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(54) Title: RNAI AGENTS FOR INHIBITING EXPRESSION OF DUX4, COMPOSITIONS THEREOF, AND METHODS OF USE

(57) Abstract: Described are RNAi agents, compositions that include RNAi agents, and methods for inhibition of a double homeobox 4 (DUX4) gene. The DUX4 RNAi agents and RNAi agent conjugates disclosed herein inhibit the expression of a DUX4 gene. Pharmaceutical compositions that include one or more DUX4 RNAi agents, optionally with one or more additional therapeutics, are also described. Delivery of the described DUX4 RNAi agents to skeletal muscle cells *in vivo*, provides for inhibition of DUX4 gene expression and a reduction in DUX4 levels, which can provide a therapeutic benefit to subjects, including human subjects, suffering from certain skeletal muscle-related diseases or disorders including Facioscapulohumeral Muscular Dystrophy (FSHD).



WO 2022/056266 A3