

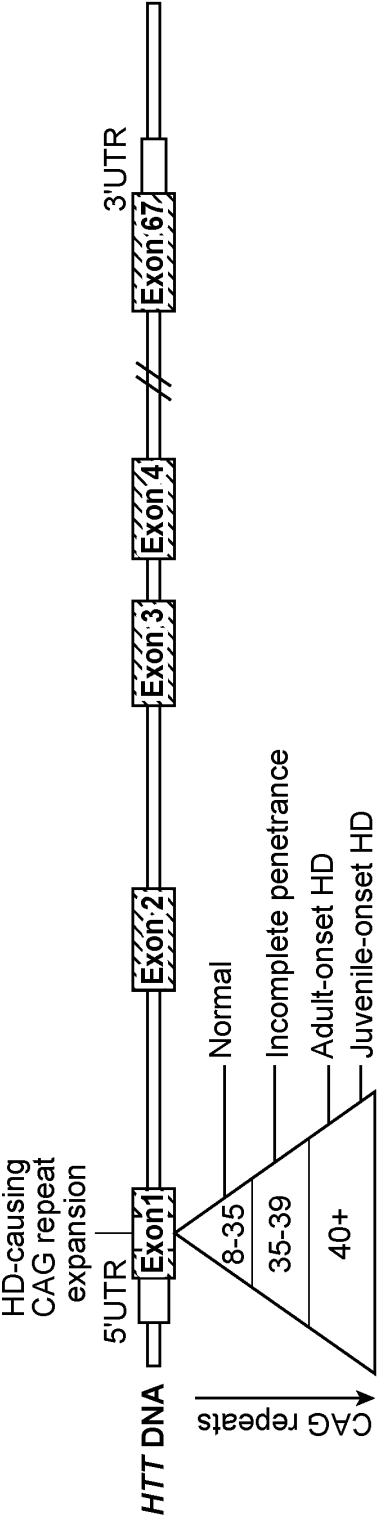


FIG. 1

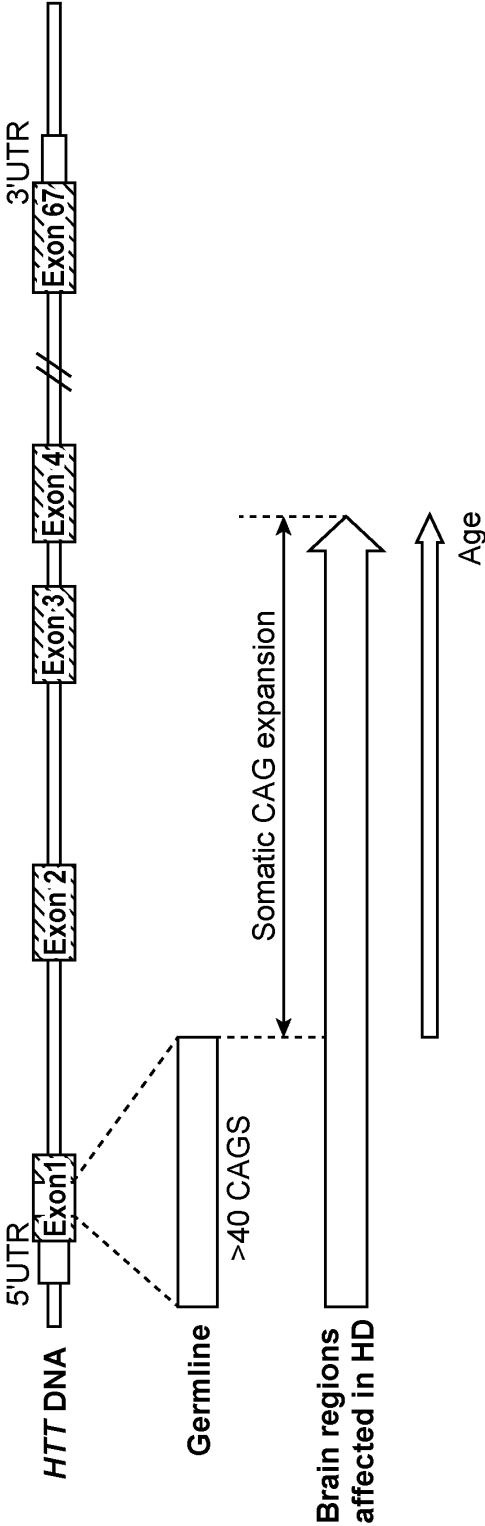


FIG. 2

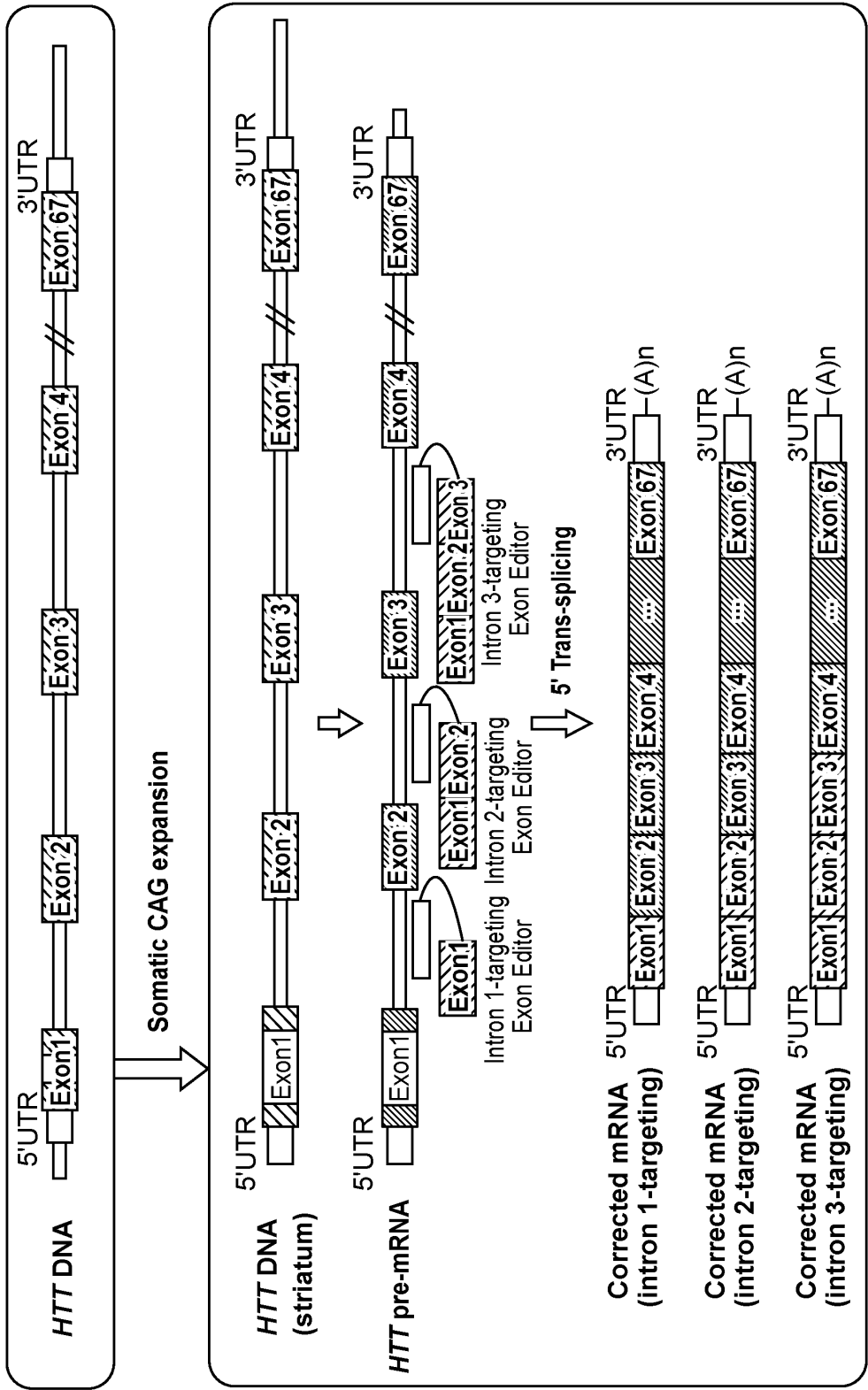


FIG. 3

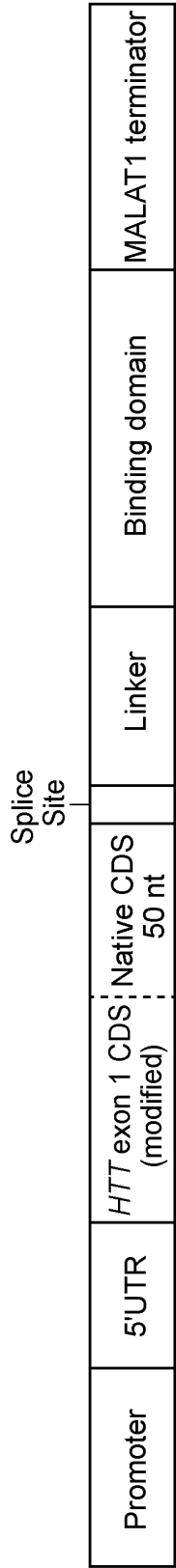


FIG. 4

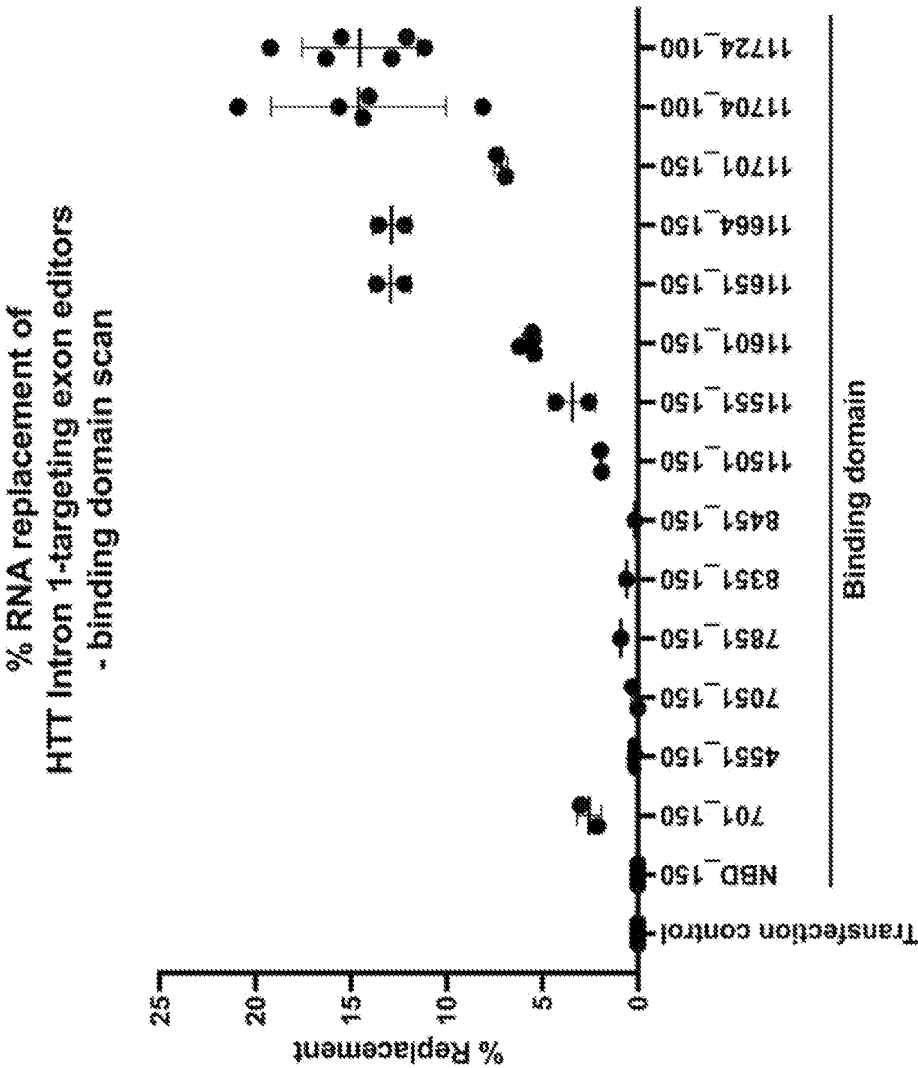


FIG. 5

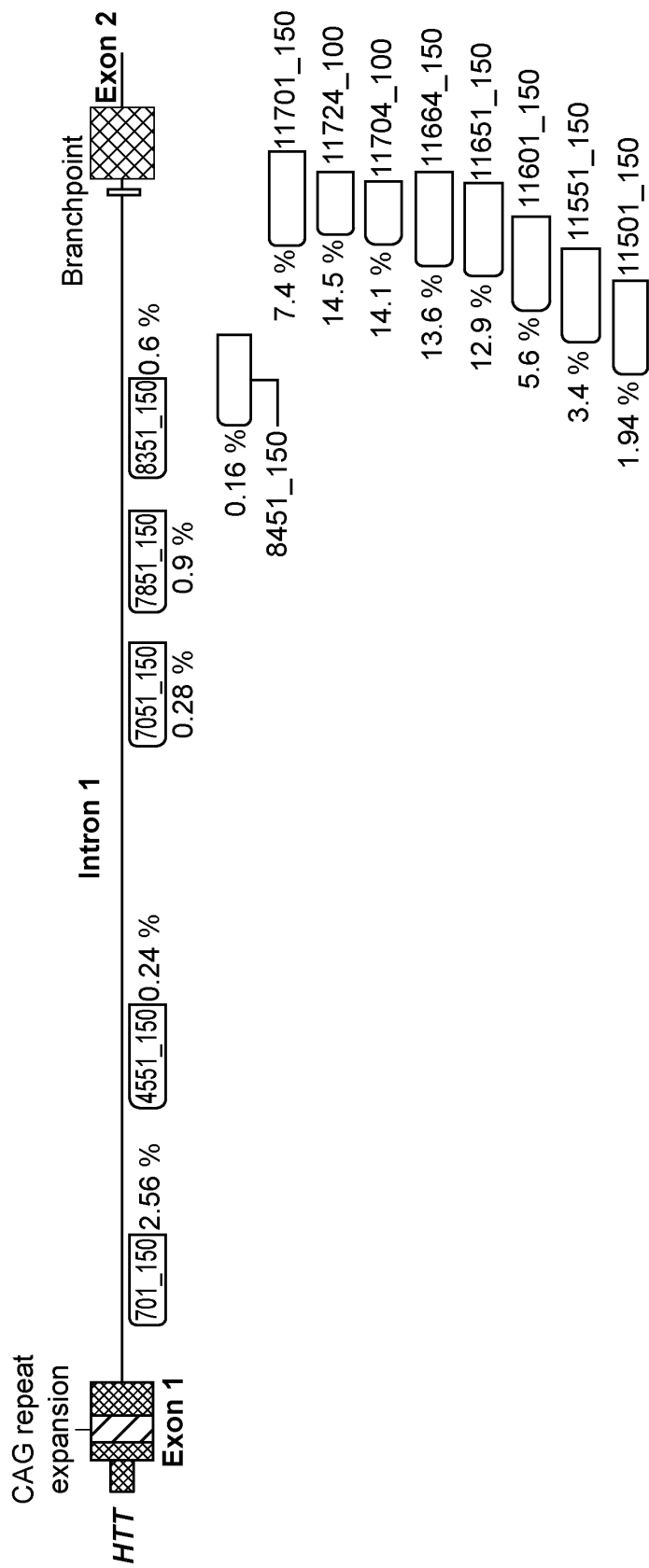


FIG. 6

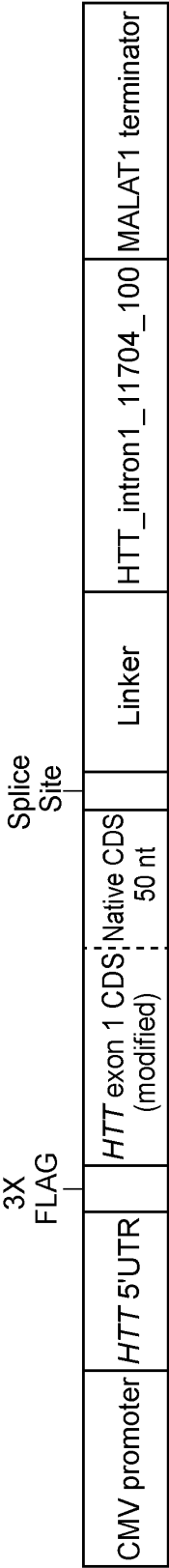


FIG. 7

% RNA Replacement by
HTT_intron1_11704_100 exon editors

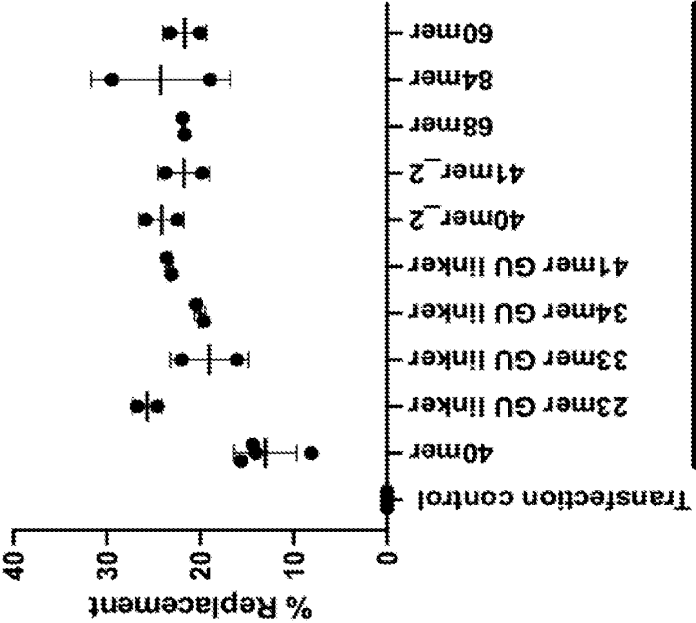


FIG. 8

Promoter	5'UTR	Splice Site				MALAT1 terminator
		exon 1 (modified)	exon 2 (modified)	Native exon 2 35 nt	Linker	

