with other genotypes are "Less Likely"37. This variant has been found to be associated with lactose intolerance in Caucasians, while other variants might play an important role in other ethnicities, including Africans and Asians.

#### YOUR RESULT

#### LESS LIKELY

People with your genotype are less likely to be lactose intolerant.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
MCM6-rs4988235	T/T	****

SEX: MALE ACC #: C4215580 DATE:



**FOOD REACTIONS** 

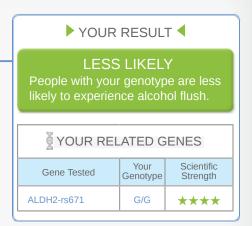


PAGE 20



# FOOD REACTIONS ALCOHOL FLUSH

Drinking alcoholic beverages is a relaxing or social activity for many, but for some it is exceedingly unpleasant due to their body's adverse reaction to alcohol. One such reaction is called alcohol flush, in which drinking even small amounts of alcohol causes a person's face to flush red and in some cases feel warm and itchy. People who flush may also experience other unpleasant symptoms, such as rapid heartbeat, nausea, or dizziness in response to alcohol. Alcohol flush is largely attributed to genetic variation in the ALDH2 gene, which encodes an enzyme critical for proper alcohol metabolism. Those who carry the inactive version of this gene are much "More Likely" to flush and experience other negative responses to alcohol 38,39, while people with other genotypes are "Less Likely" to flush. Perhaps not surprisingly, this variant is also associated with overall reduced consumption of alcohol 40. In most cases, avoiding alcohol is the best remedy for those who experience alcohol flush.





PATHWAY GENOMICS

PAGE 21

# VITAMINS AND NUTRIENTS ARE AN IMPORTANT PART OF OUR HEALTH.

There are genetic markers associated with being predisposed to lower levels of certain nutrients, which means you may want to make certain your diet has enough of the foods that contain these nutrients. Ensuring you consume the right amount of vitamins and nutrients from your diet is an important part of your health plan. The recommended daily allowances (RDA) for vitamins and nutrients in this section are based on guidance from the Institute of Medicine of the National Academies (IOM). For more information regarding RDAs, visit www.iom.edu.



#### ► YOUR RESULTS ◀

VITAMIN B2	page:22	STAY BALANCED
VITAWIIN BZ	paye.22	STAT BALANCEL

VITAMIN B6	page:23	OPTIMIZE INTAKE

VITAMIN B12	page:23	STAY BALANCED
V117 (IVIII V DIL	13	

l .		
FOLATE - FOLIC ACID	page:24	STAY BALANCED

VITAMIN A	page:25	INCONCLUSIVE

VITAMIN C	page:26	STAY BALANCED

VITAMINI D	page:26	STAY BALANCED
VIII/ (IVIII V D	13	

VITAMIN E	page:27	OPTIMIZE INTAKE



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

PAGE 22

#### YOUR NUTRITION RECOMMENDATIONS

- ✓ You have a genetic variant associated with lower vitamin B-6 levels. Be sure your diet includes foods rich in vitamin B-6, such as dark green leafy vegetables, whole grains, legumes, poultry, fish and eggs.
- ✓ Review the recommended daily allowance (RDA) for vitamin and nutrient levels for your age, gender and health status. Do not exceed recommended levels. Sometimes too much can be harmful.
- ✓ Talk to your physician, a registered dietitian, or a certified nutritionist about monitoring your nutrient levels and assessing your nutritional needs.





#### NUTRITIONAL NEEDS VITAMIN B2

Vitamin B2, or riboflavin, is a central component of flavin mononucleotide (FMN) and flavin adenine dinucleotide (FAD), both of which serve as cofactors of several critical enzymes involved in the electron transport chain, as well as in the metabolism of carbohydrates, fats and proteins<sup>41</sup>. Vitamin B2 is found in a variety of foods including milk, cheese, green leafy vegetables, legumes, beans, lean meats and fortified grains. Individuals with the T/T genotype at a variant in the MTHFR gene are likely to have increased levels of homocysteine, which are a risk factor for cardiovascular disease and stroke<sup>42,43,44</sup>. Levels of homocysteine were highest in T/T individuals with low riboflavin or vitamin B2 levels, and further, riboflavin supplementation was found to reduce homocysteine levels in these individuals<sup>45,46</sup>. Thus, individuals with the T/T genotype should "Optimize Intake" of vitamin B2 by eating foods rich in vitamin B2. On the other hand, vitamin B2 levels are likely to have a relatively small impact on homocysteine levels in people with the C/T or C/C genotypes, and hence, they should "Stay Balanced" and maintain a healthy diet.



#### STAY BALANCED

In people with your genotype, riboflavin levels have a relatively small impact on levels of homocysteine. Elevated levels of homocysteine are a risk factor for heart disease. You should maintain a healthy diet.

# YOUR RELATED GENES Gene Tested Your Genotype Scientific Strength MTHFR-rs1801133 C/C ★★★★

SEX: MALE ACC #: C4215580 DATE:



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

PAGE 23



#### NUTRITIONAL NEEDS VITAMIN B6

Vitamin B-6, also called pyridoxine, helps your body's neurological system to function properly, promotes red blood cell health, and is involved in sugar metabolism ("http://ods.od.nih.gov/factsheets/vitaminb6/"). Vitamin B-6 is found naturally in many foods, including beans, whole grains, meat, eggs and fish. Most people receive sufficient amounts of vitamin B-6 from a healthy diet, and B-6 deficiency is rare in the United States.

The genetic marker rs4654748 in the NBPF3 gene (near the ALPL gene) has been found in multiple studies to be associated with reduced levels of vitamin B-6, possibly due to faster than normal clearance of this vitamin from the bloodstream<sup>47,48</sup>. Individuals with a C/C or C/T genotype had lower levels of B-6 than those with the T/T genotype. Therefore, if your genotype is C/C or C/T, you will get a result of "Optimize Intake." If your genotype is T/T, it is suggested that you "Stay Balanced" and maintain a healthy diet. The studies we report observed associations between vitamin levels and particular genotypes; however, that does not mean that your levels are out of balance. You should ensure that you are eating a healthy diet and discuss this result with your physician. The recommended intake of vitamin B-6 for most adults is 1.3 to 1.7 milligrams per day.



#### **OPTIMIZE INTAKE**

People with your genotype are more likely to have lower blood levels of vitamin B-6. You may optimize your intake of vitamin B-6 by paying attention to your diet and eating foods rich in vitamin B-6.

### YOUR RELATED GENES

Gene Tested	Your Genotype	Scientific Strength
NBPF3-rs4654748	C/C	****



#### NUTRITIONAL NEEDS VITAMIN B12

Vitamin B-12 plays an important role in how your brain and nervous system function. It helps to keep red blood cells healthy and is a critical component for synthesis and regulation of your DNA<sup>49</sup>. Vitamin B-12 is found naturally in foods of animal origin including meat, fish, poultry, eggs and milk products. A healthy diet will typically provide sufficient B-12, although vegetarians, vegans, older people, and those with problems absorbing B-12 due to digestive system disorders may be deficient. Symptoms of vitamin B-12 deficiency can vary, but may include fatigue, weakness, bloating, or numbness and tingling in the hands and feet. The recommended intake for adults is 2.4 micrograms per day.

Multiple genetic studies have identified a marker in the gene FUT2 as being associated with lower levels of B-12 in the blood 48,50,47. This effect may be due to reduced absorption of B-12 in the gut 48. People with G/G or A/G genotypes are recommended to "Optimize Intake" because they may have lower levels of B-12. Eating foods rich in vitamin B-12 can promote healthy levels of B-12, especially for those over the age of 50. People with the A/A genotype should "Stay Balanced" and maintain a healthy diet. The studies we report observed associations between vitamin B-12 levels and particular genotypes; however, that does not mean that your levels are out of balance. You should ensure that you are eating a healthy diet and discuss this result with your physician.

#### YOUR RESULT

#### STAY BALANCED

Your genotype is not associated with lower blood levels of vitamin B12. You should stay balanced and maintain a healthy diet.

#### YOUR RELATED GENES

Gene Tested	Your Genotype	Scientific Strength
FUT2-rs602662	A/A	****

SEX: MALE ACC #: C4215580 DATE:



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

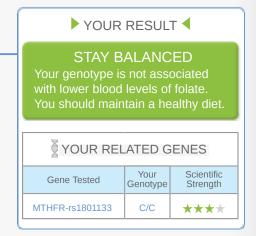
PAGE 24



#### NUTRITIONAL NEEDS FOLATE - FOLIC ACID

Folate is found in many foods, such as green leafy vegetables like chard or kale, as well as beans, lentils, fruits and fortified grains. This nutrient plays a role in protein metabolism, as well as DNA repair<sup>51</sup>. Folate can lower the blood level of homocysteine, a substance linked to cardiovascular disease at high levels<sup>52</sup>. Diets rich in folate have been associated with reduced risk of cardiovascular disease<sup>53</sup>. Folate is particularly important early in pregnancy for preventing some birth defects<sup>51</sup>. For this reason, pregnant women or women intending to become pregnant are advised an elevated recommended daily intake of 600 micrograms of folate. The recommended intake of folate for most adults is 400 micrograms per day.

