Van der Woude syndrome

Van der Woude syndrome is a condition that affects the development of the face. Many people with this disorder are born with a cleft lip, a cleft palate (an opening in the roof of the mouth), or both. Affected individuals usually have depressions (pits) near the center of the lower lip, which may appear moist due to the presence of salivary and mucous glands in the pits. Small mounds of tissue on the lower lip may also occur. In some cases, people with van der Woude syndrome have missing teeth.

People with van der Woude syndrome who have cleft lip and/or palate, like other individuals with these facial conditions, have an increased risk of delayed language development, learning disabilities, or other mild cognitive problems. The average IQ of individuals with van der Woude syndrome is not significantly different from that of the general population.

Frequency

Van der Woude syndrome is believed to occur in 1 in 35,000 to 1 in 100,000 people, based on data from Europe and Asia. Van der Woude syndrome is the most common cause of cleft lip and palate resulting from variations in a single gene, and this condition accounts for approximately 1 in 50 such cases.

Causes

Mutations in the \textit{IRF6} gene cause van der Woude syndrome. The \textit{IRF6} gene provides instructions for making a protein that plays an important role in early development. This protein is a transcription factor, which means that it attaches (binds) to specific regions of DNA and helps control the activity of particular genes.

The IRF6 protein is active in cells that give rise to tissues in the head and face. It is also involved in the development of other parts of the body, including the skin and genitals.

Mutations in the \textit{IRF6} gene that cause van der Woude syndrome prevent one copy of the gene in each cell from making any functional protein. A shortage of the IRF6 protein affects the development and maturation of tissues in the face, resulting in the signs and symptoms of van der Woude syndrome.

Inheritance Pattern

This condition is inherited in an autosomal dominant pattern, which means one copy of the altered gene in each cell is sufficient to cause the disorder. In most cases, an affected person has one parent with the condition. Occasionally, an individual who has a copy of the altered gene does not show any signs or symptoms of the disorder.
Other Names for This Condition
• cleft lip and/or palate with mucous cysts of lower lip
• lip-pit syndrome
• VDWS
• VWS

Diagnosis & Management
Genetic Testing Information
• What is genetic testing?
  /primer/testing/genetictesting
Genetic Testing Registry: Van der Woude syndrome

Other Diagnosis and Management Resources
• GeneReview: IRF6-Related Disorders
  https://www.ncbi.nlm.nih.gov/books/NBK1407

Additional Information & Resources
Health Information from MedlinePlus
• Health Topic: Cleft Lip and Palate
  https://medlineplus.gov/cleftlipandpalate.html

Genetic and Rare Diseases Information Center
• Van der Woude syndrome

Educational Resources
• Malacards: van der woude syndrome 1
  https://www.malacards.org/card/van_der_woude_syndrome_1
• March of Dimes: Cleft Lip and Cleft Palate
• Orphanet: Van der Woude syndrome
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=888
• Seattle Children's Hospital
  https://www.seattlechildrens.org/conditions/chromosomal-genetic-conditions/van-der-woude
Patient Support and Advocacy Resources

- AboutFace International
  https://www.aboutface.ca/
- American Cleft Palate-Craniofacial Association
  https://cleftline.org/
- Children's Craniofacial Association
  https://ccakids.org/

Clinical Information from GeneReviews

- IRF6-Related Disorders
  https://www.ncbi.nlm.nih.gov/books/NBK1407

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28van+der+woude+syndrome%5B
  TIA%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last
  +1080+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- VAN DER WOUDE SYNDROME 1
  http://omim.org/entry/119300

Medical Genetics Database from MedGen

- Van der Woude syndrome

Sources for This Summary

  families with van der Woude and/or popliteal pterygium syndrome: all with a mutation in the IRF6
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/14757865
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1735675/

- Kondo S, Schutte BC, Richardson RJ, Bjork BC, Knight AS, Watanabe Y, Howard E, de Lima RL,
  HH, Lidral AC, Pober BR, Moreno L, Arcos-Burgos M, Valencia C, Houdayer C, Bahau M, Moretti-
  Ferreira D, Richieri-Costa A, Dixon MJ, Murray JC. Mutations in IRF6 cause Van der Woude and
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/12219090
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3169431/

- Nopoulos P, Richman L, Andreadsen N, Murray JC, Schutte B. Cognitive dysfunction in adults with
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/17438385
  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/17539900

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/14994878

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20301581

• OMIM: VAN DER WOUDE SYNDROME 1 
  http://omim.org/entry/119300

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