FIG. 9

% RNA replacement of
HTT intron 2-targeting exon editors

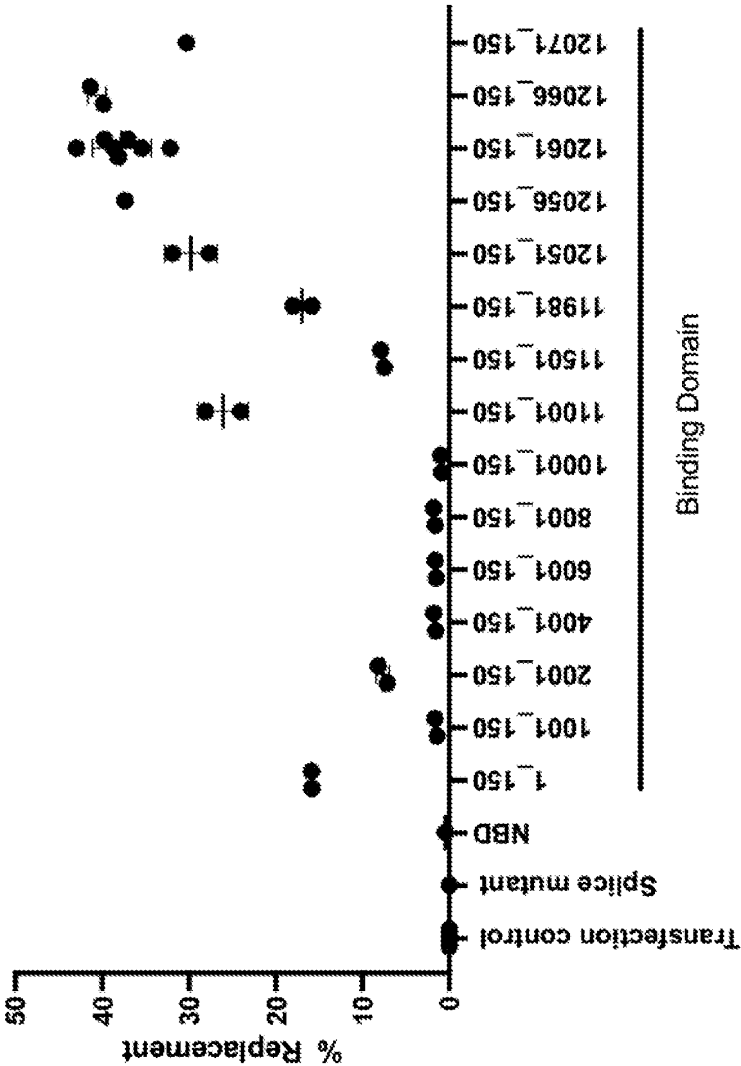


FIG. 10

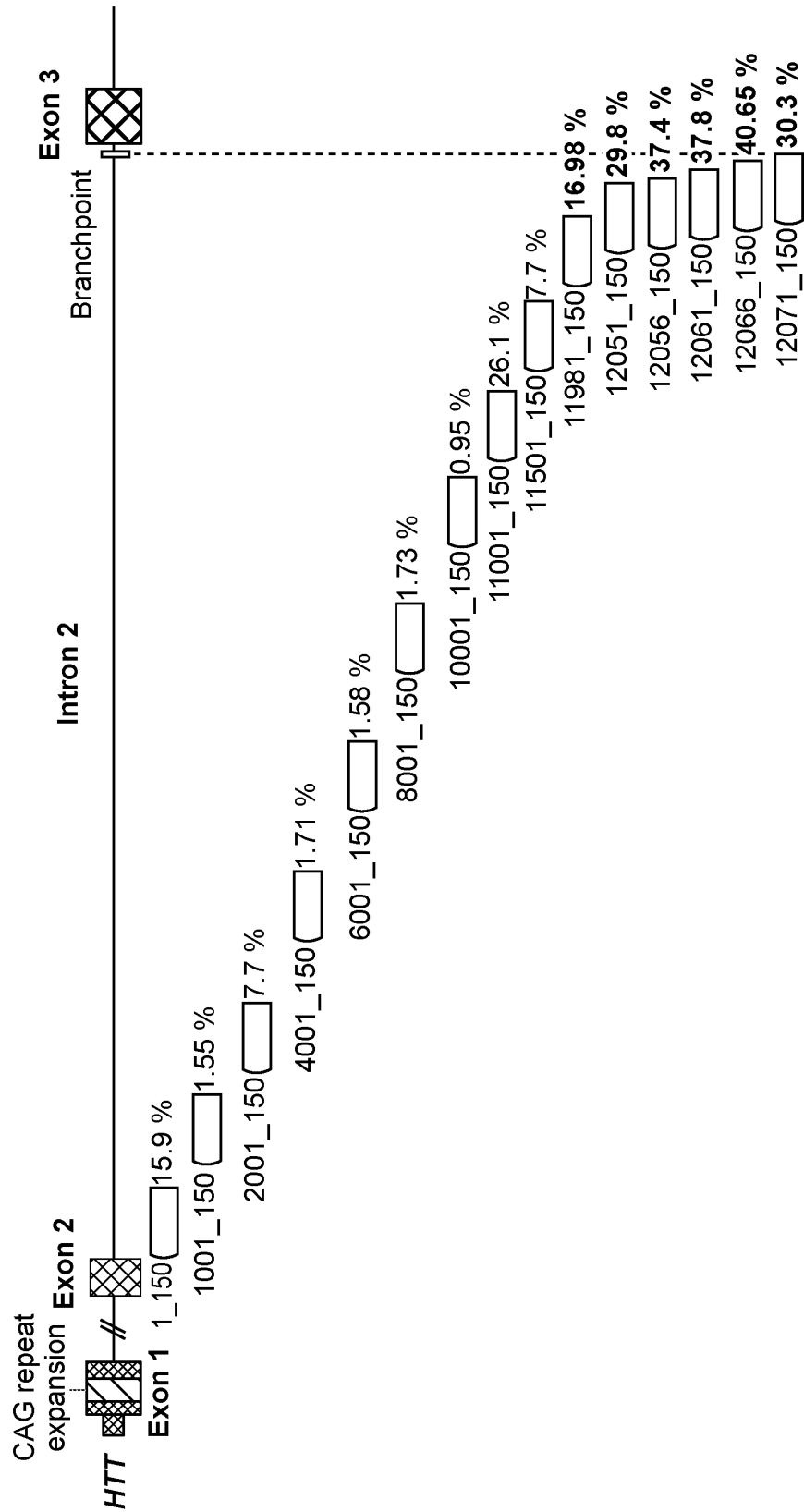


FIG. 11

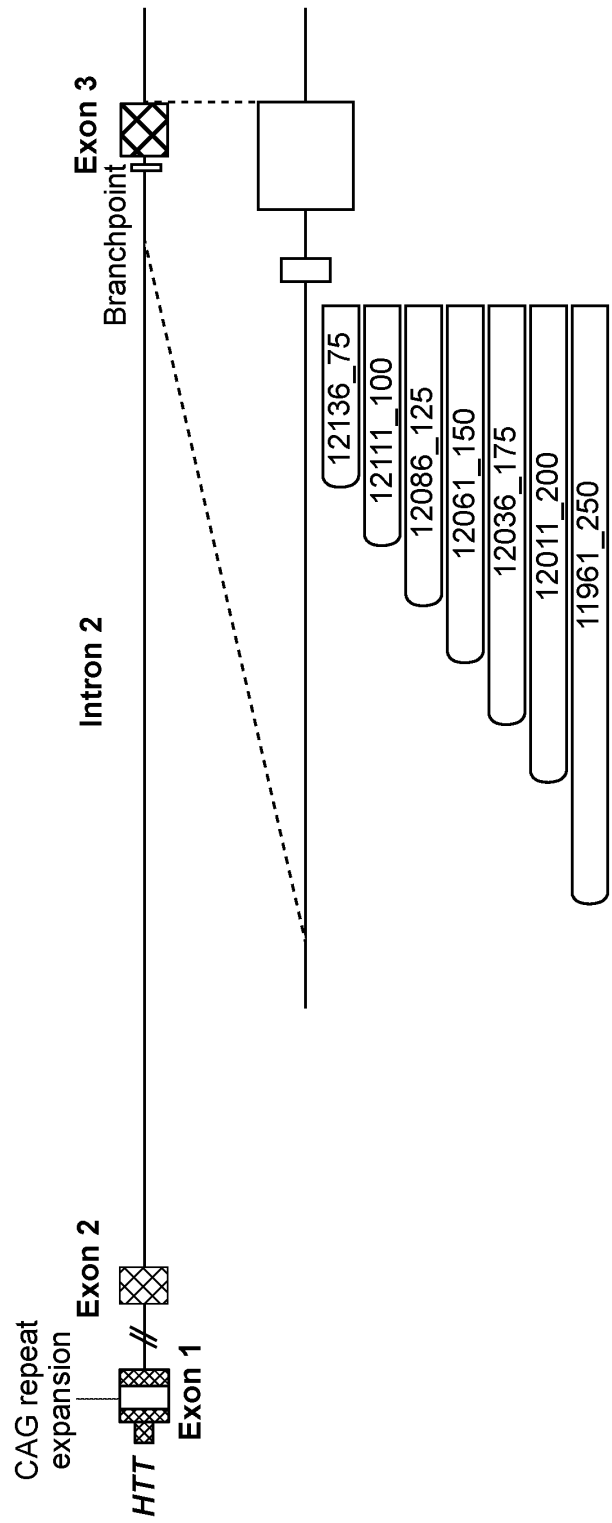


FIG. 12A

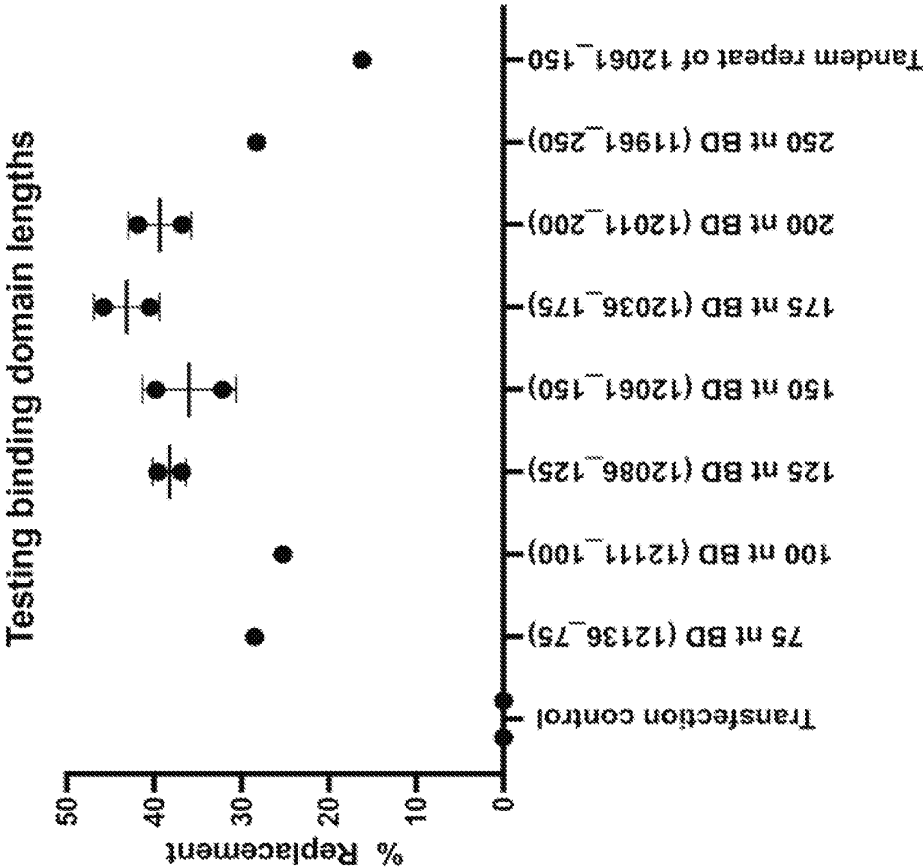


FIG. 12B

3X FLAG			Splice Site		
CMV promoter	HTT 5'UTR	exon 1 (modified)	exon 2 (modified)	Native exon 2 35 nt	
					Linker
					HTT_intron2_12061_150
					MALAT1 terminator