A relatively common variant in the MTHFR gene, known as C677T (rs1801133), has been associated with lowered folate and elevated homocysteine levels in the blood<sup>52</sup>. Hence, people with a T/T or C/T genotype should "Optimize Intake" of folate. People with the C/C genotype should "Stay Balanced" and maintain a healthy diet. The studies we report observed associations between vitamin levels and particular genotypes; however, that does not mean that your levels are out of balance. You should ensure that you are eating a healthy diet and discuss this result with your physician.



SEX: MALE ACC #: C4215580

DATE:



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

PAGE 25



#### NUTRITIONAL NEEDS VITAMIN A

Vitamin A is a nutrient that describes a number of related compounds, including retinol, retinal, and retinoic acid. Vitamin A is critical for numerous functions in the body, including healthy vision, immune system action, bone growth, reproduction, and the proper regulation of gene expression <sup>54,55,56,57</sup>. The recommended intake of vitamin A for most adults is 700 to 900 micrograms per day.

Much of the vitamin A found in your body is derived from beta-carotene, a nutrient found in some plants and foods, such as pumpkin, carrots, sweet potatoes and spinach. A genetic study has found that vitamin A conversion from beta-carotene is impaired in women carrying variants of the BCMO1 gene<sup>58</sup>. This association has not been studied in men.

Those with a result of "Optimize Intake" may bypass this effect by consuming adequate amounts of preformed vitamin A, which can be found in fortified milk and breakfast cereals, as well as in multivitamins containing retinyl palmitate or retinyl acetate 59,60. People who receive a "Stay Balanced" outcome should maintain a healthy diet. An additional outcome in this report is "Inconclusive," which means that there was not enough scientific evidence to determine how your genotype relates to the efficiency of converting beta-carotene to vitamin A. The study we report observed associations between vitamin A levels and particular genotypes. However, that does not mean that your levels are out of balance. You should eat a healthy diet and speak with your physician before making specific changes to your dietary regimen.



#### YOUR RESULT

#### **INCONCLUSIVE**

There is not enough scientific evidence to determine how you genotype relates to efficiency ir converting beta-carotene into vitamin A.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
BCMO1-rs7501331	C/C	****
BCMO1- rs12934922	T/T	****

SEX: MALE ACC #: C4215580 DATE:



**NUTRITIONAL NEEDS** 

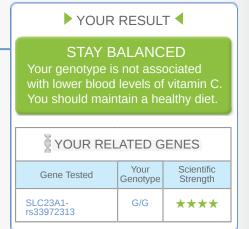
PATHWAY GENOMICS

PAGE 26



#### NUTRITIONAL NEEDS VITAMIN C

Vitamin C, or L-ascorbic acid, must be acquired from dietary sources, as humans are unable to synthesize it. Some dietary sources of vitamin C include lemons, oranges, red peppers, watermelons, strawberries and citrus juices or juices fortified with vitamin C. While a severe deficiency of vitamin C ultimately leads to scurvy, variations in vitamin C levels have also been associated with a wide range of chronic complex diseases, such as atherosclerosis, type 2 diabetes and cancer 1. These associations are thought to result from a contribution of vitamin C as an antioxidant, as well as its role in the synthesis of collagen and various hormones. After ingestion, the vitamin C in one's diet gets transported across the cell membrane via transport proteins, one of which is SLC23A1. A recent study of over 15,000 people found that the A allele of a variant in SLC23A1 was associated with decreased levels of circulating vitamin C 1. Therefore, if your genotype is A/A or A/G, you will get a result of "Optimize Intake." People with a G/G genotype should "Stay Balanced" and maintain a healthy diet.





#### NUTRITIONAL NEEDS VITAMIN D

Vitamin D is important for the absorption and utilization of calcium, which is beneficial for maintaining good bone health<sup>63</sup>. Exposure to sunlight is an important determinant of a person's vitamin D level, since there are few natural dietary sources of vitamin D. While sunscreen use blocks skin production of vitamin D, excessive sun exposure is a risk factor for skin cancer and related conditions, and is not recommended. Dietary sources of vitamin D include some fatty fish, fish liver oils, and milk or cereals fortified with vitamin D. The recommended intake of vitamin D for most adults is 600 IUs per day. About 115 IUs of vitamin D is found in one cup of vitamin D-fortified, non-fat, fluid milk.

Multiple genetic studies have identified a variant in the GC gene that codes for the vitamin D-binding protein that is associated with decreased blood levels of 25-hydroxyvitamin D, which is the major circulating form of vitamin  $D^{64,65}$ . People with the G/G or G/T genotype at this genetic marker may be susceptible to lower blood vitamin D levels due to reduced ability to transport vitamin D in the body. Therefore, these people may need to "Optimize Intake" of vitamin D. People with a T/T genotype are advised to "Stay Balanced" and maintain a healthy diet. The studies we report observed associations between vitamin D levels and certain genotypes; however, that does not mean that your levels are out of balance. You should eat a healthy diet and speak with your physician before making specific changes to your dietary regimen.



#### STAY BALANCED

Your genotype is not associated with lower levels of vitamin D (plasma 25-hydroxyvitamin D levels). However, other factors, such as diet and exposure to sunlight, play an important role in regulating levels of vitamin D in blood.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
GC-rs2282679	T/T	****

SEX: MALE ACC #: C4215580 DATE:



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

PAGE 27



# NUTRITIONAL NEEDS VITAMIN E

Vitamin E is a group of eight antioxidant molecules, of which alpha-tocopherol is the most abundant in the body. Vitamin E functions to promote a strong immune system and regulates other metabolic processes<sup>66,67</sup>. The recommended intake of vitamin E for most adults is 15 milligrams per day. Note that synthetic varieties of vitamin E found in some fortified foods and supplements are less biologically active. Sources of naturally-occurring vitamin E in foods are vegetable oils, green leafy vegetables, eggs and nuts.

One study of 3,891 individuals found that people with the A/A or A/C genotypes at an intergenic marker, rs12272004, near the APOA5 gene, had increased plasma levels of alpha-tocopherol<sup>68</sup>. Therefore, they should "Stay Balanced" and maintain a healthy diet. This is good news since increased vitamin E levels are associated with decreased frailty and disability in old age<sup>69</sup>. People with the C/C genotype were not associated with increased levels of alpha-tocopherol, and hence they would need to "Optimize Intake" of vitamin E through the increased intake of foods rich in vitamin E. Keep in mind, however, that most adults normally do not take in adequate amounts of vitamin E on a daily basis<sup>70</sup>, so keeping an eye on your vitamin E intake is good advice for anyone. The studies we report observed associations between vitamin E levels and certain genotypes; however, that does not mean that your levels are out of balance. You should eat a healthy diet and speak with your physician before making specific changes to your dietary regimen.



#### **OPTIMIZE INTAKE**

Your genotype is not associated with increased alpha-tocopherol levels, which is one compound that makes up vitamin E. You may optimize your intake of vitamin E by eating foods rich in this nutrient.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
INTERGENIC- rs12272004	C/C	****



**NUTRITIONAL NEEDS** 

PATHWAY GENOMICS

PAGE 28

#### NUTRIENTS, WEIGHT MANAGEMENT AND GENETICS

New studies continue to emerge that demonstrate links between nutrients and genetics that show benefits in health and weight loss. Some of the benefits that scientists observe in relation to genes are listed below.

Important: The genes and associated benefits listed below are not part of your genetic test. The content on this page is informational.



NUTRIENT/FOOD	POTENTIAL HEALTH & WEIGHT LOSS BENEFITS	ASSOCIATED GENE(S)
Resveratrol	Weight Loss, Decrease Weight Gain	SIRT1, PPARA, PPARG, ER
Polyphenols (tea)	Decrease Weight Gain	PPARG
Conjugated Linoleic Acid (CLA)	Fat Burning, Weight Loss	PPARA, PPARG
Ispoprenols (farnesol)	Weight Loss	PPARA, PPARG
Abietic Acid	Weight Loss	PPARG
Capsaicin (Hot Pepper)	Weight Loss, Anti-inflammatory	PPARG
Phytol (Chlorophyll)	Weight Loss	PPARA
Auraptene (Citrus)	Weight Loss	PPARA, PPARG
Isohumulone (Hops)	Weight Loss	PPARA, PPARG
Guggulsterone (Gugle)	Weight Loss	Farnesoid X Receptor
Soy/Genistein	Weight Loss	Steroid Receptors: Estrogen, Androgen, Progesterone
Diosgenin	Weight Loss	Steroid Receptors: Progesterone
Ginseng	Weight Loss	Steroid Receptors: Estrogen
Hyperforin	Weight Loss	Pregnane X Receptor
Alpha-lipoic Acid	Reduction of Overeating	AMPK Inhibitor
Anthocyanins (Pigment)	Overall Health Benefit	Adiponectin
Licorice LFO (Polyphenols)	Overall Health Benefit	FA synthase
Pomegranate Extract (Lenolenic Acid)	Overall Health Benefit	b-oxidation/PPARA

SEX: MALE ACC #: C4215580 DATE:



**EXERCISE** 

PAGE 29



# EXERCISE HAS LONG BEEN SHOWN TO PROVIDE MANY HEALTH BENEFITS.

Studies have shown a link between genetics and exercise, and how people respond to exercise for weight loss and other health benefits. A few examples of this link include the ACE and ACTN3 genes and the association with elite athlete status, as well as the LPL gene and its connection to the loss of body fat in response to exercise. A summary of your results is listed below.



