Phosphoglycerate mutase deficiency

Phosphoglycerate mutase deficiency is a disorder that primarily affects muscles used for movement (skeletal muscles). Beginning in childhood or adolescence, affected individuals experience muscle aches or cramping following strenuous physical activity. Some people with this condition also have recurrent episodes of myoglobinuria. Myoglobinuria occurs when muscle tissue breaks down abnormally and releases a protein called myoglobin, which is processed by the kidneys and released in the urine. If untreated, myoglobinuria can lead to kidney failure.

In some cases of phosphoglycerate mutase deficiency, microscopic tube-shaped structures called tubular aggregates are seen in muscle fibers. It is unclear how tubular aggregates are associated with the signs and symptoms of the disorder.

Frequency

Phosphoglycerate mutase deficiency is a rare condition; about 15 affected people have been reported in the medical literature. Most affected individuals have been African American.

Causes

Phosphoglycerate mutase deficiency is caused by mutations in the PGAM2 gene. This gene provides instructions for making an enzyme called phosphoglycerate mutase, which is involved in a critical energy-producing process in cells known as glycolysis. During glycolysis, the simple sugar glucose is broken down to produce energy.

The version of phosphoglycerate mutase produced from the PGAM2 gene is found primarily in skeletal muscle cells. Mutations in the PGAM2 gene greatly reduce the activity of phosphoglycerate mutase, which disrupts energy production in these cells. This defect underlies the muscle cramping and myoglobinuria that occur after strenuous exercise in affected individuals.

Inheritance Pattern

This condition is inherited in an autosomal recessive pattern, which means both copies of the PGAM2 gene in each cell have mutations. The parents of an individual with an autosomal recessive condition each carry one copy of the mutated gene, but they typically do not show signs and symptoms of the condition. However, people who carry one altered copy of the PGAM2 gene may have some features of phosphoglycerate mutase deficiency, including episodes of exercise-induced muscle cramping and myoglobinuria.
Other Names for This Condition

- deficiency mutase phosphoglycerate
- glycogen storage disease X
- GSD X
- GSD10
- GSDX
- myopathy due to phosphoglycerate mutase deficiency
- PGAM deficiency
- PGAMM deficiency

Diagnosis & Management

Genetic Testing Information

- What is genetic testing?
  https://primer/testing/genetictesting
- Genetic Testing Registry: Glycogen storage disease type X

Research Studies from ClinicalTrials.gov

- ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22phosphoglycerate+mutase+deficiency%22

Additional Information & Resources

Health Information from MedlinePlus

- Health Topic: Carbohydrate Metabolism Disorders
  https://medlineplus.gov/carbohydratemetabolismdisorders.html
- Health Topic: Muscle Disorders
  https://medlineplus.gov/muscledisorders.html

Genetic and Rare Diseases Information Center

- Phosphoglycerate mutase deficiency
  https://rarediseases.info.nih.gov/diseases/9964/phosphoglycerate-mutase-deficiency
Educational Resources

- MalaCards: phosphoglycerate mutase deficiency
  https://www.malacards.org/card/phosphoglycerate_mutase_deficiency
- Neuromuscular Disease Center, Washington University
  https://neuromuscular.wustl.edu/msys/glycogen.html#pgm
- Orphanet: Glycogen storage disease due to phosphoglycerate mutase deficiency
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=97234

Patient Support and Advocacy Resources

- Metabolic Support UK
  https://www.metabolicsupportuk.org/
- Muscular Dystrophy Association
  https://www.mda.org/disease/metabolic-myopathies
- Resource list from the University of Kansas Medical Center: Metabolic Conditions
  http://www.kumc.edu/gec/support/metaboli.html

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28phosphoglycerate+mutase+deficiency%5BTIAB%5D%29+OR+%28glycogen+storage+disease+type+X%5BTIAB%5D%29+OR+%28%28PGAM%5BTIAB%5D%29+AND+%28deficiency%5BTIAB%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D

Catalog of Genes and Diseases from OMIM

- GLYCOGEN STORAGE DISEASE X
  http://omim.org/entry/261670

Sources for This Summary

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/8761269

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/8447317
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1682163/

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/7603528

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