FIG. 13

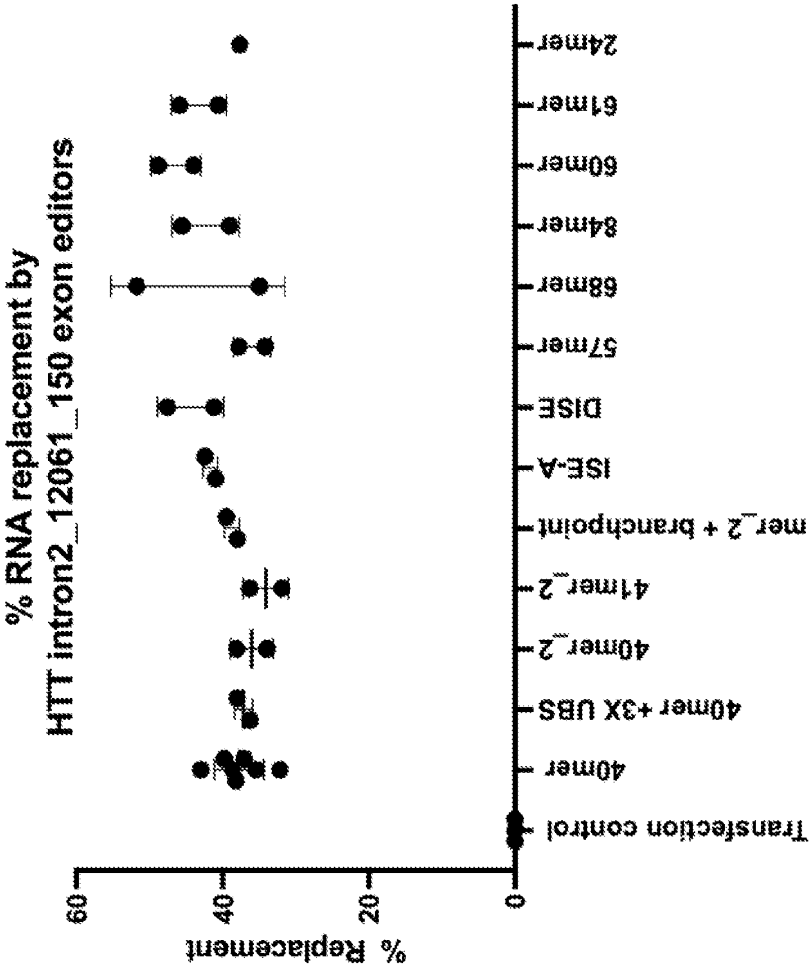


FIG. 14

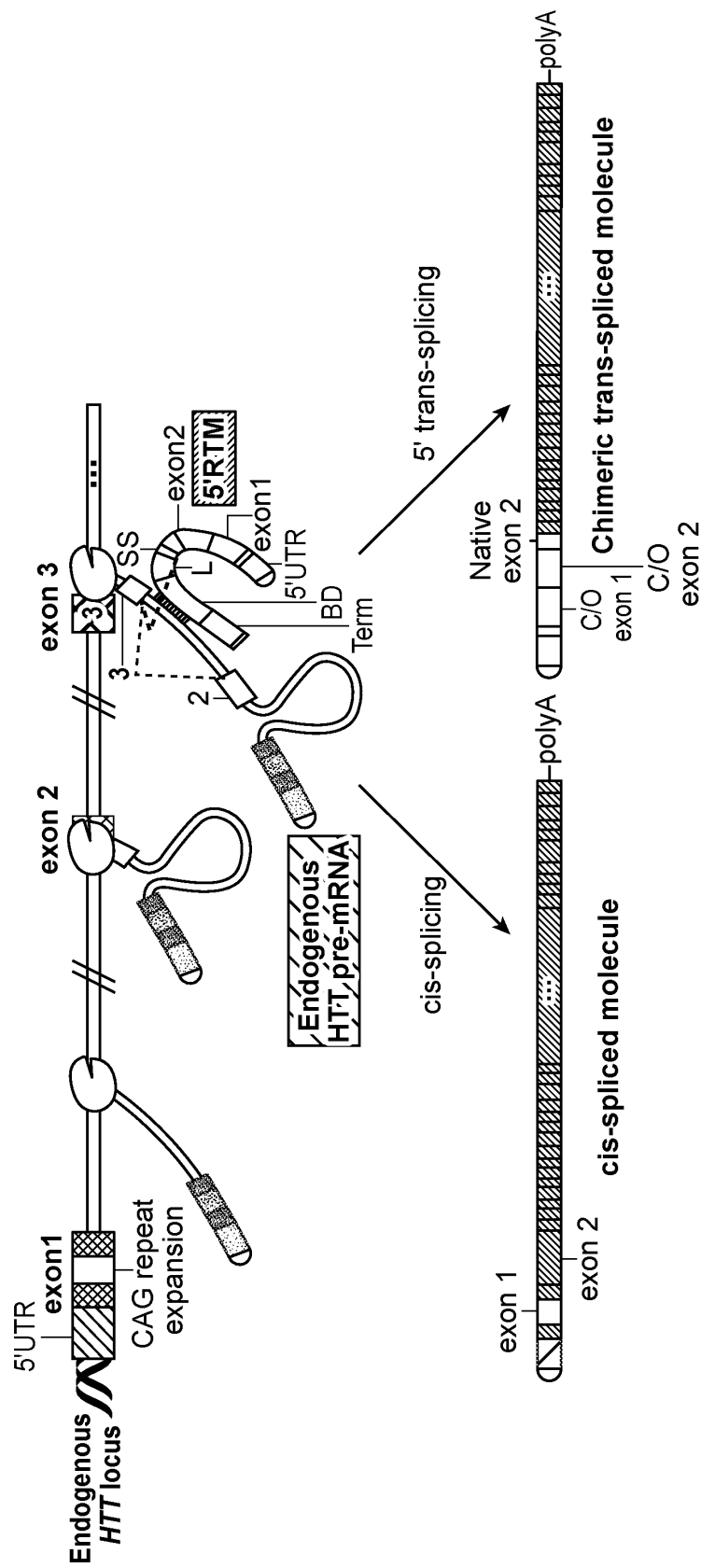


FIG. 15

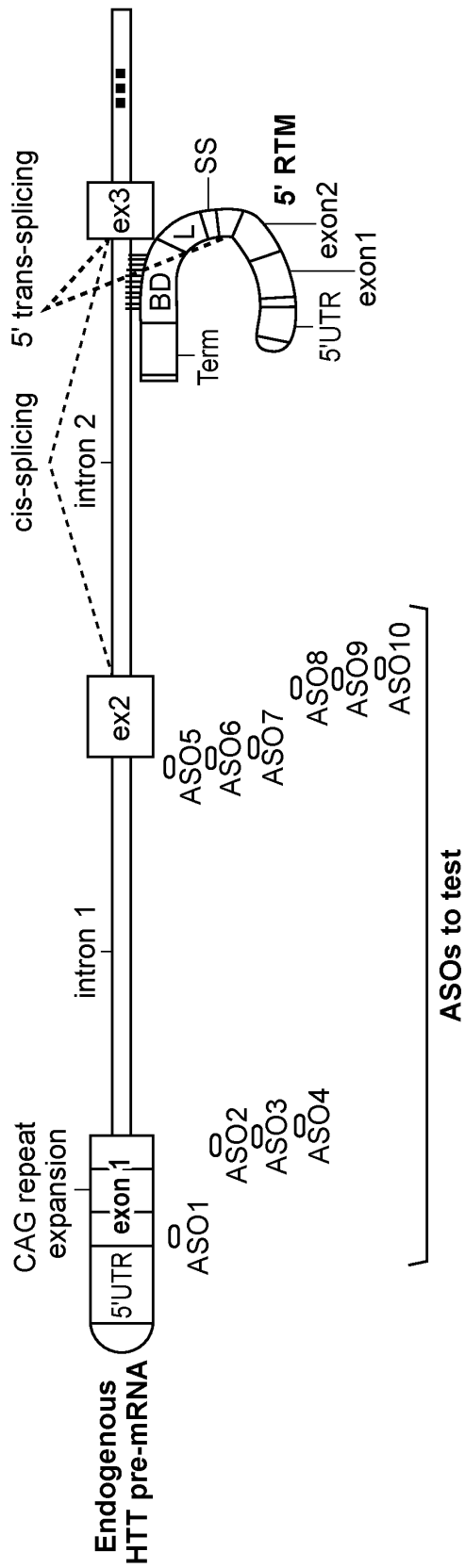


FIG. 16

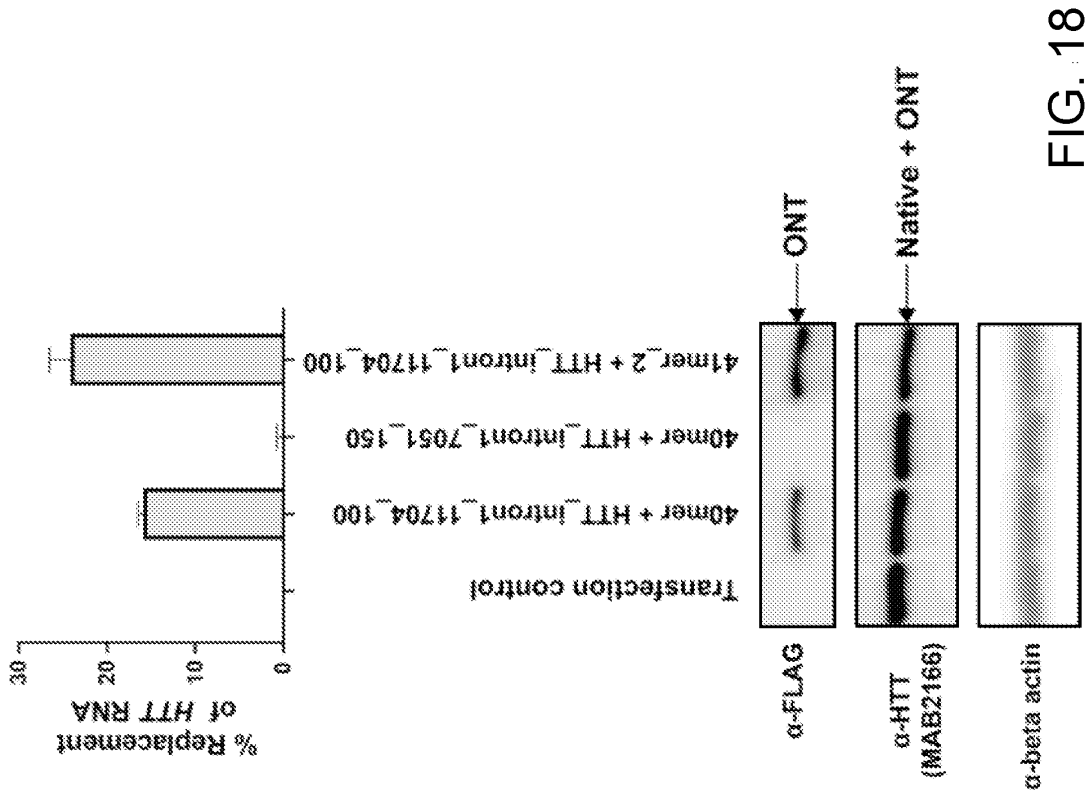


FIG. 17