#### ▶ YOUR RESULTS ◀

Be	ENDURANCE TRAINING	page:30	ENHANCED BENEFIT
3º	STRENGTH TRAINING	page:31	LESS BENEFICIAL
11	STRENOTH TRAINING	page.01	EEGG BEIVEI 101/12

Be	AEROBIC CAPACITY (VO	O2MAX) pa	age:31	DECREASED
		,		

~			
	MUSCLE POWER	naga:22	LESS MUSCLE POWER
	MUSCLE POWER	page:32	LLOO WOOCLE FOWER

Be	ACHILLES TENDINOPATHY	page:32	TYPICAL

	WEIGHT LOSS RESPONSE TO		EXERCISE STRONGLY
Be	EXERCISE	page:33	RECOMMENDED

	BLOOD PRESSURE RESPONSE		EXERCISE STRONGLY
3	TO EXERCISE	page:33	RECOMMENDED

	HDL (GOOD) CHOLESTEROL		
Be	RESPONSE TO EXERCISE	page:34	NORMAL BENEFIT

	LOSS OF BODY FAT RESPONSE		
Be	TO EXERCISE	page:34	NORMAL BENEFIT

	INSULIN SENSITIVITY RESPONSE		
3º	TO EXERCISE	page:34	ENHANCED BENEFIT



PAGE 30



#### YOUR EXERCISE RECOMMENDATIONS

- ✓ Your genetics are associated with enhanced health benefits from endurance exercises, such as mid-long distance walking, jogging and bicycling. Weight resistance exercises may be less beneficial.
- ✓ You have a genetic variant associated with being overweight. You can lower your chances by leading a physically active lifestyle.
- ✓ You have a genetic variant associated with elevated blood pressure if you are not physically active. Exercise may help you manage your blood pressure.
- ✓ Continue a vigorous exercise regimen after losing weight. You have genes that are associated with an increased chance of gaining weight back.





# EXERCISE ENDURANCE TRAINING

Endurance training is generally used to describe exercise that is done for a longer duration with moderate intensity. Most people can benefit from a combination of endurance, high intensity and resistance exercises. Some people have genetic markers that are associated with "Enhanced Benefit" from endurance training, while others will gain "Normal Benefit." The studies that were used to calculate your result tested responses to a 20-week endurance training program <sup>71,72,73</sup>. This result can be used to help tailor your exercise routine. Always consult your physician or health care provider before beginning any exercise program.



#### **ENHANCED BENEFIT**

Endurance training may provide enhanced health benefits to people with your genotype.

YOUR RELATED GENES		
Your Genotype	Scientific Strength	
C/C	****	
A/A	****	
T/C	****	
	Your Genotype C/C	

SEX: MALE ACC #: C4215580 DATE:



**EXERCISE** 

INSIG2-rs7566605

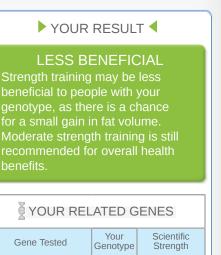


PAGE 31



# EXERCISE STRENGTH TRAINING

Strength training can be described as exercises that incorporate the use of opposing forces to build muscle. The possible outcomes in this report are "Beneficial" and "Less Beneficial." In a small study of young adult men, those with the C/G or C/C genotypes at rs7566605 were more likely to experience increased fat volume after participating in 12 weeks of resistance training, and thus strength training was "Less Beneficial" This association has not been identified in women.

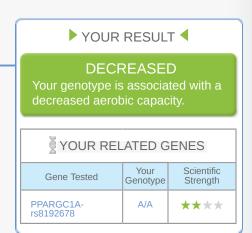


\*\*\*\*



# EXERCISE AEROBIC CAPACITY (VO2MAX)

Maximal oxygen uptake (VO2max) is widely used as the best measure of an individual's cardiorespiratory fitness. VO2max is defined as the maximum volume of oxygen per unit time that an individual uses at maximum exertion. The baseline VO2max level can vary depending on age, gender, past medical history, current health and level of physical activity. However, anyone can increase their fitness and VO2max by endurance training. Elite athletes in endurance sports, such as cross-country skiing and long-distance running, have a higher VO2max than elite athletes in power sports, such as wrestling and weightlifting 75. The rs8192678 SNP in the PPARGC1A gene, which is a key regulator of energy metabolism, was associated with baseline VO2max (L/min) in a study of 303 Spanish and British men 76. The G/G and G/A genotypes of rs8192678 were associated with "Typical" VO2max, whereas the A/A genotype was associated with a "Decreased" VO2max. This association has not been studied in women. Please remember that you can always increase your VO2max and fitness by endurance training, even if you start with a decreased aerobic capacity.







PAGE 32





#### **EXERCISE** MUSCLE POWER

Do you have a genetic variant that is found in nearly all sprinters qualified for top-level competitions like the Olympic Games? The so-called "sprinter gene" refers to the functional version of the ACTN3 gene, which contains information for making a protein found in fast-twitch muscle fibers<sup>77</sup>. The protein and the fast-twitch muscle fibers are important in generating explosive bursts of force. This is why the functional version of ACTN3 is also seen with high frequencies in other elite power-oriented athletes, such as weightlifters<sup>78</sup>. About 80% of people have at least one functional copy of the ACTN3 gene, which can lead to enhanced muscle power. However, having functional ACTN3 is only one of a myriad of genetic and non-genetic factors that contribute to the success of elite athletes. The remaining 20% of people, who do not have a functional copy of ACTN3, may have less muscle power and are less likely to be world-class sprinters or weightlifters, but their chance to excel may not be affected in sports that require other types of body performance, such as endurance and nimbleness.



#### YOUR RESULT

#### LESS MUSCLE POWER

Your genotype is rare in elite and weightlifters. However, this gene is only one of many factors determining a person's athletic

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
CTN3-re1815730	T/T	4444



#### **EXERCISE ACHILLES TENDINOPATHY**

The Achilles tendon connects your calf muscles to your heel bone. Tendinopathy describes either the inflammation or tiny tears to the tendon. People who play sports and runners who place stress on the Achilles tendon have the greatest likelihood of tendinopathy. If you have a G/G genotype you may be more "Injury-Prone," while other genotypes have a "Typical" likelihood of developing Achilles tendinopathy. In a small study, people with the G/ G genotype at rs679620 (MMP3) had 2.5 times more chance of developing Achilles tendinopathy compared to other genotypes<sup>79</sup>.



## YOUR RESULT **TYPICAL**

Your genotype is associated with a typical likelihood of Achilles

YOUR RELATED GENES		
Your Genotype	Scientific Strength	
A/A	****	
	Your Genotype	





FTO-rs1121980

PAGE 33



#### **EXERCISE** WEIGHT LOSS RESPONSE TO EXERCISE



Exercise is a large part of many weight loss plans, as it is a crucial tool for weight control for everyone. The possible outcomes in this report are "Exercise Strongly Recommended" and "Exercise Recommended." If your report shows "Exercise Strongly Recommended," your genotype has been shown to be associated with a tendency to be overweight, and exercise is strongly recommended for you. If your report shows "Exercise Recommended," you have one less risk factor for being overweight. However, this should not be taken as one less reason to exercise, because being physically active is beneficial to all people, regardless of genetic makeup. People with the "Exercise Strongly Recommended" outcome contain a specific variant (T allele) in the genetic marker rs1121890 of the FTO gene, which has been shown to be associated with increased body mass index (BMI) and waistline 80,81. However, a large study showed that people who have this variant could reduce their propensity to increased BMI by being physically active<sup>81</sup>.

#### YOUR RESULT

#### **EXERCISE STRONGLY RECOMMENDED**

You have a genetic variant that is associated with being overweight an active lifestyle are strongly

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength

C/T

\*\*\*



#### **EXERCISE** BLOOD PRESSURE RESPONSE TO EXERCISE

High blood pressure, also known as hypertension, is a common health issue. It has been estimated that a majority of people will have hypertension at some time in their lives. A genetic variant in the EDN1 gene has been shown to increase the likelihood of hypertension in people who were low in cardiorespiratory fitness, which refers to the ability of the heart and lungs to provide muscles with oxygen for physical activity<sup>82</sup>. This genetic variant did not show an effect in people who were high in cardiorespiratory fitness. If you have this variant, your result is "Exercise Strongly Recommended," since you may need to exercise to reduce your chances of hypertension. If you do not have the variant, your result is "Exercise Recommended," since exercise is still the right decision to manage other risk factors for high blood pressure you may have.

#### YOUR RESULT

#### **EXERCISE STRONGLY** RECOMMENDED

Your genotype is associated with blood pressure, if you have low fitness levels. Exercise may help you manage your blood pressure.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
EDN1-rs5370	G/T	****



PATHWAY GENOMICS

PAGE 34



#### **EXERCISE**

#### HDL (GOOD) CHOLESTEROL RESPONSE TO EXERCISE

One of the health benefits of exercise can be the improvement of your cholesterol. HDL cholesterol is known as the good cholesterol, and having more HDL is beneficial. Most people can improve their HDL levels by exercising. In the Heritage Family Study, people with the A/G and G/G genotypes were more likely to have an "Enhanced Benefit" in their HDL levels by exercising <sup>73</sup>. People with "Normal Benefit" may also increase their HDL levels by exercising, but may not experience an enhanced effect.



