CSTB gene
cystatin B

**Normal Function**

The *CSTB* gene provides instructions for making a protein called cystatin B. This protein reduces the activity of (inhibits) enzymes called cathepsins. Cathepsins help break down certain proteins in the lysosomes (compartments in the cell that digest and recycle materials). While the specific function of cystatin B is unclear, it may help protect the cells' proteins from cathepsins that leak out of the lysosomes.

One region of the *CSTB* gene has a particular repeating sequence of 12 DNA building blocks (nucleotides) written as CCCCG-CCCCG-CG. This sequence, called a dodecamer repeat, is usually repeated two or three times within a part of the gene that helps regulate cystatin B protein production.

**Health Conditions Related to Genetic Changes**

**Unverricht-Lundborg disease**

In almost all affected individuals, Unverricht-Lundborg disease is caused by an increased number of copies (expansion) of the dodecamer repeat in the *CSTB* gene. Most people with this disorder have more than 30 repeats of the dodecamer sequence in both copies of the *CSTB* gene.

In a small number of individuals, one copy of the *CSTB* gene has the expanded dodecamer repeat while the second copy carries one of nine other identified mutations. Some of these mutations substitute one protein building block (amino acid) for another amino acid in the cystatin B protein. Others result in a shortened protein that may function improperly or not at all, or cause the protein to be pieced together incorrectly. Only one individual with Unverricht-Lundborg disease has been reported to have mutations other than the dodecamer repeat expansion in both copies of the gene in each cell.

The expanded dodecamer repeat in the *CSTB* gene seems to interfere with the production of cystatin B protein. Levels of cystatin B in affected individuals are only 5 to 10 percent of normal, and cathepsin levels are significantly increased. These changes are believed to cause the signs and symptoms of Unverricht-Lundborg disease, but the specific mechanism is unknown.
Chromosomal Location

Cytogenetic Location: 21q22.3, which is the long (q) arm of chromosome 21 at position 22.3

Molecular Location: base pairs 43,773,665 to 43,776,375 on chromosome 21 (Homo sapiens Updated Annotation Release 109.20190905, GRCh38.p13) (NCBI)

Credit: Genome Decoration Page/NCBI

Other Names for This Gene

- CPI-B
- CST6
- cystatin B (stefin B)
- CYTB_HUMAN
- EPM1
- liver thiol proteinase inhibitor
- PME
- stefin B
- STFB

Additional Information & Resources

Clinical Information from GeneReviews

- Unverricht-Lundborg Disease
  https://www.ncbi.nlm.nih.gov/books/NBK1142
Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28CSTB%5BTIAB%5D%29+OR+%28cystatin+B%5BTIAB%5D%29+OR+%28EPM1%5BTIAB%5D%29+OR+%28STFB%5BTIAB%5D%29+OR+%28cystatin+B%5BTIAB%5D%29+OR+%28CPI-B%5BTIAB%5D%29+OR+%28stefin+B%5BTIAB%5D%29+OR+%28liver+thiol+proteinase+inhibitor%5BTIAB%5D%29+OR+%28Genes%5BMH%5D%29+OR+%28Genetic+Phenomena%5BMH%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+1440+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- CYSTATIN B
  http://omim.org/entry/601145

Research Resources

- Atlas of Genetics and Cytogenetics in Oncology and Haematology
  http://atlasgeneticsoncology.org/Genes/CSTBID40181ch21q22.html
- ClinVar
  https://www.ncbi.nlm.nih.gov/clinvar?term=CSTB%5Bgene%5D
- HGNC Gene Symbol Report
- Monarch Initiative
  https://monarchinitiative.org/gene/NCBIGene:1476
- NCBI Gene
- UniProt
  https://www.uniprot.org/uniprot/P04080

Sources for This Summary

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  http://omim.org/entry/601145
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Reprinted from Genetics Home Reference:  

Reviewed: June 2008  
Published: October 1, 2019  

Lister Hill National Center for Biomedical Communications  
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