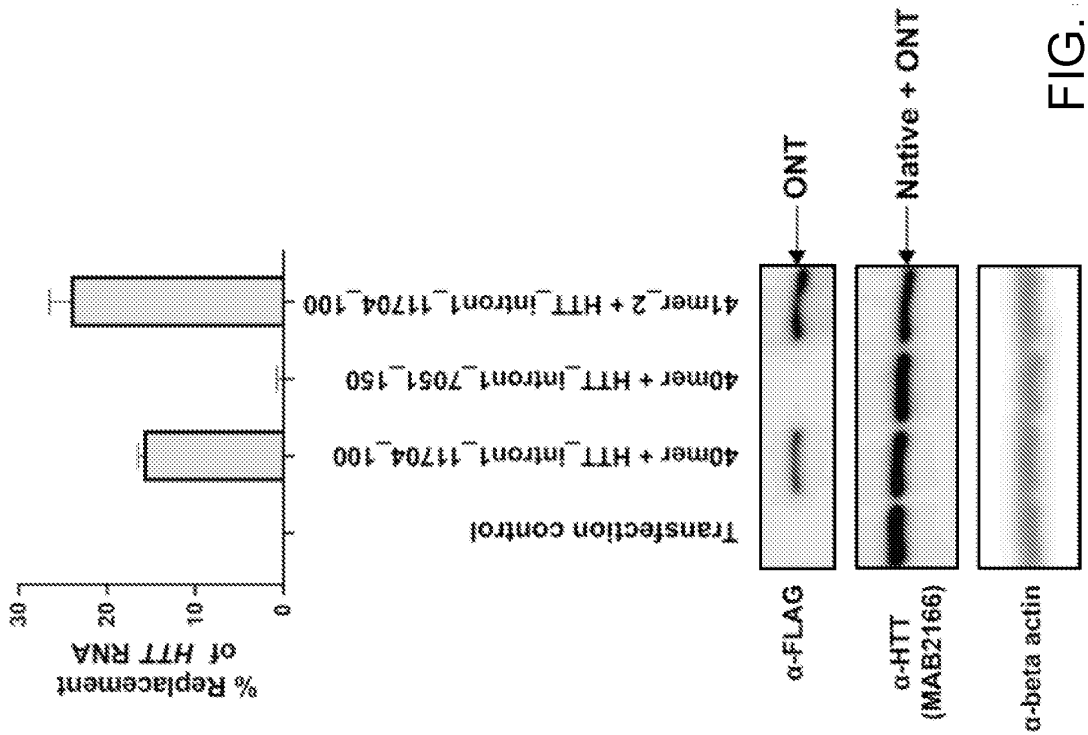


FIG. 18

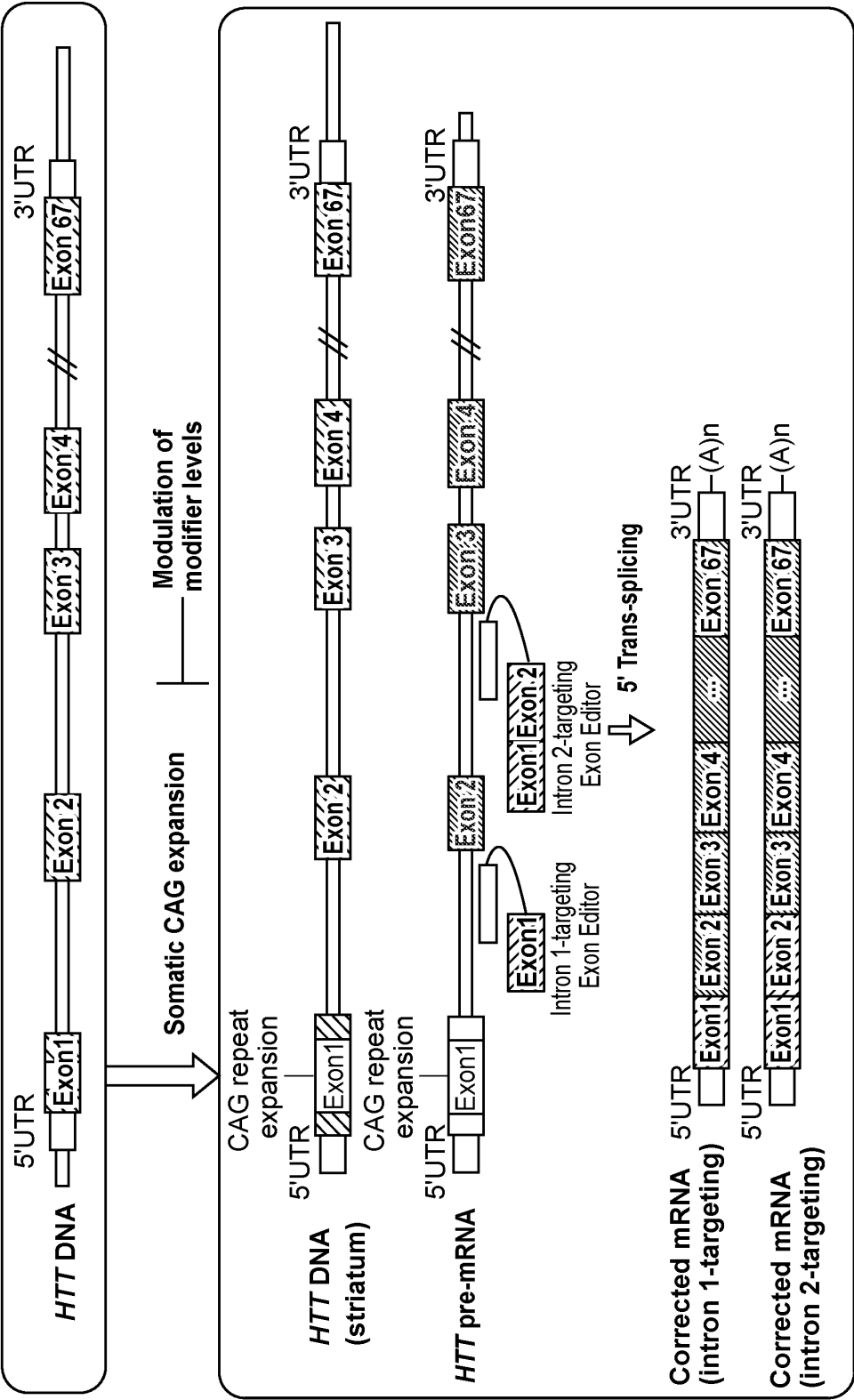


FIG. 19

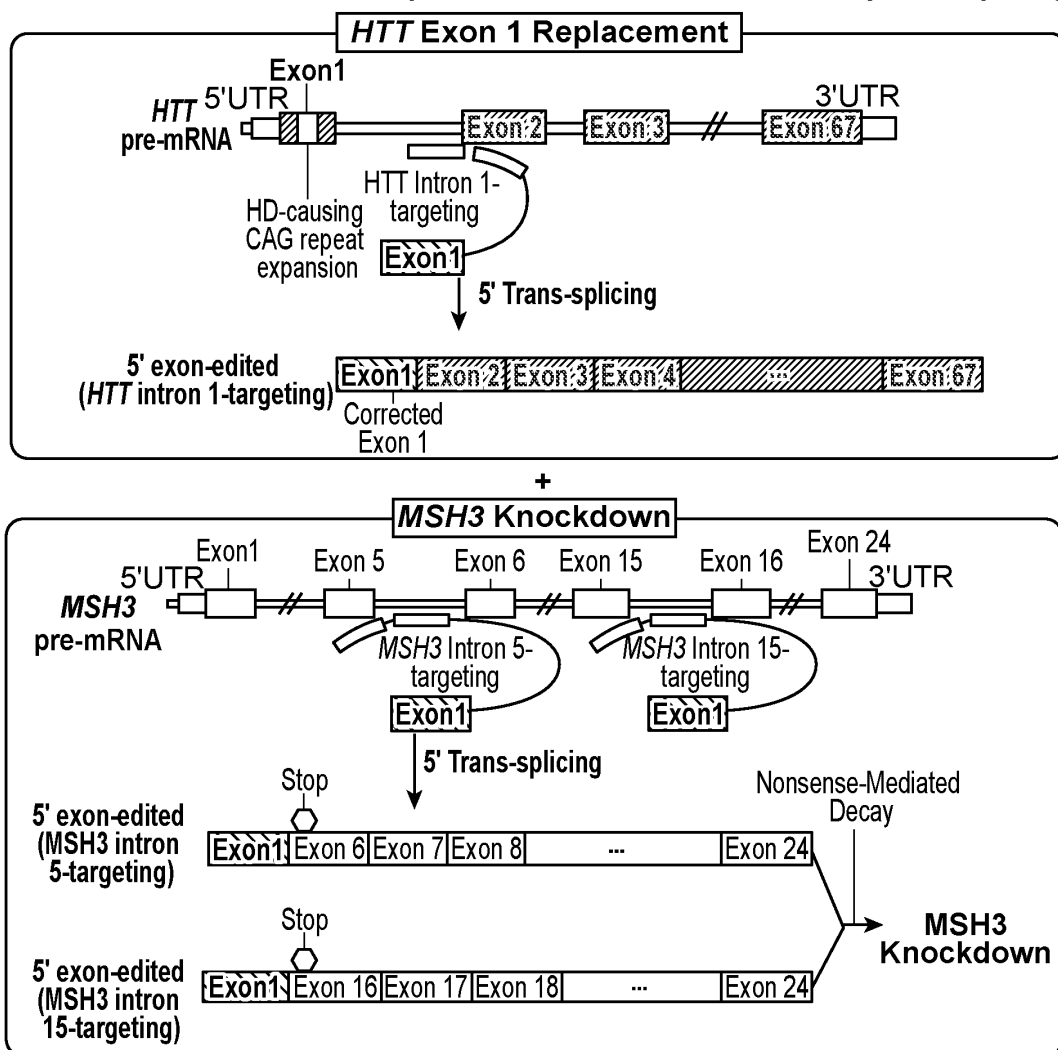
Dual action RTM: HTT exon 1 replacement + knockdown of MSH3 by trans-splicing