#### **NORMAL BENEFIT**

Your genotype is associated with a typical increase in HDL (good) cholesterol in response to a 20week endurance training program.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
PPARD-rs2016520	A/A	++++



#### **EXERCISE**

#### LOSS OF BODY FAT RESPONSE TO EXERCISE



Many people exercise to lose body fat. If you have a specific genotype in the gene LPL, you may have an "Enhanced Benefit" from exercise to lose body fat<sup>71</sup>. If you have the "Normal Benefit" genotype, you will still experience fat reduction if you exercise, but it might take more effort. The study was based on women who participated in a 20-week endurance training program. This association has not been identified in men.

#### YOUR RESULT

#### NORMAL BENEFIT

Your genotype is associated with a typical reduction in body fat mass and percent of body fat in response to exercise.

YOUR RELATED GENES		
Your Genotype	Scientific Strength	
C/C	****	
	Your Genotype	



#### **EXERCISE**

#### INSULIN SENSITIVITY RESPONSE TO EXERCISE

Insulin sensitivity is a good thing. Insulin in your body helps control your response to glucose, commonly known as sugar. Having an increased insulin sensitivity means that the body has a better ability to process sugar. The opposite of insulin sensitivity is called insulin resistance, which is linked to obesity and type 2 diabetes. Most people have a beneficial response to exercise, resulting in increased insulin sensitivity. According to a study, people with C/C or C/T genotypes, at a marker in the LIPC gene, showed an "Enhanced Benefit," compared to those with a T/T genotype<sup>72</sup>. Although people with T/T genotypes are likely to gain "Less Benefit" in insulin sensitivity from exercise training, exercise remains important in many other aspects of their health.

#### YOUR RESULT

#### **ENHANCED BENEFIT**

Your genotype is associated with enhanced insulin sensitivity in response to exercise.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
LIPC-rs1800588	C/T	****

SEX: MALE ACC #: C4215580 DATE:



YOUR BODY AND WEIGHT

PAGE 35





Your report includes how your genes may relate to your metabolism, if you are likely to maintain weight loss, as well as your predisposition for obesity. A summary of your results is provided below.

#### ▶ YOUR RESULTS ◀

OBESITY

page:37

**ABOVE AVERAGE** 

-1-

WEIGHT LOSS-REGAIN

page:37

MORE LIKELY TO GAIN WEIGHT BACK

**METABOLISM** 

page:38

NORMAL

-\$-

ADIPONECTIN LEVELS

page:38

**TYPICAL** 



SEX: MALE ACC #: C4215580 DATE:



YOUR BODY AND WEIGHT

PAGE 36





# YOUR BODY AND WEIGHT YOUR ACTUAL WEIGHT (BMI)



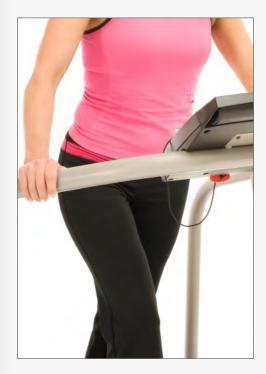
#### **Normal**

POUR BM body mass index

24.5

,	
Category	BMI range - kg/m2
Underweight	Less than 18.5
Normal	from 18.5 to 24.9
Overweight	from 25 to 29.9
Obese	from 30 to 34.9
Clinically Obese	from 35 to 39.9
Extremely Obese	40 or greater

Body mass index (BMI) is a measure of body fat based on height and weight that applies to adult men and women. BMI is usually represented in kg/m2. Your BMI was calculated using your survey responses for weight and height. If your BMI is not listed here, you may not have completed those responses in the survey. Your actual weight is a result of a combination of factors including lifestyle, environment and genetics. Your Obesity Index result is a measure of your likelihood, based on genetics, to have a BMI over 35 (clinically or extremely obese). Since your weight is affected by many factors, it is possible for your Obesity Index result to be very different than your actual weight. The important point is that the genetics of obesity do not lead to an inevitable outcome. Many people have a choice of managing lifestyle to counteract genetics. For example, some people that are of normal weight BMI can have an Obesity Index of above average or high. This example is commonly seen in someone who is controlling diet, nutrition, eating behaviors and/or exercise to manage their body weight. The opposite can also be true. Some people who have an actual BMI in the obese categories can have an Obesity Index of average, below average or low. This case can sometimes be explained by lifestyle choices, environment or other health factors that have led a person to become obese without having the genetics associated to obesity.



SEX: MALE ACC #: C4215580 DATE:



YOUR BODY AND WEIGHT

PAGE 37





## YOUR BODY AND WEIGHT OBESITY

Obesity is influenced by both genetic and environmental factors. Approximately 40 to 70% of an individual's susceptibility to obesity is inherited<sup>1</sup>. When someone reaches a body mass index (BMI) of 30 to 35 (clinically obese) or above 40 (morbidly obese), genetic factors with strong effects are likely to be involved. There are 2 possible outcomes of this test: "Average" and "Above Average". An "Above Average" outcome does not mean that you are obese, it only means that you have a higher than average genetic likelihood for a high BMI.

Your genetic predisposition to obesity is determined from your genotypes at variants in the FTO (fat mass and obesity associated) and MC4R (melanocortin-4 receptor) genes. The association of these genes to obesity is well-established. The MC4R gene is expressed in the brain's hunger center and is involved in regulating energy balance<sup>83</sup>. Rare mutations in the MC4R gene have been shown to cause a rare, inherited form of obesity. FTO is less well-understood, but is also believed to be important for controlling feeding behavior and energy balance<sup>84</sup>. Your test result includes common variants that have been confirmed in many large genetic studies (including multiple studies of over 38000 individuals) to be associated with a predisposition for high BMI and/or obesity, you can mitigate your risks by eating a proper diet, exercising and reducing stress<sup>89,90</sup>.



#### **ABOVE AVERAGE**

Your genetic profile indicates an above average predisposition for being overweight.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
FTO-rs9939609	A/T	****
MC4R-rs17782313	C/T	++++



# YOUR BODY AND WEIGHT WEIGHT LOSS-REGAIN

There are genes associated with the tendency to gain weight back after a person loses weight, and there are genes that protect a person from weight regain. In one study, people with the G/G genotype at a marker in the ADIPOQ gene were "More Likely to Gain Weight Back," while people with other genotypes were more likely to show "Weight Loss Maintained" 11 tis best after losing weight to maintain a healthy diet, exercise and nutrition plan to keep the extra pounds off and support long-term health.



# MORE LIKELY TO GAIN WEIGHT BACK You may have difficulty keeping weight off after losing weight. YOUR RELATED GENES Gene Tested Your Genotype Scientific Strength ADIPOQrs17300539 G/G \*\*\*\*\*

SEX: MALE ACC #: C4215580

DATE:



YOUR BODY AND WEIGHT

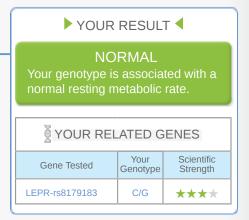
PAGE 38





## YOUR BODY AND WEIGHT METABOLISM

Metabolism describes the way your body burns energy (calories) and tends to have a strong correlation to managing your weight. Resting metabolism is how your body burns energy while at rest. People with a "Fast" metabolism can sometimes eat more food with little exercise and not gain weight. People with a "Normal" metabolism tend to require average amounts of food intake and average amounts of exercise to maintain weight. A genetic marker in the leptin receptor (LEPR) is associated with interactions in your brain that trigger how and when you burn energy. People with a C/C genotype tend to have an increased resting metabolic rate, or "Fast" metabolism, while people with C/G or G/G genotypes are not associated with an increased resting metabolic rate; therefore, they have a "Normal" metabolism. However, having this genetic variant is only one of many other genetic and non-genetic factors that contribute towards your metabolism.

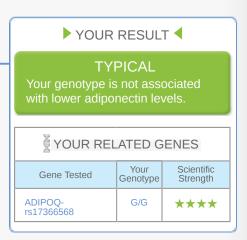




## YOUR BODY AND WEIGHT ADIPONECTIN LEVELS

Exercise is a common method of increasing your metabolism.

Adiponectin is a hormone that is produced by fat cells and functions in the body to trigger your liver and muscles to get energy from  ${\rm fat}^{93}$ . Higher levels of adiponectin are considered good for weight loss and health  $^{94}$ . Your health care provider can test your adiponectin levels. If you have low levels, losing weight may be a good way to increase your adiponectin levels  $^{93}$ . A variant in the adiponectin gene (ADIPOQ) is associated with adiponectin levels. People with A/A or A/G genotypes were associated with "Possibly Low" levels of adiponectin, while those with the G/G genotype had "Typical" levels  $^{95}$ .



