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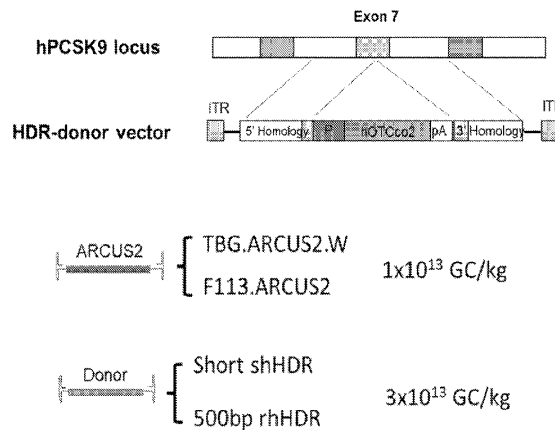
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(54) Title: METHODS FOR TREATMENT OF ORNITHINE TRANSCARBAMYLASE (OTC) DEFICIENCY

FIG 2



Total N = 14

Serotypes: AAVhu37 (n=4); AAVrh79 (n=8+2 ctl)

(57) Abstract: A dual vector system for treating ornithine transcarbamylase deficiency is provided. The system includes ((a) a gene editing AAV comprising a first AAV rh79 capsid and a first vector genome comprising a 5' ITR, a sequence encoding a meganuclease that targets PCSK9 under control of regulatory sequences that direct expression of the meganuclease in a target cell comprising a PCSK9 gene, and a 3' ITR; and (b) a donor AAV vector comprising a second AAV capsid and a second AAV rh79 vector genome comprising: a 5' ITR, a 5' homology directed recombination (HDR) arm, a transgene encoding ornithine transcarbamylase (OTC) and regulatory sequences that direct expression of the transgene in the target cell, a 3' HDR arm, and a 3' ITR.

[Continued on next page]

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