FIG. 20

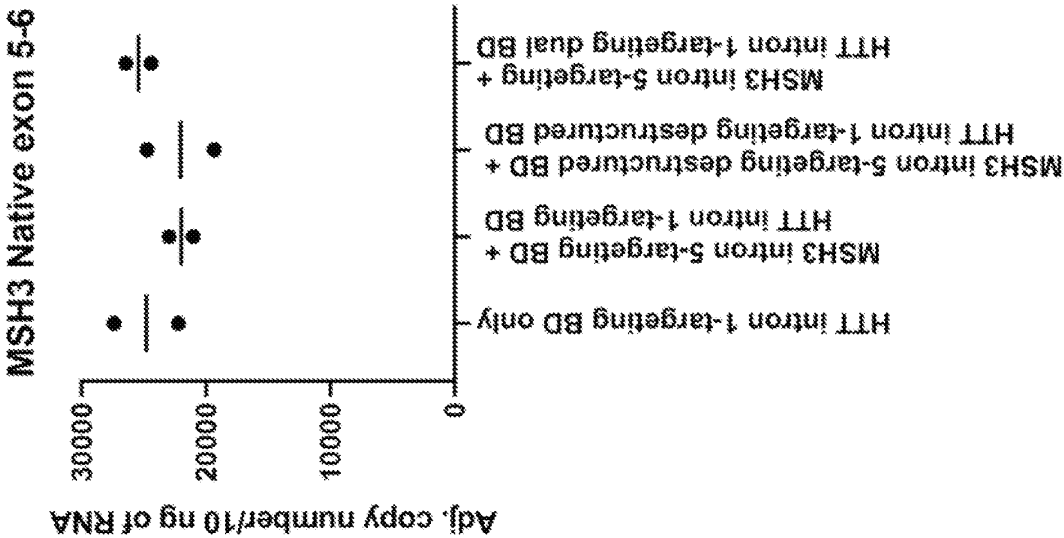


FIG. 21C

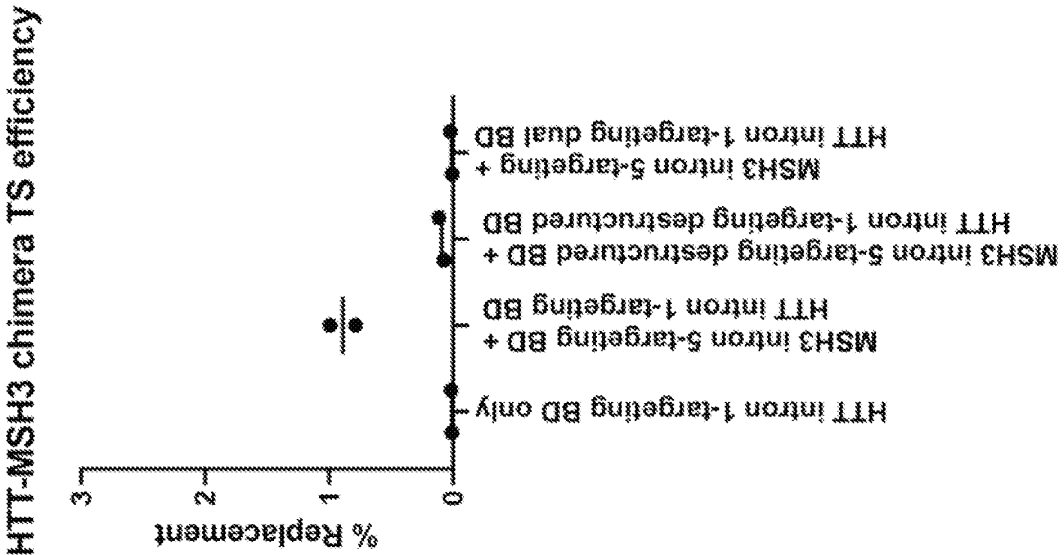


FIG. 21B

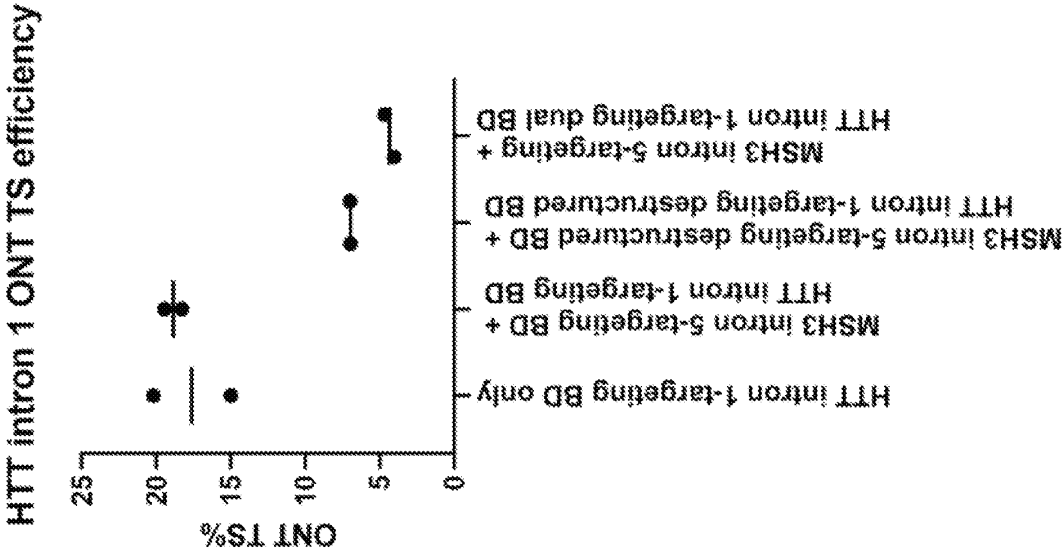


FIG. 21A

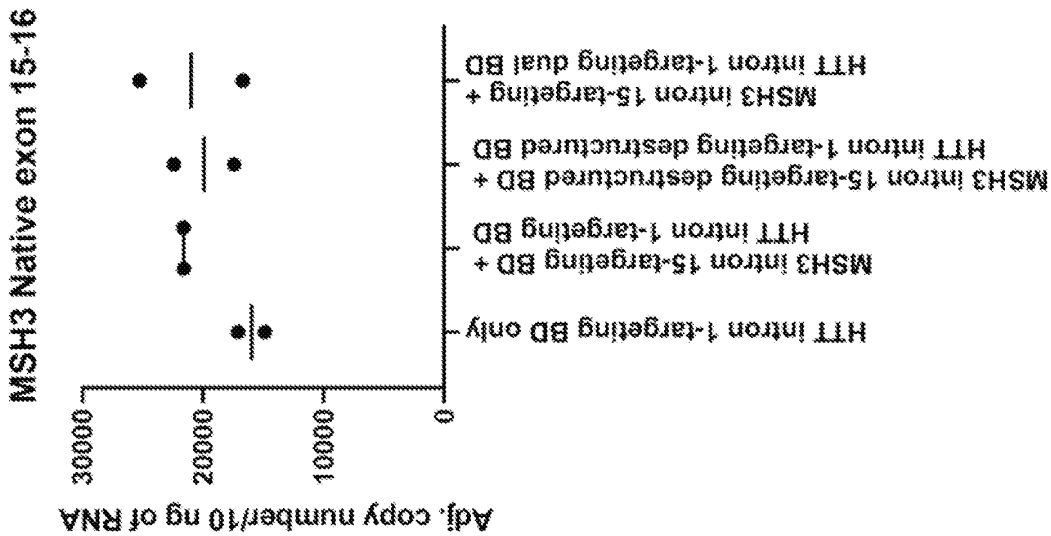


FIG. 22C

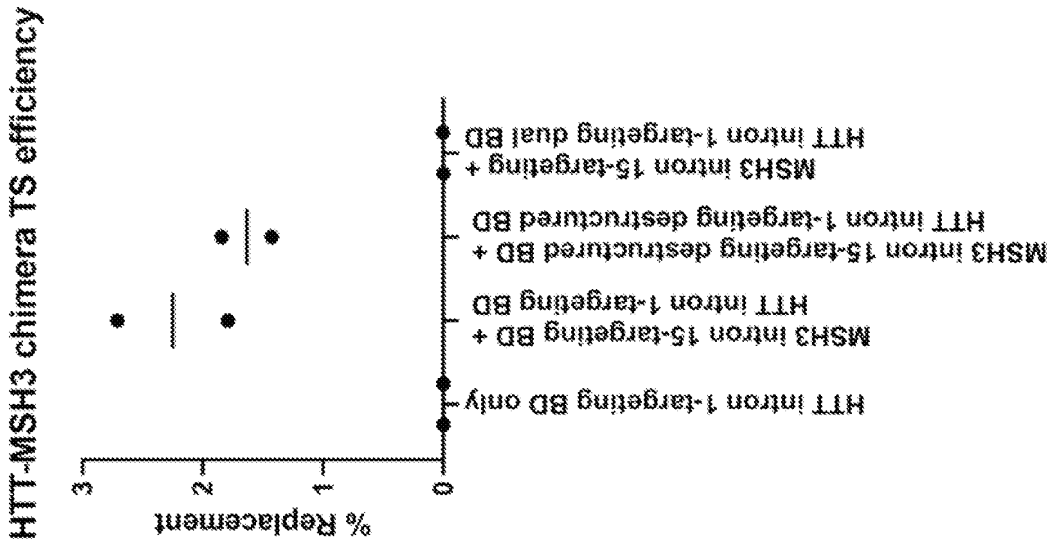