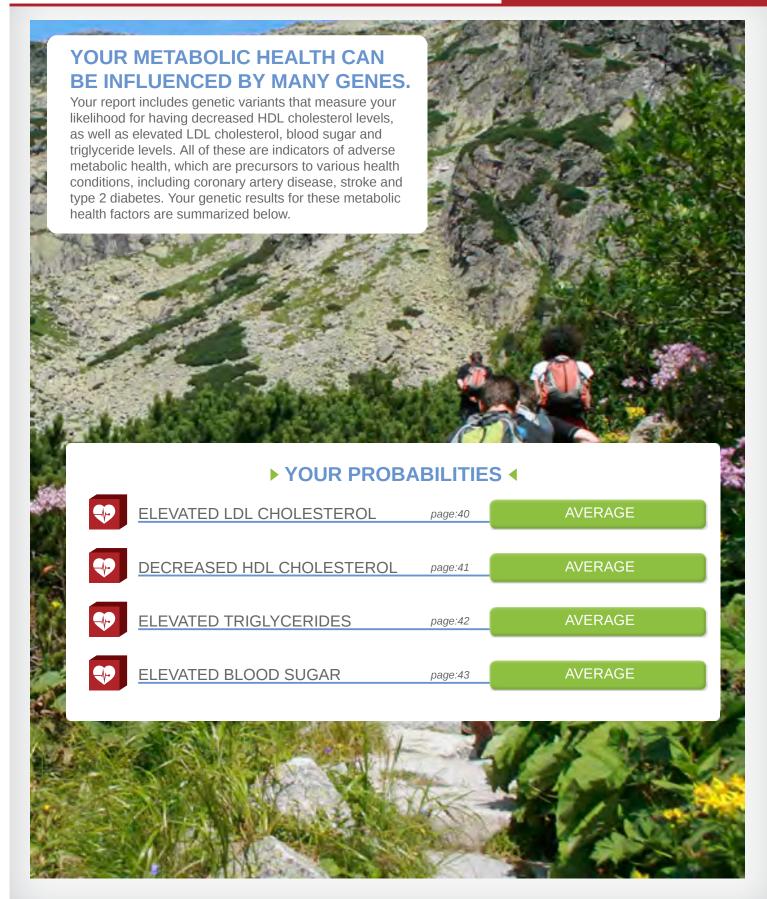
SEX: MALE ACC #: C4215580

DATE:



METABOLIC HEALTH FACTORS

PATHWAY GENOMICS



SEX: MALE ACC #: C4215580

DATE:



METABOLIC HEALTH FACTORS

PAGE 40



## YOUR HEALTH RECOMMENDATIONS

- ✓ You have a higher than average genetic risk for obesity. If you are having difficulties controlling your weight, you should take action as soon as possible to reduce your risks. Regardless of your current weight, your Pathway Fit report provides you with guidance that can help you work actively to achieve and maintain a healthy weight.
- ✓ To ensure that your diet and exercise program are helping you maintain optimal health, work with your physician to track your blood sugar and cholesterol levels.
- ✓ You have an "Above Average" genetic likelihood for some metabolic health factors, but you do not have a "High" likelihood in any category. Discuss these results with your physician.





## METABOLIC HEALTH FACTORS ELEVATED LDL CHOLESTEROL

Low-density lipoprotein (LDL) is the type of cholesterol that can become dangerous if you have too much of it. Like gunk clogging up your kitchen drain, LDL cholesterol can form plaque and build up in the walls of your arteries. This can make your arteries narrower and less flexible,;putting you at risk for conditions like a heart attack or stroke. Optimally, LDL levels should be less than 100 mg/dl. Near-optimal levels range from 100 to 129 mg/dl and borderline high from 130 to 159 mg/dl. A score greater than 160 mg/dl is high and greater than 190 mg/dl is very high. Your physician can measure your cholesterol levels.

A genetic result of "High" or "Above Average" does not mean you have elevated LDL cholesterol levels, but tells you that you may have a genetic propensity for elevated LDL cholesterol levels. On the other hand, a result of "Low" or "Below Average," tells you that you have a lower than average genetic:likelihood for elevated LDL cholesterol levels. However, you could still develop problems with your LDL levels as a result of your diet and other factors.: This report is based on genetic variants studied in over 19,000 individuals. A genetic result of "High" means that you share a similar genetic profile with individuals from the Framingham Heart Study who had elevated LDL cholesterol levels measuring. on average, above 139 mg/dl with approximately 25% of individuals measuring above 160 mg/dl<sup>8</sup>. A genetic result of "Above Average" means that you share a similar genetic profile with individuals measuring, on average, above 130 mg/ dl LDL with approximately 17% of individuals measuring above 160 mg/dl LDL cholesterol<sup>8</sup>. A genetic result of "Average" means that you share a similar genetic profile with individuals measuring, on average, near-optimal LDL cholesterol levels. Diet plays an important part in LDL levels. Processed foods and foods high in trans fat contribute to elevated LDL levels.



#### **AVERAGE**

Based on your genetic profile you have an average likelihood for elevated LDL cholesterol levels.

## YOUR RELATED GENES

Gene Tested	Your Genotype	Scientific Strength
ABCG8-rs6544713	C/C	****
LDLR-rs6511720	G/G	****
MAFB-rs6102059	C/T	****
APOB-rs515135	G/G	****
HMGCR- rs3846663	T/T	****
HNF1A-rs2650000	C/C	****
INTERGENIC- rs1501908	G/C	****
CELSR2- rs12740374	G/T	****
PCSK9- rs11206510	T/T	***
NCAN-rs10401969	T/T	****

SEX: MALE ACC #: C4215580 DATE:



MÉTABOLIC HEALTH FACTORS PATHWAY GENOMICS

PAGE 41



## METABOLIC HEALTH FACTORS DECREASED HDL CHOLESTEROL

High-density lipoprotein (HDL) cholesterol is known as good cholesterol, because high levels of HDL cholesterol seem to protect against heart attack, while low levels of HDL cholesterol (less than 40 mg/dL) increase the risk of heart disease <sup>96</sup>. While multiple mechanisms are known to account for this, the major one is thought to be the role of HDL in transporting excess cholesterol away from the arteries and back to the liver, where it is passed from the body <sup>97</sup>. Your HDL cholesterol can be measured with a simple blood test. In men, typical HDL cholesterol levels range from 40 to 50 mg/dl. In women, female hormones cause typical HDL cholesterol levels to range from 50 to 60 mg/dl; however, after menopause there is a tendency for decreased HDL cholesterol levels. Foods containing trans fats can lower HDL cholesterol levels, which is unhealthy. Cholesterol levels should be monitored by your physician.

A genetic result of "High" or "Above Average" does not mean you have decreased HDL cholesterol levels, but tells you that you may have a high propensity for decreased HDL cholesterol levels. On the other hand, a result of "Low" or "Below Average," tells you that you have a lower than average propensity for decreased HDL cholesterol levels. Our genetic testing is based on genetic variants studied in over 19,000 individuals. A result of "High" means that you share a similar genetic profile with individuals from the Framingham Heart Study who had decreased HDL cholesterol levels measuring, on average, below 46 mg/dl with approximately 37% of individuals measuring below 40 mg/dl<sup>8</sup>. On the other hand, a result of "Above Average" means that you share a similar genetic profile with individuals measuring, on average, below 50 mg/dl HDL cholesterol with approximately 30% of individuals measuring below 40 mg/dl HDL cholesterol<sup>8</sup>.

## YOUR PROBABILITY

#### **AVERAGE**

Based on your genetic profile you have an average likelihood for decreased HDL cholesterol levels.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
ZNF259-rs964184	C/C	****
PLTP-rs7679	T/T	****
LIPG-rs4939883	C/T	****
GALNT2- rs4846914	A/G	****
TTC39B-rs471364	A/A	****
ANGPTL4- rs2967605	G/A	****
CETP-rs247616	C/T	****
KCTD10- rs2338104	C/G	****
LCAT-rs2271293	G/G	****
ABCA1-rs1883025	G/G	****
HNF4A-rs1800961	C/C	****
FADS1-rs174547	T/T	****
LPL-rs12678919	A/A	****
LIPC-rs10468017	C/T	****

SEX: MALE ACC #: C4215580

DATE:



METABOLIC HEALTH FACTORS

PATHWAY

PAGE 42



# METABOLIC HEALTH FACTORS ELEVATED TRIGLYCERIDES

Triglyceride is the chemical term for fat as it is stored in your body. People with elevated triglycerides are at risk of conditions, such as coronary artery disease or type 2 diabetes. Having higher triglycerides is often associated with poor lifestyle choices, such as lack of exercise, excessive alcohol consumption, cigarette smoking, excessive refined carbohydrate consumption and being overweight. A normal triglyceride score is under 150 mg/dl. Triglyceride levels in the range of 150 to 199 mg/dl are defined as borderline high, with over 200 mg/dl considered high and over 500 mg/dl very high. Your triglyceride levels can be monitored by your physician.

A result of "High" or "Above Average" does not mean you have elevated triglyceride levels, but tells you that you may have a propensity for elevated triglycerides levels. On the other hand, a genetic test result of "Low" or "Below Average," tells you that you have a lower than average likelihood for elevated triglyceride levels. The genetic test is based on genetic variants studied in over 19,000 individuals. A genetic result of "High" means that you share a similar genetic profile with individuals from the Framingham Heart Study who had elevated triglyceride levels measuring on average above 150 mg/dl with approximately 31% of individuals measuring above 200 mg/dl<sup>8</sup>.



