F8 gene
coaqulation factor VIII

Normal Function

The F8 gene provides instructions for making a protein called coagulation factor VIII. Coagulation factors are a group of related proteins that are essential for the formation of blood clots. After an injury, clots protect the body by sealing off damaged blood vessels and preventing further blood loss.

Coagulation factor VIII is made chiefly by cells in the liver. This protein circulates in the bloodstream in an inactive form, bound to another molecule called von Willebrand factor, until an injury that damages blood vessels occurs. In response to injury, coagulation factor VIII is activated and separates from von Willebrand factor. The active protein (sometimes written as coagulation factor VIIIa) interacts with another coagulation factor called factor IX. This interaction sets off a chain of additional chemical reactions that form a blood clot.

Health Conditions Related to Genetic Changes

Hemophilia

Mutations in the F8 gene cause hemophilia A, the most common form of this bleeding disorder. More than 1,300 alterations in this gene have been identified. Some of these mutations change single DNA building blocks (base pairs) in the gene, while others delete or insert multiple base pairs. The most common mutation in people with severe hemophilia A is a rearrangement of genetic material called an inversion. This inversion involves a large segment of the F8 gene.

Mutations in the F8 gene lead to the production of an abnormal version of coagulation factor VIII or reduce the amount of this protein. The altered or missing protein cannot participate effectively in the blood clotting process. As a result, blood clots cannot form properly in response to injury. These problems with blood clotting lead to excessive bleeding that can be difficult to control. Some mutations, such as the large inversion described above, almost completely eliminate the activity of coagulation factor VIII and result in severe hemophilia. Other mutations reduce but do not eliminate the protein’s activity, resulting in mild or moderate hemophilia.
Chromosomal Location

Cytogenetic Location: Xq28, which is the long (q) arm of the X chromosome at position 28

Molecular Location: base pairs 154,835,788 to 155,022,723 on the X chromosome (Homo sapiens Annotation Release 109, GRCh38.p12) (NCBI)

Credit: Genome Decoration Page/NCBI

Other Names for This Gene

• AHF
• antihemophilic factor
• coagulation factor VIII, procoagulant component
• coagulation factor VIII, procoagulant component (hemophilia A)
• DXS1253E
• FA8_HUMAN
• Factor VIII F8B
• FVIII
• HEMA
• procoagulant component

Additional Information & Resources

Educational Resources
• Biochemistry (fifth edition, 2002): Diagram of the blood clotting cascade
  https://www.ncbi.nlm.nih.gov/books/NBK22589/?rendertype=figure&id=A1401

Clinical Information from GeneReviews
• Hemophilia A
  https://www.ncbi.nlm.nih.gov/books/NBK1404
Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28F8%5BTIAB%5D%29+OR+%28coagulation+factor+VIII%5BTI%5D%29+OR+%28factor+VIII%5BMAJR%5D%29%29+AND+%28%28Genes%5BMH%5D%29+OR+%28Genetic+Phenomena%5BMH%5D%29%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+720+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- COAGULATION FACTOR VIII
  http://omim.org/entry/300841

Research Resources

- Atlas of Genetics and Cytogenetics in Oncology and Haematology
  http://atlasgeneticsoncology.org/Genes/GC_F8.html

- ClinVar
  https://www.ncbi.nlm.nih.gov/clinvar?term=F8%5Bgene%5D

- HGNC Gene Symbol Report

- Monarch Initiative
  https://monarchinitiative.org/gene/NCBIGene:2157

- NCBI Gene

- UniProt
  https://www.uniprot.org/uniprot/P00451

Sources for This Summary


  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/11950963 
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1187163/

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/15931172

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Reprinted from Genetics Home Reference: 

Reviewed: May 2010
Published: May 28, 2019

Lister Hill National Center for Biomedical Communications
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