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(54) Title: COMPOSITIONS AND METHODS FOR TREATING DISORDERS ASSOCIATED WITH LOSS-OF-FUNCTION MUTATIONS IN SCN2A

(57) Abstract: Provided herein are methods for increasing levels of SCN2A protein in a cell, comprising contacting the cell with an antisense oligonucleotide that enhances splicing at a splice site of a retained intron in an intron-retaining SCN2A mRNA or pre-mRNA, wherein the retained intron is selected from among intron 1, 2, 3, 4, 5, 11, 13, 17 and 24 and wherein the antisense oligonucleotide comprises a sequence of nucleobases that is complementary to a target region in the SCN2A mRNA or pre-mRNA. Also provided are antisense oligonucleotides for use in such methods. Also provided are methods for treating disorders associated with a heterozygous loss-of-function mutation in SCN2A, comprising administering to the subject such antisense oligonucleotides.



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