Argininosuccinic aciduría

Argininosuccinic aciduría is an inherited disorder that causes ammonia to accumulate in the blood. Ammonia, which is formed when proteins are broken down in the body, is toxic if the levels become too high. The nervous system is especially sensitive to the effects of excess ammonia.

Argininosuccinic aciduría usually becomes evident in the first few days of life. An infant with argininosuccinic aciduría may be lacking in energy (lethargic) or unwilling to eat, and have poorly controlled breathing rate or body temperature. Some babies with this disorder experience seizures or unusual body movements, or go into a coma. Complications from argininosuccinic aciduría may include developmental delay and intellectual disability. Progressive liver damage, skin lesions, and brittle hair may also be seen.

Occasionally, an individual may inherit a mild form of the disorder in which ammonia accumulates in the bloodstream only during periods of illness or other stress.

Frequency

Argininosuccinic aciduría occurs in approximately 1 in 70,000 newborns.

Causes

Mutations in the ASL gene cause argininosuccinic aciduría.

Argininosuccinic aciduría belongs to a class of genetic diseases called urea cycle disorders. The urea cycle is a sequence of reactions that occur in liver cells. It processes excess nitrogen, generated when protein is used by the body, to make a compound called urea that is excreted by the kidneys.

In argininosuccinic aciduría, the enzyme that starts a specific reaction within the urea cycle is damaged or missing. The urea cycle cannot proceed normally, and nitrogen accumulates in the bloodstream in the form of ammonia.

Ammonia is especially damaging to the nervous system, so argininosuccinic aciduría causes neurological problems as well as eventual damage to the liver.

Inheritance Pattern

This condition is inherited in an autosomal recessive pattern, which means both copies of the 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.
Other Names for This Condition

- Argininosuccinate lyase deficiency
- argininosuccinic acidemia
- Argininosuccinicaciduria
- argininosuccinyl-CoA lyase deficiency
- arginosuccinase deficiency
- ASA
- ASAuria
- ASL deficiency

Diagnosis & Management

Formal Diagnostic Criteria
- ACT Sheet: Increased Citrulline
  https://www.ncbi.nlm.nih.gov/books/NBK55827/bin/Citrullinemia.pdf

Formal Treatment/Management Guidelines
- New England Consortium of Metabolic Programs: Acute Illness Protocol

Genetic Testing Information
- What is genetic testing?
  primer/testing/genetictesting
- Genetic Testing Registry: Argininosuccinate lyase deficiency

Research Studies from ClinicalTrials.gov
- ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22argininosuccinic+aciduria%22+OR+%22amino+acid+metabolism%2C+inborn+errors%22

Other Diagnosis and Management Resources
- Baby’s First Test
  https://www.babysfirsttest.org/newborn-screening/conditions/argininosuccinic-aciduria
- GeneReview: Argininosuccinate Lyase Deficiency
  https://www.ncbi.nlm.nih.gov/books/NBK51784
GeneReview: Urea Cycle Disorders Overview
https://www.ncbi.nlm.nih.gov/books/NBK1217

MedlinePlus Encyclopedia: Hereditary urea cycle abnormality
https://medlineplus.gov/ency/article/000372.htm

National Organization for Rare Disorders (NORD) Physician Guide: Urea Cycle Disorders
https://rarediseases.org/physician-guide/urea-cycle-disorders/

Additional Information & Resources

Health Information from MedlinePlus

- Encyclopedia: Hereditary urea cycle abnormality
  https://medlineplus.gov/ency/article/000372.htm
- Health Topic: Amino Acid Metabolism Disorders
  https://medlineplus.gov/aminoacidmetabolismdisorders.html
- Health Topic: Genetic Brain Disorders
  https://medlineplus.gov/geneticbraindisorders.html
- Health Topic: Newborn Screening
  https://medlineplus.gov/newbornscreening.html

Genetic and Rare Diseases Information Center

- Argininosuccinic aciduria
  https://rarediseases.info.nih.gov/diseases/5843/argininosuccinic-aciduria

Educational Resources

- MalaCards: argininosuccinic aciduria
  https://www.malacards.org/card/argininosuccinic_aciduria
- Orphanet: Argininosuccinic aciduria
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=23
- Screening, Technology and Research in Genetics
  http://www.newbornscreening.info/Parents/aminoaciddisorders/ASAL.html
- Vanderbilt Children’s Hospital
- Virginia Department of Health
Patient Support and Advocacy Resources

- Metabolic Support UK
  https://www.metabolicsupportuk.org/
- National Organization for Rare Disorders (NORD)
  https://rarediseases.org/rare-diseases/argininosuccinic-aciduria/
- National Urea Cycle Disorders Foundation
  http://www.nucdf.org/
- Urea Cycle Disorders Consortium
  https://www.rarediseasesnetwork.org/cms/ucdc/Learn-More/Disorder-Definitions

Clinical Information from GeneReviews

- Argininosuccinate Lyase Deficiency
  https://www.ncbi.nlm.nih.gov/books/NBK51784
- Urea Cycle Disorders Overview
  https://www.ncbi.nlm.nih.gov/books/NBK1217

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28argininosuccinic+aciduria%5BTIAB%5D%29+OR+%28argininosuccinate+lyase+deficiency%5BTIAB%5D%29+OR+%28asl+deficiency%5BTIAB%5D%29%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+2160+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- ARGININOSUCCINIC ACIDURIA
  http://omim.org/entry/207900

Sources for This Summary

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  https://rarediseases.org/rare-diseases/argininosuccinic-aciduria/

• National Urea Cycle Disorders Foundation
  http://www.nucdf.org/

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