FIG. 22B

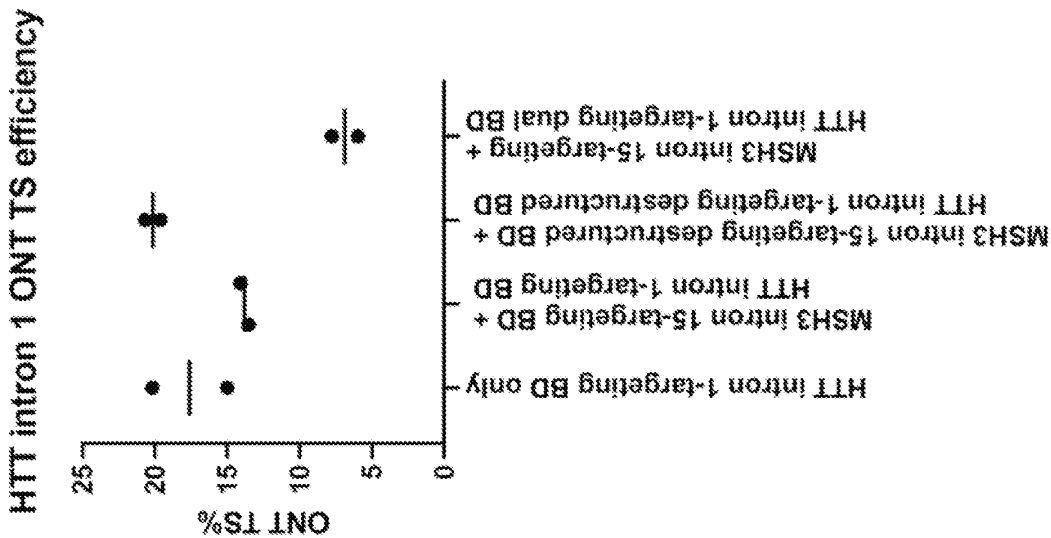


FIG. 22A

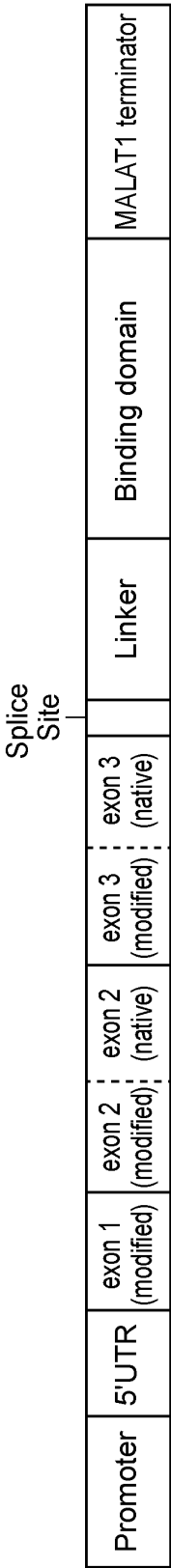


FIG. 23

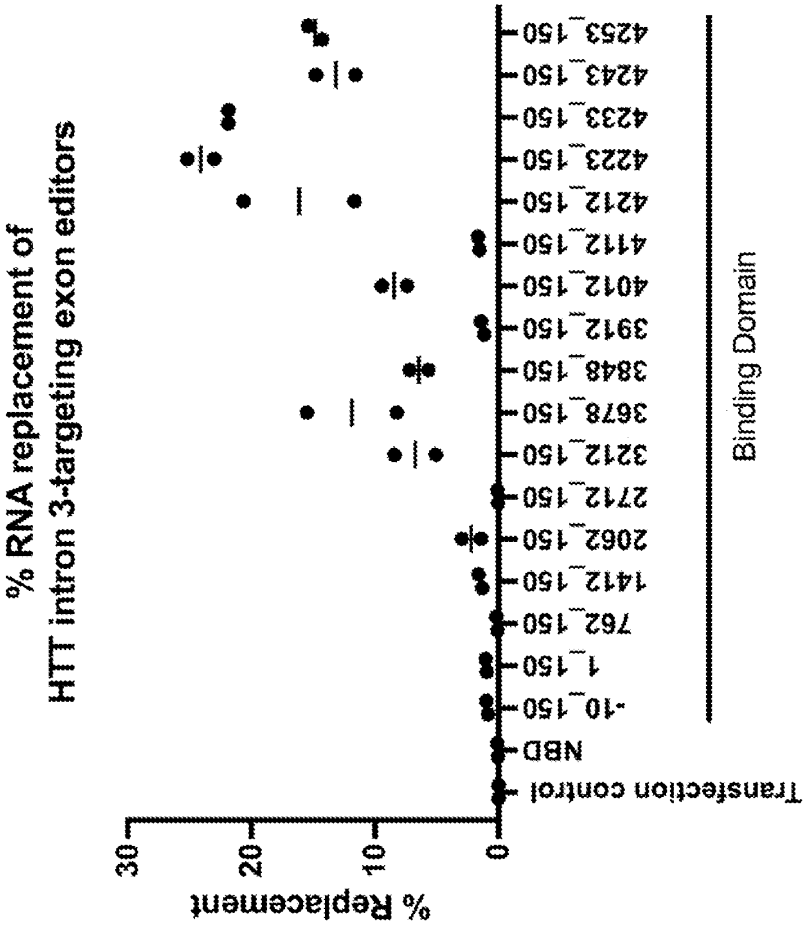


FIG. 24

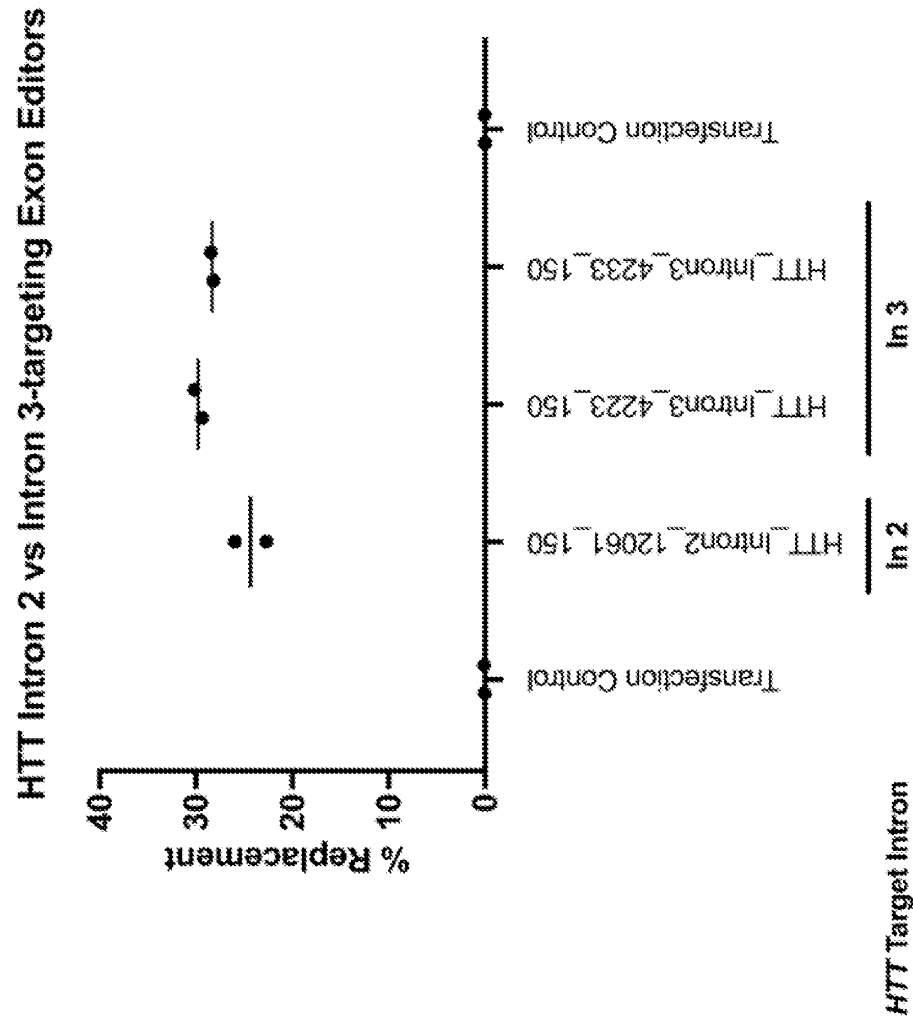


FIG. 25

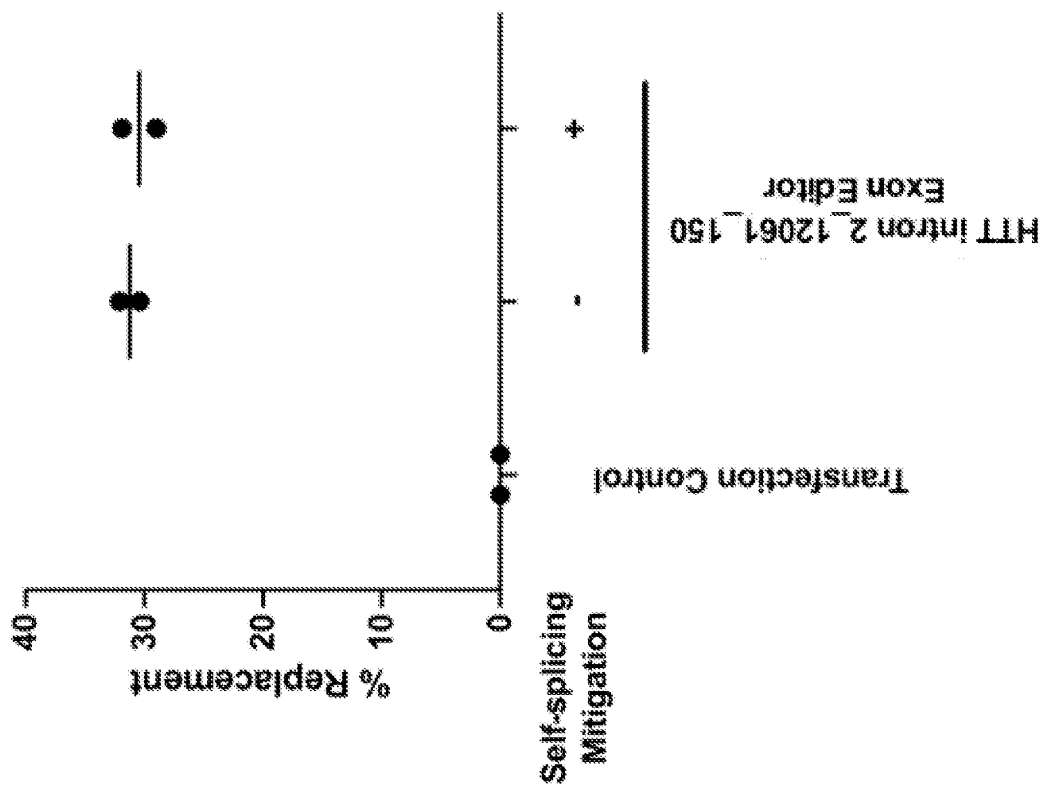


