SCN10A gene
sodium voltage-gated channel alpha subunit 10

Normal Function

The SCN10A gene belongs to a family of genes that provide instructions for making sodium channels. These channels, which transport positively charged sodium atoms (sodium ions) into cells, play a key role in a cell's ability to generate and transmit electrical signals.

The SCN10A gene provides instructions for making one part (the alpha subunit) of a sodium channel called NaV1.8. NaV1.8 sodium channels are found in nerve cells called nociceptors that transmit pain signals. Nociceptors are part of the peripheral nervous system, which connects the brain and spinal cord to cells that detect sensations such as touch, smell, and pain. Nociceptors are primarily involved in transmitting pain signals. The centers of nociceptors, known as the cell bodies, are located in a part of the spinal cord called the dorsal root ganglion. Fibers called axons extend from the cell bodies, reaching throughout the body to receive sensory information. In addition to nociceptors, NaV1.8 sodium channels have also been found in heart muscle cells where, by controlling the flow of sodium ions, they likely play a role in maintaining a normal heart rhythm.

Health Conditions Related to Genetic Changes

Small fiber neuropathy

Mutations in the SCN10A gene account for approximately 5 percent of cases of small fiber neuropathy, a condition characterized by severe pain attacks and a reduced ability to differentiate between hot and cold. The mutations that cause this condition change single protein building blocks (amino acids) in the alpha subunit of the NaV1.8 sodium channel. Many of the mutations result in NaV1.8 sodium channels that open more easily than usual, increasing the flow of sodium ions that produce nerve impulses within nociceptors. This increase in sodium ions enhances transmission of pain signals, causing individuals to be more sensitive to stimulation that might otherwise not cause pain. In this condition, the small fibers that extend from the nociceptors and transmit pain signals (axons) degenerate over time. The cause of this degeneration is unknown, but it likely accounts for signs and symptoms such as the loss of temperature differentiation.

Other disorders

Certain common variants (polymorphisms) in the SCN10A gene have been found to increase the risk of developing an irregular heartbeat (arrhythmia). These polymorphisms lead to the production of an altered NaV1.8 sodium channel that can
disrupt the electrical signals that control the heartbeat. Specifically, changes in the *SCN10A* gene are associated with a type of arrhythmia known as heart block. Heart block occurs when the heart's electrical signals are slowed down or interrupted. It is unknown how changes to the NaV1.8 sodium channel lead to heart block.

**Chromosomal Location**

Cytogenetic Location: 3p22.2, which is the short (p) arm of chromosome 3 at position 22.2

Molecular Location: base pairs 38,697,110 to 38,794,010 on chromosome 3 (Homo sapiens Updated Annotation Release 109.20200228, GRCh38.p13) (NCBI)

Credit: Genome Decoration Page/NCBI

**Other Names for This Gene**

- hPN3
- Nav1.8
- peripheral nerve sodium channel 3
- PN3
- SCNAA_HUMAN
- sodium channel protein type 10 subunit alpha
- sodium channel protein type X subunit alpha
- sodium channel, voltage gated, type X alpha subunit
- sodium channel, voltage-gated, type X, alpha polypeptide
- sodium channel, voltage-gated, type X, alpha subunit
- voltage-gated sodium channel subunit alpha Nav1.8
Additional Information & Resources

Educational Resources

- American Heart Association: About Arrhythmia
  https://www.heart.org/en/health-topics/arrhythmia/about-arrhythmia

- Biochemistry (fifth edition, 2002): The Sodium Channel (figure)
  https://www.ncbi.nlm.nih.gov/books/NBK22509/figure/A1820/

- National Heart Lung and Blood Institute: Conduction Disorders
  https://www.nhlbi.nih.gov/health-topics/conduction-disorders

  https://www.ncbi.nlm.nih.gov/books/NBK10965/

- Washington University, St. Louis Neuromuscular Disease Center: Voltage-Gated Sodium Channels
  https://neuromuscular.wustl.edu/mother/chan.html#nachvg

Scientific Articles on PubMed

- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28SCN10A%5BTIAB%5D%29+OR+%28Nav1.8%5BTIAB%5D%29+AND+%28%28Genes%5BMH%5D%29+OR+%28Genetic+Phenomena%5BMH%5D%29%29+AND+english%5BLa%5D+AND+human%5Bmh%5D+AND+&22%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM

- SODIUM CHANNEL, VOLTAGE-GATED, TYPE X, ALPHA SUBUNIT
  http://omim.org/entry/604427

Research Resources

- ClinVar

- HGNC Gene Symbol Report

- Monarch Initiative
  https://monarchinitiative.org/gene/NCBIGene:6336

- NCBI Gene

- UniProt
  https://www.uniprot.org/uniprot/Q9Y5Y9
Sources for This Summary

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20062061

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/23115331
  Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3511073/

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/21646736

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