## YOUR PROBABILITY

#### **AVERAGE**

Based on your genetic profile you have an average likelihood for elevated triglyceride levels.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
ZNF259-rs964184	C/C	****
XKR6-rs7819412	A/A	****
PLTP-rs7679	T/T	****
APOB-rs7557067	A/G	****
MLXIPL-rs714052	T/T	****
TRIB1-rs2954029	A/T	****
FADS1-rs174547	T/T	****
NCAN-rs17216525	C/C	****
LPL-rs12678919	A/A	****
GCKR-rs1260326	C/T	****
ANGPTL3- rs10889353	A/A	****

SEX: MALE ACC #: C4215580

DATE:



METABOLIC HEALTH FACTORS

PAGE 43





# METABOLIC HEALTH FACTORS ELEVATED BLOOD SUGAR



Elevated blood sugar is a health condition that results from higher than normal levels of the sugar (glucose) in the blood plasma. High blood sugar levels are measured as a reading greater than 140 mg/dl or a fasting plasma glucose level of greater than 100 mg/dl. High blood sugar levels often indicate a condition called insulin resistance and can lead to type 2 diabetes. Your physician can directly measure blood sugar or you can use a blood test at home to check your blood sugar.

A genetic result of "High" or "Above Average" does not mean you have elevated blood sugar levels, but tells you that you may have a genetic propensity for elevated blood sugar levels. On the other hand, a result of "Low" or "Below Average," tells you that you have a lower than average genetic likelihood for elevated blood sugar levels. This report is based on genetic variants identified in a study of more than 100,000 individuals<sup>9,98</sup>.

## YOUR PROBABILITY

#### **AVERAGE**

Based on your genetic profile you have an average likelihood for elevated blood sugar levels.

YOUR RELATED GENES		
Gene Tested	Your Genotype	Scientific Strength
MADD-rs7944584	A/A	****
TCF7L2-rs7903146	C/T	****
GCKR-rs780094	G/A	****
GLIS3-rs7034200	A/A	****
G6PC2-rs560887	A/A	****
GCK-rs4607517	G/G	****
PROX1-rs340874	C/T	****
FADS1-rs174550	T/T	****
SLC2A2- rs11920090	T/T	****
ADCY5- rs11708067	A/A	****
CRY2-rs11605924	A/A	****
ADRA2A- rs10885122	G/G	****
MTNR1B- rs10830963	C/G	****

**END OF REPORT** 

SEX: MALE ACC #: C4215580

DATE:



**APPENDIX** 

PAGE 44



The scientific studies referenced in this report are provided below and can be referenced at www.pubmed.gov. All of these papers were published in peer-reviewed journals. PubMed is a service managed by the National Institutes of Health (NIH), a part of the U.S. Department of Health and Human Services, and it tracks more than 19 million citations for biomedical articles and scientific research.

- 1. O'Rahilly S et al. Human Obesity: A Heritable Neurobehavioral Disorder That Is Highly Sensitive To Environmental Conditions. *Diabetes* **57**, 2905-10 (2008).
- 2. Ordovas JM et al. Dietary Fat Intake Determines The Effect Of A Common Polymorphism In The Hepatic Lipase Gene Promoter On High-density Lipoprotein Metabolism: Evidence Of A Strong Dose Effect In This Gene-nutrient Interaction In The Framingham Study. *Circulation* **106**, 2315-21 (2002).
- 3. Junyent M et al. Novel Variants At KCTD10, MVK, And MMAB Genes Interact With Dietary Carbohydrates To Modulate HDL-cholesterol Concentrations In The Genetics Of Lipid Lowering Drugs And Diet Network Study. *The American Journal Of Clinical Nutrition* **90**, 686-94 (2009).
- 4. Sonestedt E et al. Fat And Carbohydrate Intake Modify The Association Between Genetic Variation In The FTO Genotype And Obesity. *The American Journal Of Clinical Nutrition* **90**, 1418-25 (2009).
- 5. Corella D et al. APOA2, Dietary Fat, And Body Mass Index: Replication Of A Gene-diet Interaction In 3 Independent Populations. *Archives Of Internal Medicine* **169**, 1897-906 (2009).
- 6. Warodomwichit D et al. ADIPOQ Polymorphisms, Monounsaturated Fatty Acids, And Obesity Risk: The GOLDN Study. *Obesity (Silver Spring, Md.)* **17**, 510-7 (2009).
- 7. Memisoglu A et al. Interaction Between A Peroxisome Proliferator-activated Receptor Gamma Gene Polymorphism And Dietary Fat Intake In Relation To Body Mass. *Human Molecular Genetics* **12**, 2923-9 (2003).
- 8. Kathiresan S et al. Common Variants At 30 Loci Contribute To Polygenic Dyslipidemia. Nature Genetics 41, 56-65 (2009).
- 9. Dupuis J et al. New Genetic Loci Implicated In Fasting Glucose Homeostasis And Their Impact On Type 2 Diabetes Risk. *Nature Genetics* **42**, 105-16 (2010).
- 10. Glaser C et al. Genetic Variation In Polyunsaturated Fatty Acid Metabolism And Its Potential Relevance For Human Development And Health. *Maternal & Child Nutrition* **7 Suppl 2**, 27-40 (2011).
- 11. Simopoulos AP. The Importance Of The Omega-6/omega-3 Fatty Acid Ratio In Cardiovascular Disease And Other Chronic Diseases. *Experimental Biology And Medicine (Maywood, N.J.)* **233**, 674-88 (2008).
- 12. Tanaka T et al. Genome-wide Association Study Of Plasma Polyunsaturated Fatty Acids In The InCHIANTI Study. *PLoS Genetics* **5**, e1000338 (2009).
- 13. Lemaitre RN et al. Genetic Loci Associated With Plasma Phospholipid N-3 Fatty Acids: A Meta-analysis Of Genome-wide Association Studies From The CHARGE Consortium. *PLoS Genetics* **7**, e1002193 (2011).
- 14. Epstein LH et al. Food Reinforcement, The Dopamine D2 Receptor Genotype, And Energy Intake In Obese And Nonobese Humans. *Behavioral Neuroscience* **121**, 877-86 (2007).
- 15. Doehring A et al. Genetic Diagnostics Of Functional Variants Of The Human Dopamine D2 Receptor Gene. *Psychiatric Genetics* **19**, 259-68 (2009).
- 16. Eny KM et al. Dopamine D2 Receptor Genotype (C957T) And Habitual Consumption Of Sugars In A Free-living Population Of Men And Women. *Journal Of Nutrigenetics And Nutrigenomics* **2**, 235-42 (2009).
- 17. de Krom M et al. Common Genetic Variations In CCK, Leptin, And Leptin Receptor Genes Are Associated With Specific Human Eating Patterns. *Diabetes* **56**, 276-80 (2007).

SEX: MALE ACC #: C4215580 DATE:



### **APPENDIX**



- 18. Bouchard L et al. Neuromedin Beta: A Strong Candidate Gene Linking Eating Behaviors And Susceptibility To Obesity. *The American Journal Of Clinical Nutrition* **80**, 1478-86 (2004).
- 19. Frayling TM et al. A Common Variant In The FTO Gene Is Associated With Body Mass Index And Predisposes To Childhood And Adult Obesity. *Science (New York, N.Y.)* **316**, 889-94 (2007).
- 20. Wardle J et al. Obesity Associated Genetic Variation In FTO Is Associated With Diminished Satiety. *The Journal Of Clinical Endocrinology And Metabolism* **93**, 3640-3 (2008).
- 21. den Hoed M et al. Postprandial Responses In Hunger And Satiety Are Associated With The Rs9939609 Single Nucleotide Polymorphism In FTO. *The American Journal Of Clinical Nutrition* **90**, 1426-32 (2009).
- 22. Dotson CD et al. Variation In The Gene TAS2R38 Is Associated With The Eating Behavior Disinhibition In Old Order Amish Women. *Appetite* **54**, 93-9 (2010).
- 23. Epstein LH et al. Food Reinforcement. *Appetite* 46, 22-5 (2006).
- 24. Eny KM et al. Genetic Variant In The Glucose Transporter Type 2 Is Associated With Higher Intakes Of Sugars In Two Distinct Populations. *Physiological Genomics* **33**, 355-60 (2008).
- 25. Cornelis MC et al. Coffee, Caffeine, And Coronary Heart Disease. *Current Opinion In Clinical Nutrition And Metabolic Care* **10**, 745-51 (2007).
- 26. Sachse C et al. Functional Significance Of A C-->A Polymorphism In Intron 1 Of The Cytochrome P450 CYP1A2 Gene Tested With Caffeine. *British Journal Of Clinical Pharmacology* **47**, 445-9 (1999).
- 27. Djordjevic N et al. Induction Of CYP1A2 By Heavy Coffee Consumption Is Associated With The CYP1A2 -163C>A Polymorphism. *European Journal Of Clinical Pharmacology* **66**, 697-703 (2010).
- 28. Gunes A et al. Variation In CYP1A2 Activity And Its Clinical Implications: Influence Of Environmental Factors And Genetic Polymorphisms. *Pharmacogenomics* **9**, 625-37 (2008).
- 29. Zhou SF et al. Structure, Function, Regulation And Polymorphism And The Clinical Significance Of Human Cytochrome P450 1A2. *Drug Metabolism Reviews* **42**, 268-354 (2010).
- 30. Kim UK et al. Positional Cloning Of The Human Quantitative Trait Locus Underlying Taste Sensitivity To Phenylthiocarbamide. *Science (New York, N.Y.)* **299**, 1221-5 (2003).
- 31. Reed DR et al. The Perception Of Quinine Taste Intensity Is Associated With Common Genetic Variants In A Bitter Receptor Cluster On Chromosome 12. *Human Molecular Genetics* **19**, 4278-85 (2010).
- 32. Hayes JE et al. Explaining Variability In Sodium Intake Through Oral Sensory Phenotype, Salt Sensation And Liking. *Physiology & Behavior* **100**, 369-80 (2010).
- 33. Fushan AA et al. Allelic Polymorphism Within The TAS1R3 Promoter Is Associated With Human Taste Sensitivity To Sucrose. *Current Biology: CB* **19**, 1288-93 (2009).
- 34. Fang L et al. The Human Lactase Persistence-associated SNP -13910\*T Enables In Vivo Functional Persistence Of Lactase Promoter-reporter Transgene Expression. *Human Genetics*, (2012).
- 35. Olds LC et al. Lactase Persistence DNA Variant Enhances Lactase Promoter Activity In Vitro: Functional Role As A Cis Regulatory Element. *Human Molecular Genetics* **12**, 2333-40 (2003).
- 36. Troelsen JT et al. An Upstream Polymorphism Associated With Lactase Persistence Has Increased Enhancer Activity. *Gastroenterology* **125**, 1686-94 (2003).
- 37. Enattah NS et al. Identification Of A Variant Associated With Adult-type Hypolactasia. Nature Genetics 30, 233-7 (2002).