FIG. 26

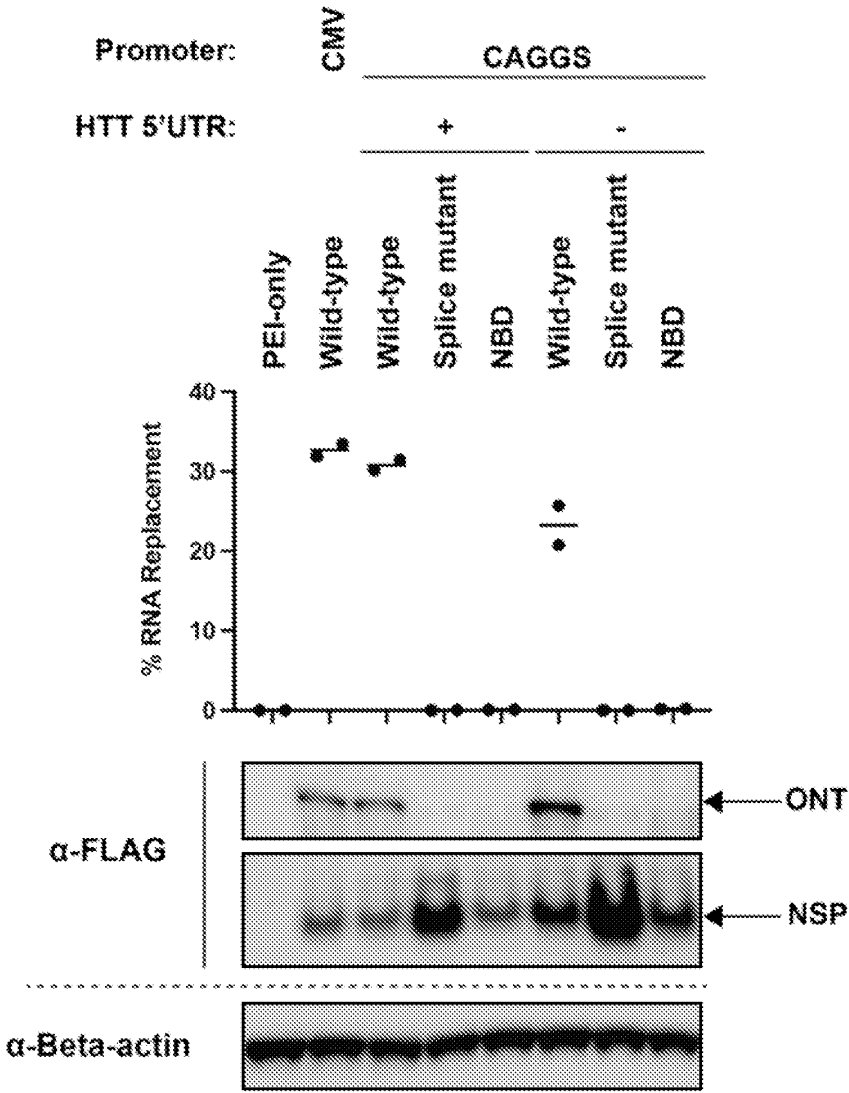


FIG. 27

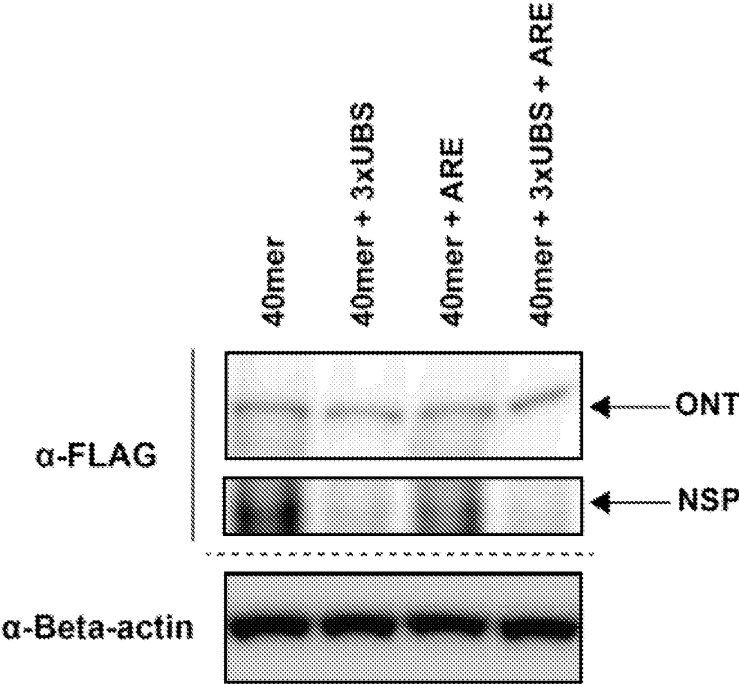


FIG. 28

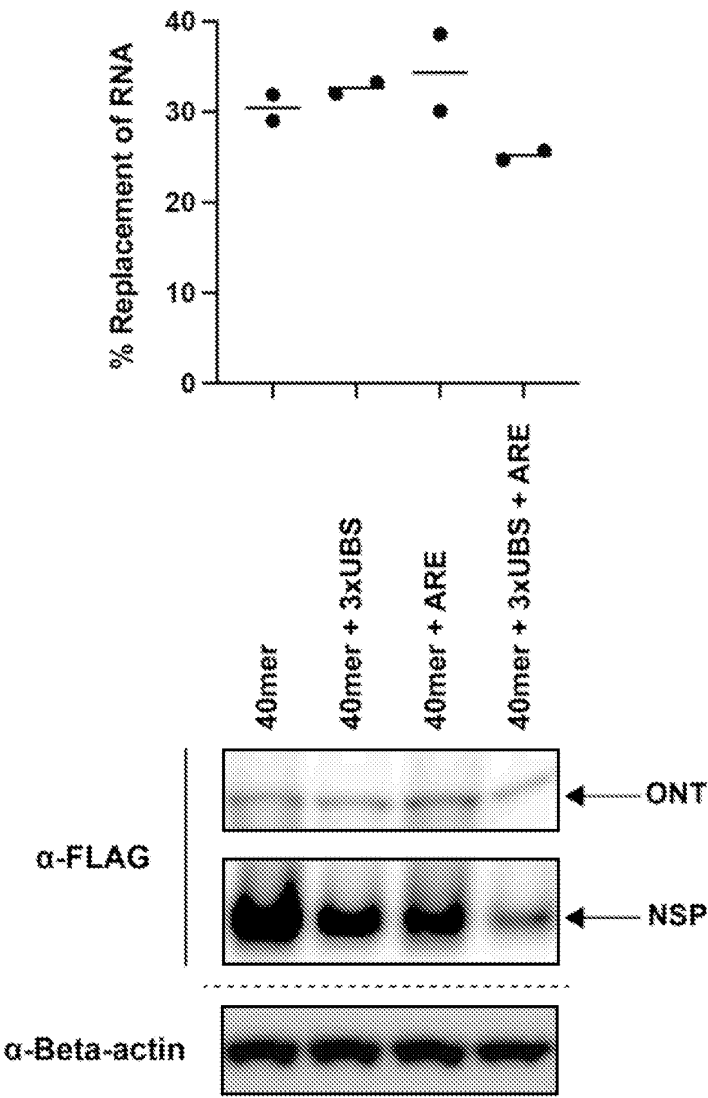


FIG. 29

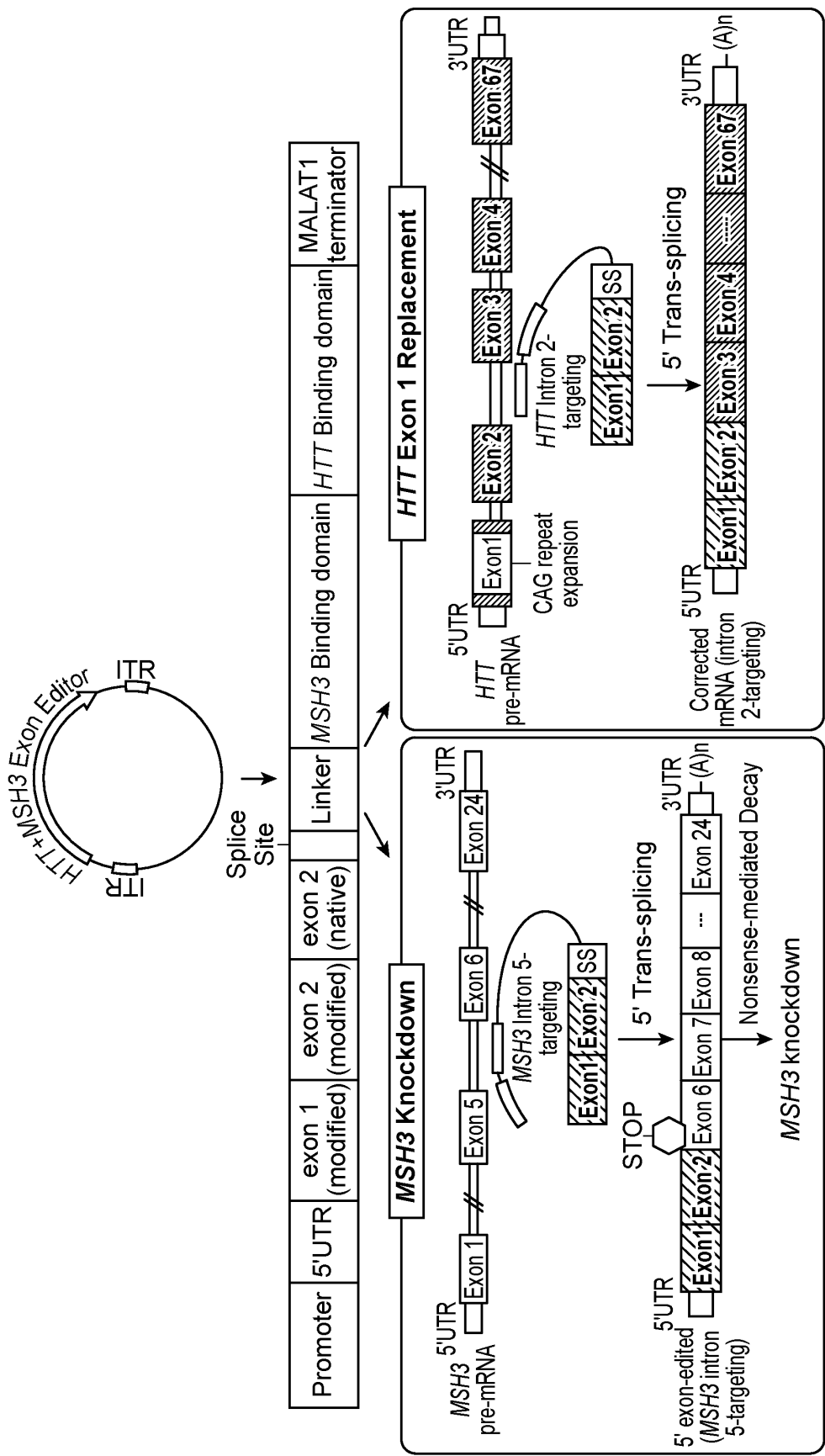
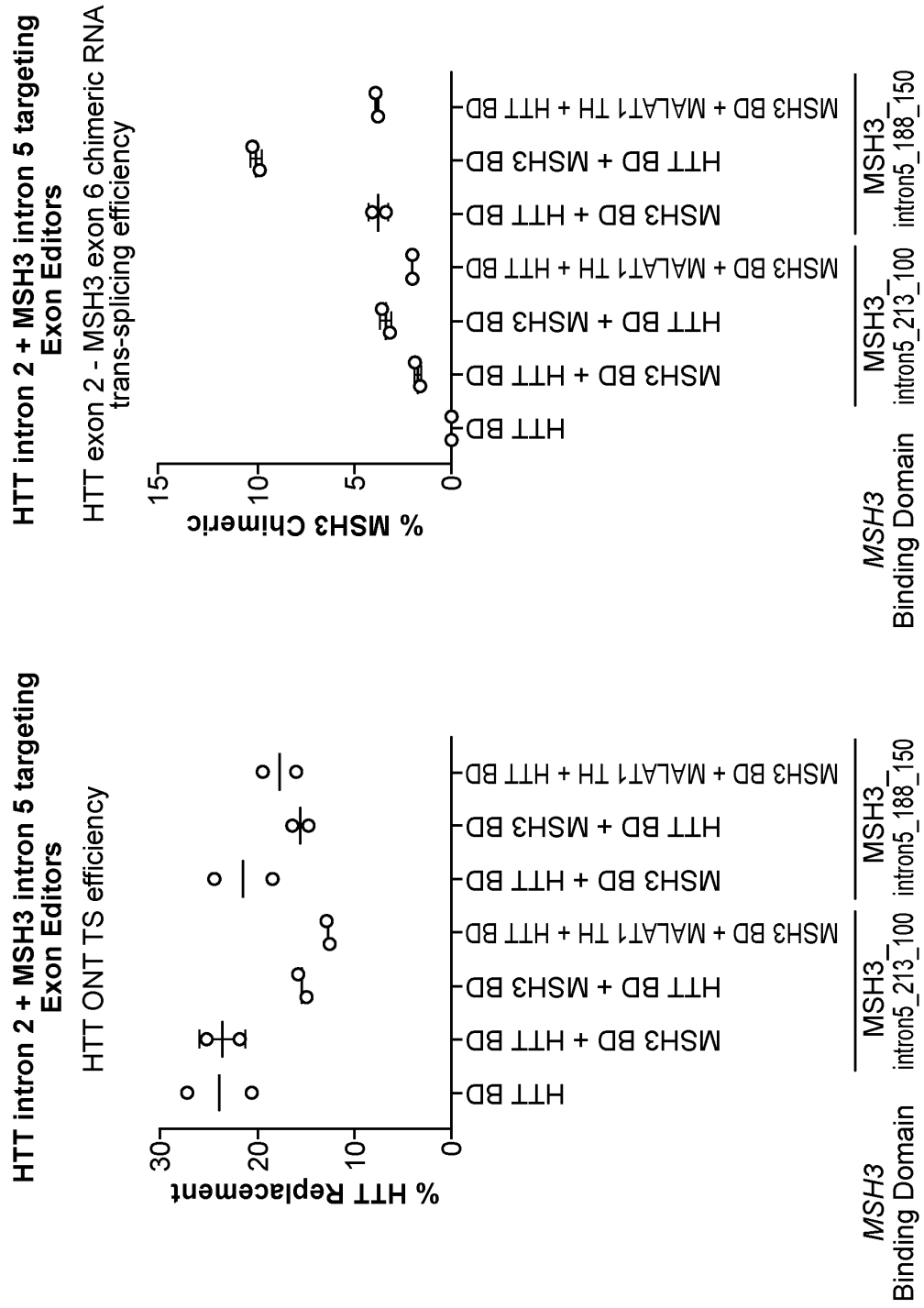


