



Pendred syndrome

Pendred syndrome is a disorder typically associated with hearing loss and a thyroid condition called a goiter. A goiter is an enlargement of the thyroid gland, which is a butterfly-shaped organ at the base of the neck that produces hormones. If a goiter develops in a person with Pendred syndrome, it usually forms between late childhood and early adulthood. In most cases, this enlargement does not cause the thyroid to malfunction.

In most people with Pendred syndrome, severe to profound hearing loss caused by changes in the inner ear (sensorineural hearing loss) is evident at birth. Less commonly, hearing loss does not develop until later in infancy or early childhood. Some affected individuals also have problems with balance caused by dysfunction of the vestibular system, which is the part of the inner ear that helps maintain the body's balance and orientation.

An inner ear abnormality called an enlarged vestibular aqueduct (EVA) is a characteristic feature of Pendred syndrome. The vestibular aqueduct is a bony canal that connects the inner ear with the inside of the skull. Some affected individuals also have an abnormally shaped cochlea, which is a snail-shaped structure in the inner ear that helps process sound. The combination of an enlarged vestibular aqueduct and an abnormally shaped cochlea is known as Mondini malformation.

Pendred syndrome shares features with other hearing loss and thyroid conditions, and it is unclear whether they are best considered as separate disorders or as a spectrum of related signs and symptoms. These conditions include a form of nonsyndromic hearing loss (hearing loss that does not affect other parts of the body) called DFNB4, and, in a small number of people, a form of congenital hypothyroidism resulting from an abnormally small thyroid gland (thyroid hypoplasia). All of these conditions are caused by mutations in the same gene.

Frequency

The prevalence of Pendred syndrome is unknown. However, researchers estimate that it accounts for 7 to 8 percent of all hearing loss that is present from birth (congenital hearing loss).

Causes

Mutations in the *SLC26A4* gene cause about half of all cases of Pendred syndrome. The *SLC26A4* gene provides instructions for making a protein called pendrin. This protein transports negatively charged particles (ions), including chloride, iodide, and bicarbonate, into and out of cells. Although the function of pendrin is not fully understood, this protein is important for maintaining the proper levels of ions in the

thyroid and the inner ear. Mutations in the *SLC26A4* gene alter the structure or function of pendrin, which disrupts ion transport. An imbalance of particular ions disrupts the development and function of the thyroid gland and structures in the inner ear, which leads to the characteristic features of Pendred syndrome.

In people with Pendred syndrome who do not have mutations in the *SLC26A4* gene, the cause of the condition is unknown. Researchers suspect that other genetic and environmental factors may influence the condition.

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

- autosomal recessive sensorineural hearing impairment, enlarged vestibular aqueduct, and goiter
- deafness with goiter
- goiter-deafness syndrome
- Pendred's syndrome

Diagnosis & Management

Formal Diagnostic Criteria

- ACT Sheet: Congenital hearing loss >~30db
https://www.ncbi.nlm.nih.gov/books/NBK55827/bin/Hearing_Loss.pdf

Genetic Testing Information

- What is genetic testing?
[/primer/testing/geneticTesting](#)
- Genetic Testing Registry: Pendred syndrome
<https://www.ncbi.nlm.nih.gov/gtr/conditions/C0271829/>

Other Diagnosis and Management Resources

- Children's Hospital of Philadelphia, Center for Childhood Communication
<https://www.chop.edu/centers-programs/center-childhood-communication>
- GeneReview: Pendred Syndrome/Nonsyndromic Enlarged Vestibular Aqueduct
<https://www.ncbi.nlm.nih.gov/books/NBK1467>

- MedlinePlus Encyclopedia: Goiter
<https://medlineplus.gov/ency/article/001178.htm>
- MedlinePlus Encyclopedia: Hearing Loss
<https://medlineplus.gov/ency/article/003044.htm>

Additional Information & Resources

Health Information from MedlinePlus

- Encyclopedia: Goiter
<https://medlineplus.gov/ency/article/001178.htm>
- Encyclopedia: Hearing Loss
<https://medlineplus.gov/ency/article/003044.htm>
- Health Topic: Hearing Disorders and Deafness
<https://medlineplus.gov/hearingdisordersanddeafness.html>
- Health Topic: Thyroid Diseases
<https://medlineplus.gov/thyroiddiseases.html>

Genetic and Rare Diseases Information Center

- Pendred syndrome
<https://rarediseases.info.nih.gov/diseases/4271/pendred-syndrome>

Additional NIH Resources

- National Institute on Deafness and Other Communication Disorders: Enlarged Vestibular Aqueducts and Childhood Hearing Loss
<https://www.nidcd.nih.gov/health/enlarged-vestibular-aqueducts-and-childhood-hearing-loss>
- National Institute on Deafness and Other Communication Disorders: Genetics of Enlarged Vestibular Aqueducts (EVA)
<https://www.nidcd.nih.gov/health/genetics-enlarged-vestibular-aqueducts-eva>
- National Institute on Deafness and Other Communication Disorders: Pendred Syndrome
<https://www.nidcd.nih.gov/health/pendred-syndrome>

Educational Resources

- American Speech-Language-Hearing Association: Large Vestibular Aqueduct (LVA) Disorders
<https://www.asha.org/content.aspx?id=8589967424>
- American Thyroid Association: Goiter
<http://www.thyroid.org/goiter/>
- Laurent Clerc National Deaf Education Center, Gallaudet University
<https://www3.gallaudet.edu/clerc-center.html>

- MalaCards: pendred syndrome
https://www.malacards.org/card/pendred_syndrome
- Orphanet: Pendred syndrome
https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=705

Patient Support and Advocacy Resources

- American Society for Deaf Children
<http://deafchildren.org/>
- Contact a Family (UK)
<https://contact.org.uk/medical-information/conditions/p/pendred-syndrome/>
- National Association of the Deaf
<https://www.nad.org/>
- Resource List from the University of Kansas Medical Center
<http://www.kumc.edu/gec/support/hearing.html>

Clinical Information from GeneReviews

- Pendred Syndrome/Nonsyndromic Enlarged Vestibular Aqueduct
<https://www.ncbi.nlm.nih.gov/books/NBK1467>

Scientific Articles on PubMed

- PubMed
<https://www.ncbi.nlm.nih.gov/pubmed?term=%28pendred+syndrome%5BTIAB%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+1800+days%22%5Bdp%5D>

Catalog of Genes and Diseases from OMIM

- PENDRED SYNDROME
<http://omim.org/entry/274600>

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Reprinted from Genetics Home Reference:

<https://ghr.nlm.nih.gov/condition/pendred-syndrome>

Reviewed: March 2016

Published: May 14, 2019

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