Mitochondrial encephalomyopathy, lactic acidosis, and stroke-like episodes

Mitochondrial encephalomyopathy, lactic acidosis, and stroke-like episodes (MELAS) is a condition that affects many of the body’s systems, particularly the brain and nervous system (encephalo-) and muscles (myopathy). The signs and symptoms of this disorder most often appear in childhood following a period of normal development, although they can begin at any age. Early symptoms may include muscle weakness and pain, recurrent headaches, loss of appetite, vomiting, and seizures. Most affected individuals experience stroke-like episodes beginning before age 40. These episodes often involve temporary muscle weakness on one side of the body (hemiparesis), altered consciousness, vision abnormalities, seizures, and severe headaches resembling migraines. Repeated stroke-like episodes can progressively damage the brain, leading to vision loss, problems with movement, and a loss of intellectual function (dementia).

Most people with MELAS have a buildup of lactic acid in their bodies, a condition called lactic acidosis. Increased acidity in the blood can lead to vomiting, abdominal pain, extreme tiredness (fatigue), muscle weakness, and difficulty breathing. Less commonly, people with MELAS may experience involuntary muscle spasms (myoclonus), impaired muscle coordination (ataxia), hearing loss, heart and kidney problems, diabetes, and hormonal imbalances.

Frequency

The exact incidence of MELAS is unknown. It is one of the more common conditions in a group known as mitochondrial diseases. Together, mitochondrial diseases occur in about 1 in 4,000 people.

Genetic Changes

MELAS can result from mutations in one of several genes, including $MT-ND1$, $MT-ND5$, $MT-TH$, $MT-TL1$, and $MT-TV$. These genes are found in the DNA of cellular structures called mitochondria, which convert the energy from food into a form that cells can use. Although most DNA is packaged in chromosomes within the nucleus, mitochondria also have a small amount of their own DNA, known as mitochondrial DNA or mtDNA.

Some of the genes related to MELAS provide instructions for making proteins involved in normal mitochondrial function. These proteins are part of a large enzyme complex in mitochondria that helps convert oxygen, fats, and simple sugars to energy. Other genes associated with this disorder provide instructions for making molecules called transfer RNAs (tRNAs), which are chemical cousins of DNA. These molecules help assemble protein building blocks called amino acids into full-length, functioning proteins within mitochondria.
Mutations in a particular transfer RNA gene, \( MT-TL1 \), cause more than 80 percent of all cases of MELAS. These mutations impair the ability of mitochondria to make proteins, use oxygen, and produce energy. Researchers have not determined how changes in mtDNA lead to the specific signs and symptoms of MELAS. They continue to investigate the effects of mitochondrial gene mutations in different tissues, particularly in the brain.

**Inheritance Pattern**

This condition is inherited in a mitochondrial pattern, which is also known as maternal inheritance. This pattern of inheritance applies to genes contained in mtDNA. Because egg cells, but not sperm cells, contribute mitochondria to the developing embryo, children can only inherit disorders resulting from mtDNA mutations from their mother. These disorders can appear in every generation of a family and can affect both males and females, but fathers do not pass traits associated with changes in mtDNA to their children.

In most cases, people with MELAS inherit an altered mitochondrial gene from their mother. Less commonly, the disorder results from a new mutation in a mitochondrial gene and occurs in people with no family history of MELAS.

**Other Names for This Condition**

- MELAS
- MELAS syndrome
- mitochondrial myopathy, encephalopathy, lactic acidosis, and stroke-like episodes
- mitochondrial myopathy, lactic acidosis, stroke-like episode
- myopathy, mitochondrial-encephalopathy-lactic acidosis-stroke

**Diagnosis & Management**

**Genetic Testing**

- Genetic Testing Registry: Juvenile myopathy, encephalopathy, lactic acidosis AND stroke  

**Other Diagnosis and Management Resources**

- GeneReview: MELAS  
- GeneReview: Mitochondrial Disorders Overview  
- MedlinePlus Encyclopedia: Lactic acidosis  
  [https://medlineplus.gov/ency/article/000391.htm](https://medlineplus.gov/ency/article/000391.htm)
• MedlinePlus Encyclopedia: Stroke
   https://medlineplus.gov/ency/article/000726.htm

