

4685 Centennial Blvd. Colorado Springs, CO 80919

Telephone: 888-843-5832 TIN: 47-2642690 Lab Director: Leslie Douglas, PhD, HCLD(MD), ABB

Fax: 719-548-8220 CLIA#: 06D2019763

Patient: Sample, Sally (1/27/64) **Provider: Jane Doe, MD** ApoE Genotyping Test Sample Collected Sample Received Sample Tested Test Reported 01/04/2025 01/06/2025 01/01/2025 01/03/2025 Sample type: Super Floss – Full Mouth Test performed by: L. Douglas Test ID: 58893 Your results indicate an ApoE genotype of: ApoE 3/4

ApoE 2/2: Rarest ApoE genotype with the lowest risk for Alzheimer's disease. ApoE_{2/2} is most efficient at removing metals, including mercury from the body; but may clear dietary fat more slowly and therefore may be at greater risk for vascular disease, diabetes, and high cholesterol.

ApoE 2/3: A heterozygous hybrid of two alleles of the same gene. In this case, the associated risk for ApoE 2/2and ApoE 3/3 are combined and averaged.

ApoE 2/4: A heterozygous hybrid of two alleles of the same gene. In this case, the associated risk for ApoE 2/2 and ApoE 4/4 are combined and averaged.

ApoE 3/3: Most common ApoE genotype found in the human population, suggesting "average risk" of diabetes, neurodegenerative and cardiovascular disorders, including Alzheimer's disease, high cholesterol and stroke.

ApoE 3/4: A heterozygous hybrid of two alleles of the same gene. In this case, the associated risk for ApoE 3/3 and ApoE 4/4 are combined and averaged.

ApoE 4/4: Genotype most often in individuals with neurological, cerebral, and cardiovascular disease; the least efficient at removing mercury and heavy metals like barium, aluminum, lead, cadmium, arsenic, copper, and uranium from the body.

Interpretation of Results Disclaimer: DNA Connexions is not a clinical diagnostic laboratory and cannot provide a diagnosis for disease and/or subsequent treatment. These results are from DNA PCR testing, and indicate the presence of targeted human DNA to predict the relative genetic susceptibility to gluten intolerance and/or Celiac Disease. The verbiage is supplied as a courtesy to health care providers to aide in an overall assessment. This information alone should not be used to diagnose or treat a health problem or disease. All reported results are intended for research purposes only and consultation with a qualified health care provider is required.

General Information

Apolipoprotein E (ApoE) is a molecule composed of protein and fats. The ApoE gene creates the precursors to cholesterol molecules, known as chylomicrons, which are the microscopic particles of emulsified fat found in our blood and lymphatic tissues. Chylomicrons are formed during the digestion of fats and are essential for the normal formation and digestion of the fatty proteins that are often referred to as "HDL" and "LDL" (high-density lipoproteins and low-density lipoproteins). ApoE helps carry and metabolize cholesterol and fat; both major components of all cells in the human body; particularly within the nervous system.

The human ApoE gene contains 299 amino acids residues which are joined together by flexible "hinges". There are three common forms of the ApoE gene: ApoE₂, ApoE₃, and ApoE₄; each with differences at amino acid 112/299 and 158/299: (ApoE₂ Cysteine112, Cysteine158); (ApoE₃ Cysteine112, Arginine158); and (ApoE₄ Arginine112, Arginine158). These differences alter the gene's protein and lipid structure. The amino acid Cysteine removes toxins and heavy metals from the

central nervous system (CNS) more efficiently than the amino acid Arginine does.

Every person inherits two copies of the gene; creating one of six possible ApoE genotypes. These are ApoE_{2/2}, ApoE _{2/3}, ApoE _{2/4}, ApoE _{3/3}, ApoE _{3/4}, and ApoE _{4/4}. Defects in these variants have been linked to the body's ability to break down and remove cholesterol, triglycerides, mercury, and other heavy metals like barium, aluminum, lead, cadmium, arsenic, copper, and uranium.

The ApoE gene has recently been studied for its role in several abnormal processes which are not directly related to fat, triglycerides, and



Figure 2. Distribution of observed and expected genotype frequencies of the apolipoprotein E gene polymorphism.

cholesterol transport. These include amyotrophic lateral sclerosis (ALS); multiple sclerosis (MS); and other disorders including insomnia and depression. In 1993, ApoE₄ was identified as a genetic risk factor for Alzheimer's disease (AD), and may be associated with the accelerated development and progression of several other neurodegenerative diseases.

REFERENCES:

Carson B.L. et al. Toxicology and Biological Monitoring of Metals in Humans, Lewis Publications, Chelsea MI pp. 16-20 1986.

- Godfrey ME, Wojcik DP, Krone CA. Apolipoprotein E genotyping as a potential biomarker for mercury neurotoxicity. J Alzheimers Dis. 2003 Jun;5(3):189-95. PubMed PMID: 12897404.
- Main BF, Jones PJ, MacGillivray RT, Banfield DK. Apolipoprotein E genotyping using the polymerase chain reaction and allele-specific oligonucleotide primers. J Lipid Res. 1991 Jan;32(1):183-7. PubMed PMID: 2010690.
- Robeson RH, Siegel AM, Dunckley T. Genomic and Proteomic Biomarker Discovery in Neurological Disease. Biomark Insights. 2008 Feb 9;3:73-86. PubMed PMID: 19578496; PubMed Central PMCID: PMC2688365.

Scriver C.A. et al The Metabolic Basis of Inherited Disease, 6th ed. McGraw-Hill, New York NY, pp 2349-50 on PFK deficiency. 1989.

Suzuki T. et al eds, Advances in Mercury Toxicology, Plenum Press, New York, 1991.