SEX: MALE ACC #: C4215580

DATE:



## **APPENDIX**



- 38. Matsuo K et al. Alcohol Dehydrogenase 2 His47Arg Polymorphism Influences Drinking Habit Independently Of Aldehyde Dehydrogenase 2 Glu487Lys Polymorphism: Analysis Of 2,299 Japanese Subjects. *Cancer Epidemiology, Biomarkers & Prevention: A Publication Of The American Association For Cancer Research, Cosponsored By The American Society Of Preventive Oncology* **15**, 1009-13 (2006).
- 39. Tanaka F et al. Polymorphism Of Alcohol-metabolizing Genes Affects Drinking Behavior And Alcoholic Liver Disease In Japanese Men. *Alcoholism, Clinical And Experimental Research* **21**, 596-601 (1997).
- 40. Higuchi S et al. Influence Of Genetic Variations Of Ethanol-metabolizing Enzymes On Phenotypes Of Alcohol-related Disorders. *Annals Of The New York Academy Of Sciences* **1025**, 472-80 (2004).
- 41. Powers HJ. Riboflavin (vitamin B-2) And Health. The American Journal Of Clinical Nutrition 77, 1352-60 (2003).
- 42. McNulty H et al. Homocysteine, B-vitamins And CVD. The Proceedings Of The Nutrition Society 67, 232-7 (2008).
- 43. Hustad S et al. The Methylenetetrahydrofolate Reductase 677C-->T Polymorphism As A Modulator Of A B Vitamin Network With Major Effects On Homocysteine Metabolism. *American Journal Of Human Genetics* **80**, 846-55 (2007).
- 44. Yazdanpanah N et al. Low Dietary Riboflavin But Not Folate Predicts Increased Fracture Risk In Postmenopausal Women Homozygous For The MTHFR 677 T Allele. *Journal Of Bone And Mineral Research : The Official Journal Of The American Society For Bone And Mineral Research* 23, 86-94 (2008).
- 45. Horigan G et al. Riboflavin Lowers Blood Pressure In Cardiovascular Disease Patients Homozygous For The 677C-->T Polymorphism In MTHFR. *Journal Of Hypertension* **28**, 478-86 (2010).
- 46. McNulty H et al. Riboflavin Lowers Homocysteine In Individuals Homozygous For The MTHFR 677C->T Polymorphism. *Circulation* **113**, 74-80 (2006).
- 47. Tanaka T et al. Genome-wide Association Study Of Vitamin B6, Vitamin B12, Folate, And Homocysteine Blood Concentrations. *American Journal Of Human Genetics* **84**, 477-82 (2009).
- 48. Hazra A et al. Genome-wide Significant Predictors Of Metabolites In The One-carbon Metabolism Pathway. *Human Molecular Genetics* **18**, 4677-87 (2009).
- 49. Zittoun J et al. Modern Clinical Testing Strategies In Cobalamin And Folate Deficiency. *Seminars In Hematology* **36**, 35-46 (1999).
- 50. Hazra A et al. Common Variants Of FUT2 Are Associated With Plasma Vitamin B12 Levels. *Nature Genetics* **40**, 1160-2 (2008).
- 51. Bailey LB et al. Folate Metabolism And Requirements. The Journal Of Nutrition 129, 779-82 (1999).
- 52. Yang QH et al. Prevalence And Effects Of Gene-gene And Gene-nutrient Interactions On Serum Folate And Serum Total Homocysteine Concentrations In The United States: Findings From The Third National Health And Nutrition Examination Survey DNA Bank. *The American Journal Of Clinical Nutrition* **88**, 232-46 (2008).
- 53. Voutilainen S et al. Low Dietary Folate Intake Is Associated With An Excess Incidence Of Acute Coronary Events: The Kuopio Ischemic Heart Disease Risk Factor Study. *Circulation* **103**, 2674-80 (2001).
- 54. Gerster H. Vitamin A--functions, Dietary Requirements And Safety In Humans. *International Journal For Vitamin And Nutrition Research. Internationale Zeitschrift Fur Vitamin- Und Ernahrungsforschung. Journal International De Vitaminologie Et De Nutrition* 67, 71-90 (1997).
- 55. Semba RD. The Role Of Vitamin A And Related Retinoids In Immune Function. Nutrition Reviews 56, S38-48 (1998).
- 56. Dawson MI. The Importance Of Vitamin A In Nutrition. Current Pharmaceutical Design 6, 311-25 (2000).
- 57. Ross AC et al. The Function Of Vitamin A In Cellular Growth And Differentiation, And Its Roles During Pregnancy And Lactation. *Advances In Experimental Medicine And Biology* **352**, 187-200 (1994).

SEX: MALE ACC #: C4215580

DATE:



## **APPENDIX**



- 58. Leung WC et al. Two Common Single Nucleotide Polymorphisms In The Gene Encoding Beta-carotene 15,15'-monoxygenase Alter Beta-carotene Metabolism In Female Volunteers. *FASEB Journal : Official Publication Of The Federation Of American Societies For Experimental Biology* 23, 1041-53 (2009).
- 59. Witschi JC et al. Preformed Vitamin A, Carotene, And Total Vitamin A Activity In Usual Adult Diets. *Journal Of The American Dietetic Association* **57**, 13-6 (1970).
- 60. Solomons NW et al. Plant Sources Of Provitamin A And Human Nutriture. Nutrition Reviews 51, 199-204 (1993).
- 61. Cahill LE et al. Vitamin C Transporter Gene Polymorphisms, Dietary Vitamin C And Serum Ascorbic Acid. *Journal Of Nutrigenetics And Nutrigenomics* **2**, 292-301 (2009).
- 62. Timpson NJ et al. Genetic Variation At The SLC23A1 Locus Is Associated With Circulating Concentrations Of L-ascorbic Acid (vitamin C): Evidence From 5 Independent Studies With >15,000 Participants. *The American Journal Of Clinical Nutrition* **92**, 375-82 (2010).
- 63. Holick MF. Vitamin D And Bone Health. The Journal Of Nutrition 126, 1159S-64S (1996).
- 64. Ahn J et al. Vitamin D-related Genes, Serum Vitamin D Concentrations And Prostate Cancer Risk. *Carcinogenesis* **30**, 769-76 (2009).
- 65. Wang TJ et al. Common Genetic Determinants Of Vitamin D Insufficiency: A Genome-wide Association Study. *Lancet* **376**, 180-8 (2010).
- 66. Beharka A et al. Vitamin E Status And Immune Function. Methods In Enzymology 282, 247-63 (1997).
- 67. Morrissey PA et al. Optimal Nutrition: Vitamin E. The Proceedings Of The Nutrition Society 58, 459-68 (1999).
- 68. Ferrucci L et al. Common Variation In The Beta-carotene 15,15'-monooxygenase 1 Gene Affects Circulating Levels Of Carotenoids: A Genome-wide Association Study. *American Journal Of Human Genetics* **84**, 123-33 (2009).
- 69. Bartali B et al. Serum Micronutrient Concentrations And Decline In Physical Function Among Older Persons. *JAMA : The Journal Of The American Medical Association* **299**, 308-15 (2008).
- 70. Maras JE et al. Intake Of Alpha-tocopherol Is Limited Among US Adults. *Journal Of The American Dietetic Association* **104**, 567-75 (2004).
- 71. Garenc C et al. Evidence Of LPL Gene-exercise Interaction For Body Fat And LPL Activity: The HERITAGE Family Study. *Journal Of Applied Physiology (Bethesda, Md. : 1985)* **91**, 1334-40 (2001).
- 72. Teran-Garcia M et al. Hepatic Lipase Gene Variant -514C>T Is Associated With Lipoprotein And Insulin Sensitivity Response To Regular Exercise: The HERITAGE Family Study. *Diabetes* **54**, 2251-5 (2005).
- 73. Hautala AJ et al. Peroxisome Proliferator-activated Receptor-delta Polymorphisms Are Associated With Physical Performance And Plasma Lipids: The HERITAGE Family Study. *American Journal Of Physiology. Heart And Circulatory Physiology* **292**, H2498-505 (2007).
- 74. Orkunoglu-Suer FE et al. INSIG2 Gene Polymorphism Is Associated With Increased Subcutaneous Fat In Women And Poor Response To Resistance Training In Men. *BMC Medical Genetics* **9**, 117 (2008).
- 75. Saltin B et al. Maximal Oxygen Uptake In Athletes. Journal Of Applied Physiology 23, 353-8 (1967).
- 76. Lucia A et al. PPARGC1A Genotype (Gly482Ser) Predicts Exceptional Endurance Capacity In European Men. *Journal Of Applied Physiology (Bethesda, Md. : 1985)* **99**, 344-8 (2005).
- 77. Yang N et al. ACTN3 Genotype Is Associated With Human Elite Athletic Performance. *American Journal Of Human Genetics* **73**, 627-31 (2003).