• National Organization for Rare Disorders (NORD) Physician Guide: Mitochondrial Myopathies
   https://rarediseases.org/physician-guide/mitochondrial-myopathy/

General Information from MedlinePlus

• Diagnostic Tests
   https://medlineplus.gov/diagnostictests.html

• Drug Therapy
   https://medlineplus.gov/drugtherapy.html

• Genetic Counseling
   https://medlineplus.gov/geneticcounseling.html

• Palliative Care
   https://medlineplus.gov/palliativecare.html

• Surgery and Rehabilitation
   https://medlineplus.gov/surgeryandrehabilitation.html

Additional Information & Resources

MedlinePlus

• Encyclopedia: Lactic acidosis
   https://medlineplus.gov/ency/article/000391.htm

• Encyclopedia: Stroke
   https://medlineplus.gov/ency/article/000726.htm

• Health Topic: Genetic Brain Disorders
   https://medlineplus.gov/geneticbraindisorders.html

• Health Topic: Mitochondrial Diseases
   https://medlineplus.gov/mitochondrialdiseases.html

• Health Topic: Neuromuscular Disorders
   https://medlineplus.gov/neuromusculardisorders.html

• Health Topic: Stroke
   https://medlineplus.gov/stroke.html

Genetic and Rare Diseases Information Center

• Mitochondrial encephalomyopathy lactic acidosis and stroke-like episodes
Additional NIH Resources

• National Institute of Neurological Disorders and Stroke
  https://www.ninds.nih.gov/Disorders/All-Disorders/Mitochondrial-myopathy-
  Information-Page

Educational Resources

• Disease InfoSearch: Mitochondrial encephalomyopathy lactic acidosis and stroke-
  like episodes
  http://www.diseaseinfosearch.org/Mitochondrial+encephalomyopathy+lactic
  +acidosis+and+stroke-like+episodes/4831

• Kennedy Krieger Institute
  https://www.kennedykrieger.org/patient-care/diagnoses-disorders/mitochondrial-
  disorders

• MalaCards: mitochondrial myopathy, encephalopathy, lactic acidosis, and stroke-
  like episodes
  http://www.malacards.org/card/mitochondrial_myopathy_encephalopathy_la
  ctic_acidosis_and_stroke_like_episodes

• Neuromuscular Disease Center, Washington University
  https://neuromuscular.wustl.edu/mitosyn.html#melas

• Orphanet: MELAS
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=550

Patient Support and Advocacy Resources

• Children Living with Inherited Metabolic Diseases (CLIMB) (UK)
  http://www.climb.org.uk/

• Children’s Mitochondrial Disease Network (UK)
  http://www.cmdn.org.uk/

• MitoAction
  http://www.mitoaction.org

• Muscular Dystrophy Association: Facts About Mitochondrial Myopathies

• National Organization for Rare Disorders (NORD)
  https://rarediseases.org/rare-diseases/melas-syndrome/

• Resource List from the University of Kansas Medical Center
  http://www.kumc.edu/gec/support/mitochon.html

• United Mitochondrial Disease Foundation
  http://www.umdf.org/
GeneReviews
- MELAS
  https://www.ncbi.nlm.nih.gov/books/NBK1233
- Mitochondrial Disorders Overview
  https://www.ncbi.nlm.nih.gov/books/NBK1224

ClinicalTrials.gov
- ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22mitochondrial+encephalomyopathy%2C+lactic+acidosis+and+stroke-like+episodes%22+OR+%22MELAS+Syndrome%22

Scientific Articles on PubMed
- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28mitochondrial+encephalomyopathy,+lactic+acidosis,+and+stroke-like+episodes%29+OR+%28MELAS+syndrome%5BMAJR%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+720+days%22%5Bdp%5D

OMIM
- MITOCHONDRIAL MYOPATHY, ENCEPHALOPATHY, LACTIC ACIDOSIS, AND STROKE-LIKE EPISODES
  http://omim.org/entry/540000

Sources for This Summary
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20301411
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/22328278
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/16400302

page 5


Reprinted from Genetics Home Reference:

Reviewed: December 2013
Published: July 24, 2018

Lister Hill National Center for Biomedical Communications
U.S. National Library of Medicine
National Institutes of Health
Department of Health & Human Services