Early-onset primary dystonia

Early-onset primary dystonia is a condition characterized by progressive problems with movement, typically beginning in childhood. Dystonia is a movement disorder that involves involuntary tensing of the muscles (muscle contractions), twisting of specific body parts such as an arm or a leg, rhythmic shaking (tremors), and other uncontrolled movements. A primary dystonia is one that occurs without other neurological symptoms, such as seizures or a loss of intellectual function (dementia). Early-onset primary dystonia does not affect a person's intelligence.

On average, the signs and symptoms of early-onset primary dystonia appear around age 12. Abnormal muscle spasms in an arm or a leg are usually the first sign. These unusual movements initially occur while a person is doing a specific action, such as writing or walking. In some affected people, dystonia later spreads to other parts of the body and may occur at rest. The abnormal movements persist throughout life, but they do not usually cause pain.

The signs and symptoms of early-onset primary dystonia vary from person to person, even among affected members of the same family. The mildest cases affect only a single part of the body, causing isolated problems such as a writer's cramp in the hand. Severe cases involve abnormal movements affecting many regions of the body.

Frequency

Early-onset primary dystonia is among the most common forms of childhood dystonia. This disorder occurs most frequently in people of Ashkenazi (central and eastern European) Jewish heritage, affecting 1 in 3,000 to 9,000 people in this population. The condition is less common among people with other backgrounds; it is estimated to affect 1 in 10,000 to 30,000 non-Jewish people worldwide.

Causes

A particular mutation in the TOR1A gene (also known as DYT1) is responsible for most cases of early-onset primary dystonia. The TOR1A gene provides instructions for making a protein called torsinA. Although little is known about its function, this protein may help process and transport other proteins within cells. It appears to be critical for the normal development and function of nerve cells in the brain.

A mutation in the TOR1A gene alters the structure of torsinA. The altered protein's effect on the function of nerve cells in the brain is unclear. People with early-onset primary dystonia do not have a loss of nerve cells or obvious changes in the structure of the brain that would explain the abnormal muscle contractions. Instead, the altered torsinA protein may have subtle effects on the connections between nerve cells and likely disrupts chemical signaling between nerve cells that control movement.
Researchers are working to determine how a change in this protein leads to the characteristic features of this disorder.

**Inheritance Pattern**

Mutations in the \( \text{TOR1A} \) gene are inherited in an autosomal dominant pattern, which means one of the two copies of the gene is altered in each cell. Many people who have a mutation in this gene are not affected by the disorder and may never know they have the mutation. Only 30 to 40 percent of people who inherit a \( \text{TOR1A} \) mutation will ever develop signs and symptoms of early-onset primary dystonia.

Everyone who has been diagnosed with early-onset primary dystonia has inherited a \( \text{TOR1A} \) mutation from one parent. The parent may or may not have signs and symptoms of the condition, and other family members may or may not be affected.

**Other Names for This Condition**

- Dystonia musculorum deformans 1
- DYT1
- Early-onset generalized torsion dystonia
- Oppenheim dystonia
- Oppenheim's dystonia
- Primary torsion dystonia

**Diagnosis & Management**

**Genetic Testing Information**

- What is genetic testing? [link]
- Genetic Testing Registry: Dystonia 1 [link]

**Research Studies from ClinicalTrials.gov**

- ClinicalTrials.gov [link]

**Other Diagnosis and Management Resources**

- GeneReview: DYT1 Early-Onset Isolated Dystonia [link]
- MedlinePlus Encyclopedia: Movement - uncontrolled or slow [link]
Additional Information & Resources

Health Information from MedlinePlus

• Encyclopedia: Movement - uncontrolled or slow
  https://medlineplus.gov/ency/article/003197.htm

• Health Topic: Dystonia
  https://medlineplus.gov/dystonia.html

Genetic and Rare Diseases Information Center

• DYT-TOR1A
  https://rarediseases.info.nih.gov/diseases/2027/dyt-tor1a

Additional NIH Resources

• National Institute for Neurological Disorders and Stroke: Dystonia Fact Sheet
  https://www.ninds.nih.gov/Disorders/All-Disorders/Dystonias-Information-Page

Educational Resources

• MalaCards: dyt1 early-onset isolated dystonia
  https://www.malacards.org/card/dyt1_early_onset_isolated_dystonia

• Merck Manual Home Edition for Patients and Caregivers

• Orphanet: Early-onset generalized limb-onset dystonia
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=256

• The Bachmann-Strauss Dystonia & Parkinson Foundation: Dystonia Questions and Answers
  http://www.dystonia-parkinson.org/sites/default/files/Dystonia_Medical_Brochure_FINAL_LOW_RES.pdf

Patient Support and Advocacy Resources

• Dystonia Medical Research Foundation
  https://dystonia-foundation.org/what-is-dystonia/types-dystonia/generalized/

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

• The Norton & Elaine Sarnoff Center for Jewish Genetics
  https://www.juf.org/

Clinical Information from GeneReviews

• DYT1 Early-Onset Isolated Dystonia
  https://www.ncbi.nlm.nih.gov/books/NBK1492
Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28Dystonic+Disorders%5BMAJR%29%5D%29+AND+%28early-onset+primary+dystonia%5BMTJ%5D%29+AND+%28early-onset+torsion+dystonia%5BMTJ%5D%29+AND+oppenheim’s+dystonia%5BMTJ%5D%29+AND+english%5BLa%5D+AND+human%5Bmh%29%5D+AND+%22last+3600+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- DYSTONIA 1, TORSION, AUTOSOMAL DOMINANT
  http://omim.org/entry/128100

Sources for This Summary


  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1693547/


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