SEX: MALE ACC #: C4215580 DATE:



**APPENDIX** 



- 78. Druzhevskaya AM et al. Association Of The ACTN3 R577X Polymorphism With Power Athlete Status In Russians. *European Journal Of Applied Physiology* **103**, 631-4 (2008).
- 79. Raleigh SM et al. Variants Within The MMP3 Gene Are Associated With Achilles Tendinopathy: Possible Interaction With The COL5A1 Gene. *British Journal Of Sports Medicine* **43**, 514-20 (2009).
- 80. Li S et al. Cumulative Effects And Predictive Value Of Common Obesity-susceptibility Variants Identified By Genome-wide Association Studies. *The American Journal Of Clinical Nutrition* **91**, 184-90 (2010).
- 81. Vimaleswaran KS et al. Physical Activity Attenuates The Body Mass Index-increasing Influence Of Genetic Variation In The FTO Gene. *The American Journal Of Clinical Nutrition* **90**, 425-8 (2009).
- 82. Rankinen T et al. Effect Of Endothelin 1 Genotype On Blood Pressure Is Dependent On Physical Activity Or Fitness Levels. *Hypertension* **50**, 1120-5 (2007).
- 83. Tao YX. The Melanocortin-4 Receptor: Physiology, Pharmacology, And Pathophysiology. *Endocrine Reviews* **31**, 506-43 (2010).
- 84. Fawcett KA et al. The Genetics Of Obesity: FTO Leads The Way. Trends In Genetics: TIG 26, 266-74 (2010).
- 85. Loos RJ et al. Common Variants Near MC4R Are Associated With Fat Mass, Weight And Risk Of Obesity. *Nature Genetics* **40**, 768-75 (2008).
- 86. Willer CJ et al. Six New Loci Associated With Body Mass Index Highlight A Neuronal Influence On Body Weight Regulation. *Nature Genetics* **41**, 25-34 (2009).
- 87. Meyre D et al. Genome-wide Association Study For Early-onset And Morbid Adult Obesity Identifies Three New Risk Loci In European Populations. *Nature Genetics* **41**, 157-9 (2009).
- 88. Cho YS et al. A Large-scale Genome-wide Association Study Of Asian Populations Uncovers Genetic Factors Influencing Eight Quantitative Traits. *Nature Genetics* **41**, 527-34 (2009).
- 89. Leskinen T et al. Leisure-time Physical Activity And High-risk Fat: A Longitudinal Population-based Twin Study. *International Journal Of Obesity (2005)* **33**, 1211-8 (2009).
- 90. Swinburn BA et al. Diet, Nutrition And The Prevention Of Excess Weight Gain And Obesity. *Public Health Nutrition* **7**, 123-46 (2004).
- 91. Goyenechea E et al. The 11391 G/A Polymorphism Of The Adiponectin Gene Promoter Is Associated With Metabolic Syndrome Traits And The Outcome Of An Energy-restricted Diet In Obese Subjects. *Hormone And Metabolic Research = Hormon-Und Stoffwechselforschung = Hormones Et Metabolisme* **41**, 55-61 (2009).
- 92. Loos RJ et al. Polymorphisms In The Leptin And Leptin Receptor Genes In Relation To Resting Metabolic Rate And Respiratory Quotient In The Québec Family Study. *International Journal Of Obesity (2005)* **30**, 183-90 (2006).
- 93. Puglisi MJ et al. Modulation Of C-reactive Protein, Tumor Necrosis Factor-alpha, And Adiponectin By Diet, Exercise, And Weight Loss. *The Journal Of Nutrition* **138**, 2293-6 (2008).
- 94. Qi Y et al. Adiponectin Acts In The Brain To Decrease Body Weight. Nature Medicine 10, 524-9 (2004).
- 95. Heid IM et al. Clear Detection Of ADIPOQ Locus As The Major Gene For Plasma Adiponectin: Results Of Genome-wide Association Analyses Including 4659 European Individuals. *Atherosclerosis* **208**, 412-20 (2010).
- 96. Natarajan P et al. High-density Lipoprotein And Coronary Heart Disease: Current And Future Therapies. *Journal Of The American College Of Cardiology* **55**, 1283-99 (2010).
- 97. Alwaili K et al. High-density Lipoproteins And Cardiovascular Disease: 2010 Update. *Expert Review Of Cardiovascular Therapy* **8**, 413-23 (2010).

SEX: MALE ACC #: C4215580 DATE:



## APPENDIX

PATHWAY GENOMICS

PAGE 49

98. Renström F et al. Genetic Predisposition To Long-term Nondiabetic Deteriorations In Glucose Homeostasis: Ten-year Follow-up Of The GLACIER Study. *Diabetes* **60**, 345-54 (2011).

SEX: MALE ACC #: C4215580

DATE:



**APPENDIX** 

PATHWAY GENOMICS

PAGE 50

## **Risks & Limitations**

## **Risks**

#### Risk of Laboratory Error

Pathway is a certified laboratory under the federal Clinical Laboratory Improvement Amendments of 1988 (CLIA) with standard and effective procedures in place for handling samples. However, laboratory error can occur, which might lead to incorrect results. Examples include, but are not limited to, a sample or DNA mislabeling or contamination, failure to obtain an interpretable report, and any other operational laboratory error. I understand that sometimes Pathway's laboratory may need a second sample to complete my testing.

## Risk of laboratory technical problems

Pathway's CLIA-certified laboratory also has standard and effective procedures in place to protect against technical and operational problems. However, such problems may still occur and examples include, but are not limited to, failure to obtain an interpretable result for a particular SNP. Sometimes it is not possible to obtain a testing result for a particular mutation or marker due to circumstances beyond Pathway's control, in which case it may not be possible for Pathway to conclusively report on a genetic change that might cause or be predictive of a condition. This may mean that Pathway cannot report my results for a particular health trait or condition, carrier status result, drug response, or other phenotype. Pathway may re-test my sample in order to obtain these results, but upon re-testing the results may still not be obtained. As with all medical laboratory testing, there is a small chance that the laboratory could report false positive or false negative results. A false positive result means that a genotype is reported as being present when it is actually not present. A false negative result means that a genotype is not reported as being present. A tested individual may wish to pursue further testing to verify any results.

### Limitations

The purpose of this test is to provide information about how a tested individual's genes affect their metabolism, weight, exercise, energy use, eating behavior, diet and nutritional choices. Tested individuals should not change their diet, physical activity, or any medical treatments they are currently using based on genetic testing results without consulting their personal health care provider.

Tested individuals may find that their experience is not consistent with Pathway's selected peer-reviewed scientific research findings of relative improvement for the study group(s). The science in this area is still developing and many personal health factors affect diet and health. Since subjects in the scientific studies referenced in this report may have had personal health and other factors different from those of tested individuals, results from these studies may not be representative of the results experienced by tested individuals. Further, some recommendations may or may not be attainable, depending on the tested individual's physical ability or other personal health factors. A limitation of this testing is that most scientific studies have been performed in Caucasian populations only. The interpretations and recommendations are done in the context of Caucasian studies, but the results may or may not be relevant to tested individuals of different or mixed ethnicities.

The association between genetic mutations and the information in your Pathway Fit® report is an active area of scientific research, and future scientific discoveries might alter our understanding of how this information is related to your diet, nutrition, and exercise.

Based on test results and other medical knowledge of the tested individual, health care providers might consider additional independent testing, or consult another health care provider or genetic counselor.

SEX: MALE ACC #: C4215580 DATE:



**APPENDIX** 

PATHWAY GENOMICS

PAGE 5

## **Change History**

There are three ways in which this genetic report might get modified. A description of those methods is below, followed by a table listing all modifications to this report.

Corrected	A report is annotated as "Corrected" if there was an error in genotypic data or the algorithms for interpretation of genotypic data that changes a patient's result from a previous report. When a correction is issued, we will notate it along with the date on the summary and the details pages, and communicate the correction to the ordering physician via phone or email. Details of the correction will be provided in the Appendix, along with the date that it occurred.
Updated	A report is annotated as "Updated" when or if we make a substantial modification to the descriptive content in one of our reports, which is typically done to improve clarity or precision. Some updates, such as grammatical corrections or typos, may not be annotated at all, and we will not always send out communications about an update. Details of the update will be provided in the Appendix, along with the date that it occurred.
New	A report is annotated as "New" if we add a new condition or significant piece of content to a report. When a report is amended, we will notate it along with the date on the summary and details page, though we may remove the amended annotation after 6 months from the point it was added. Details of the amendment will be provided in the Appendix, along with the date that it occurred.