FIG. 30



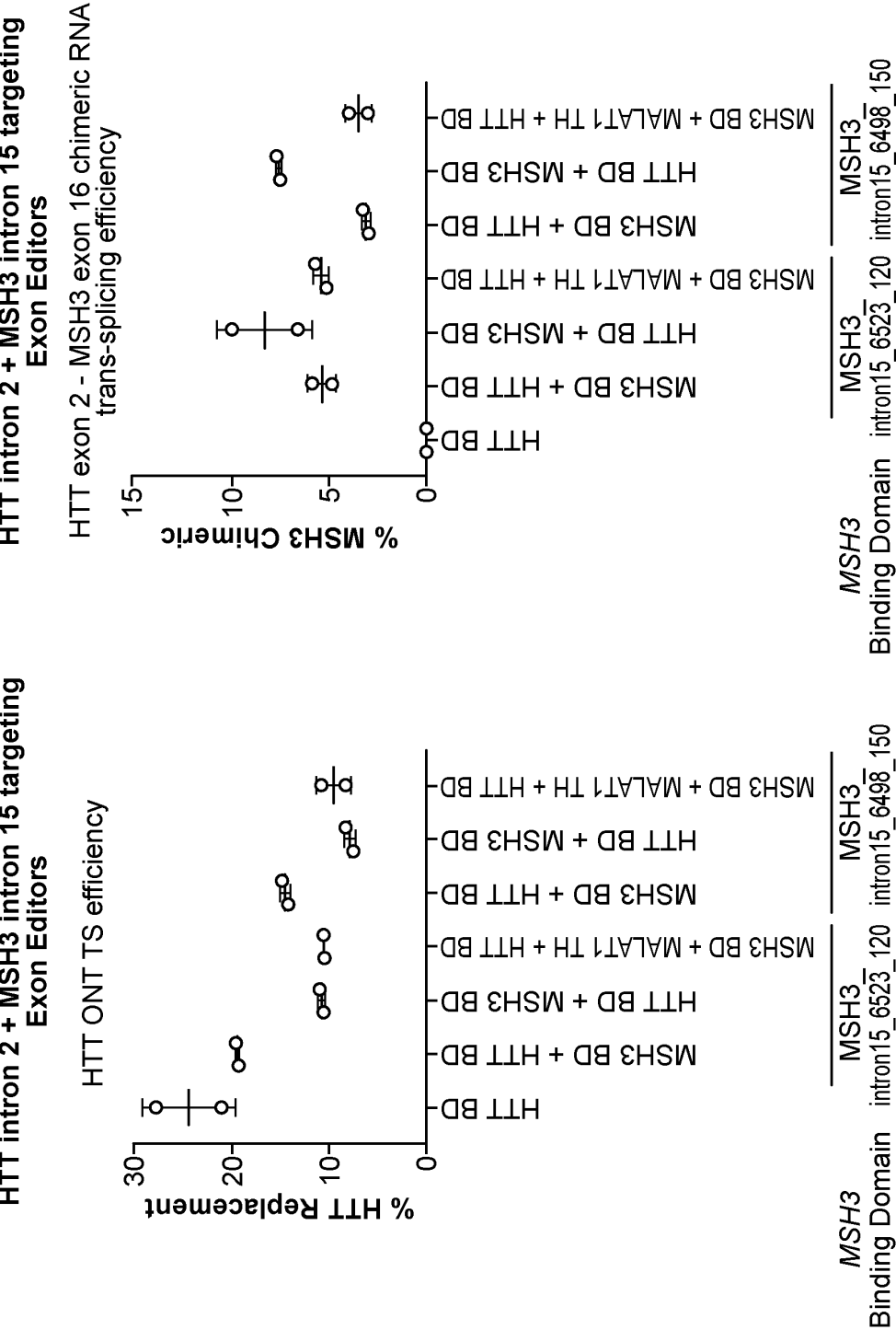


FIG. 32A

FIG. 32B

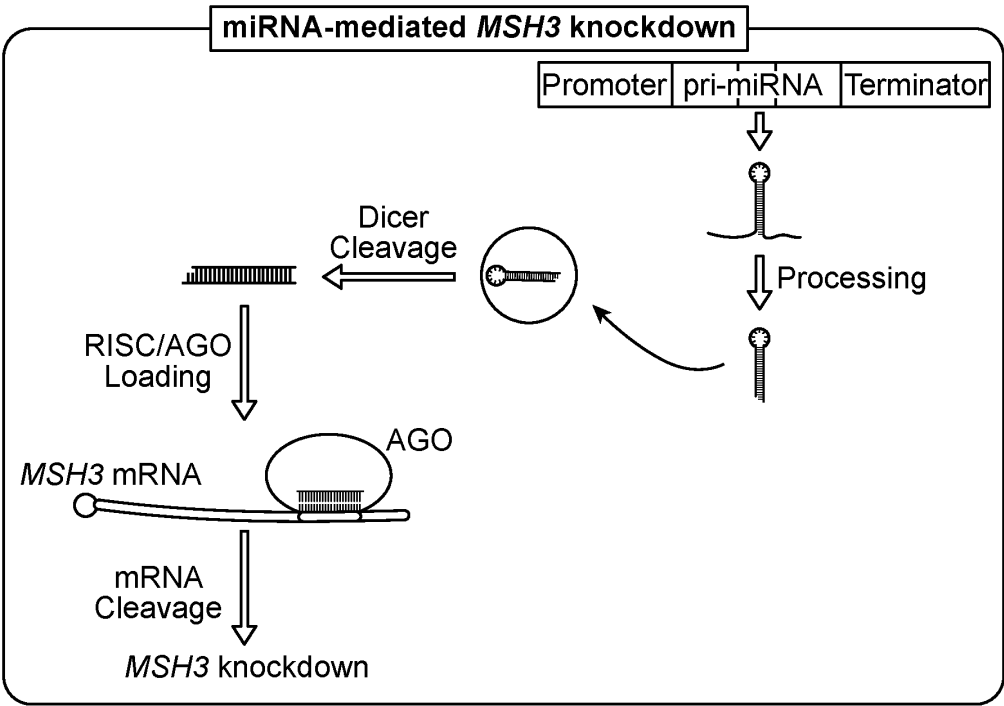


FIG. 33

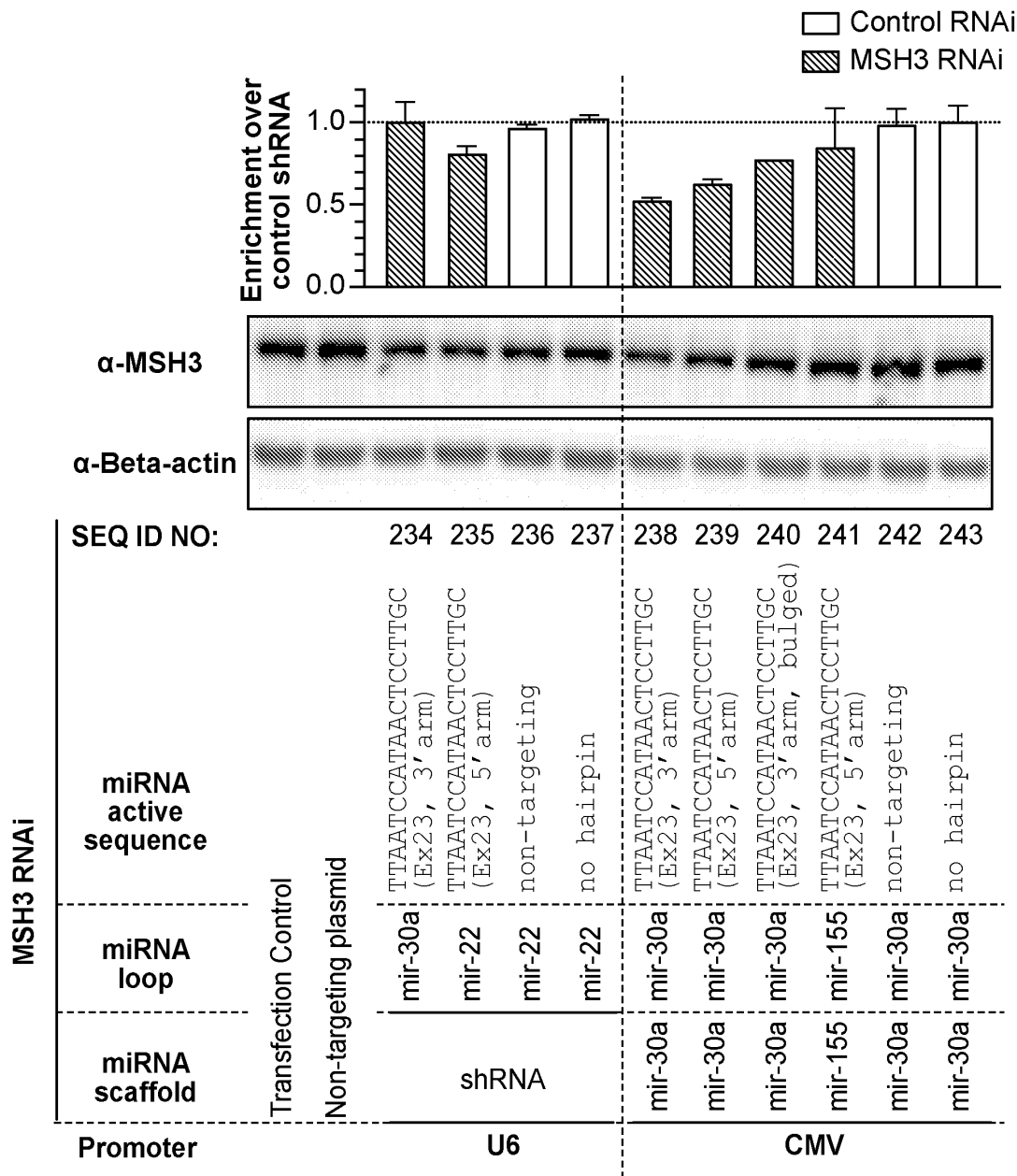


FIG. 34

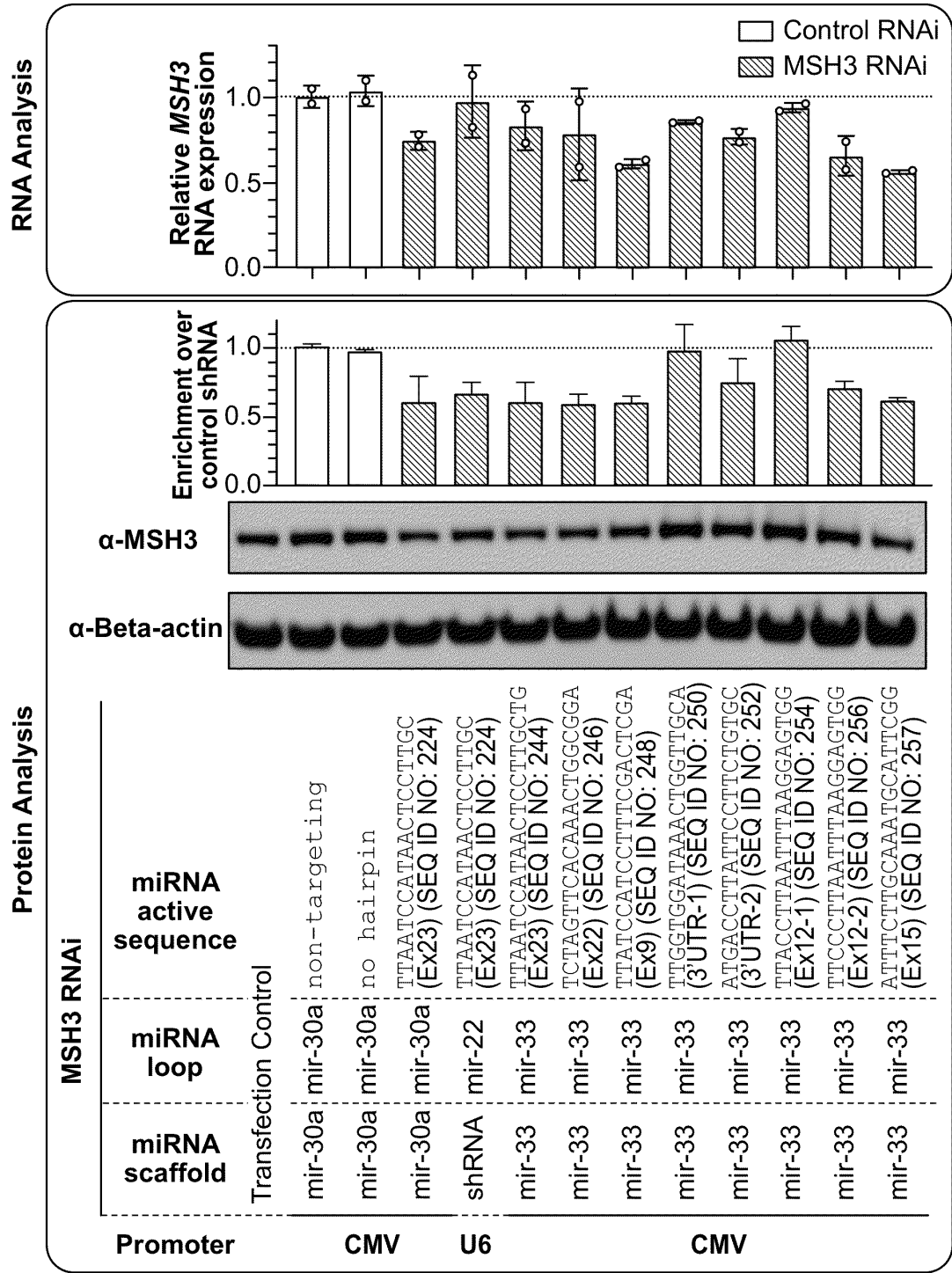


FIG. 35

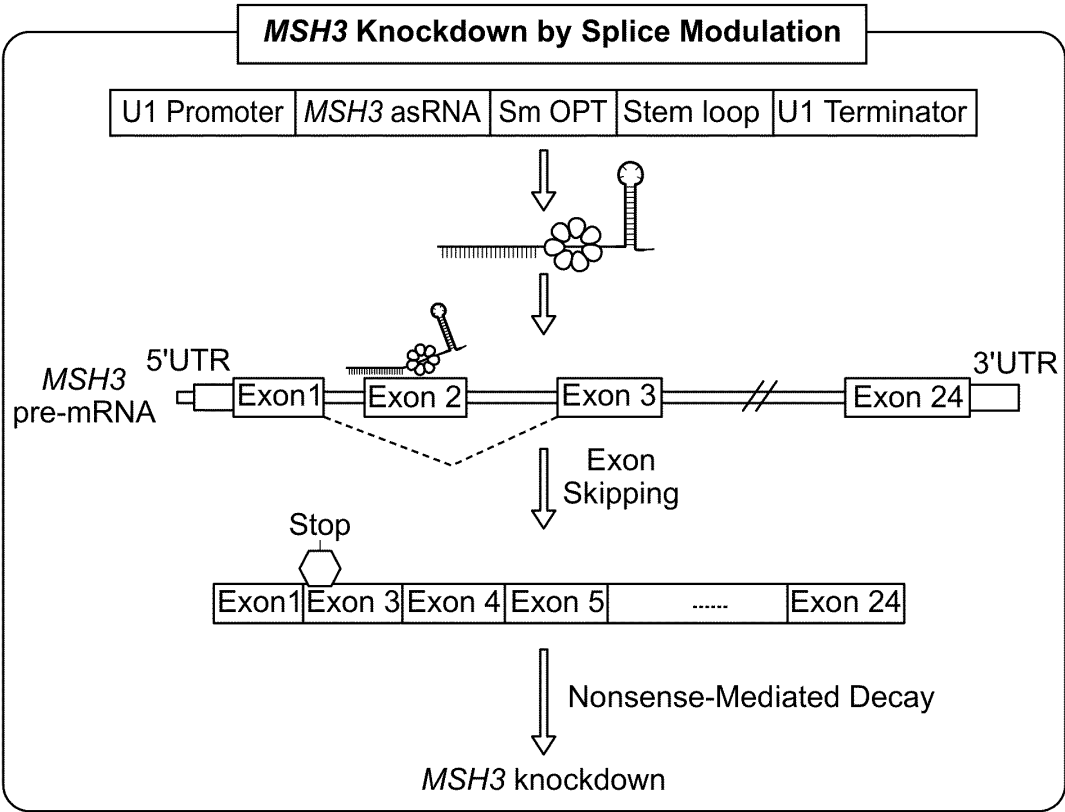


FIG. 36

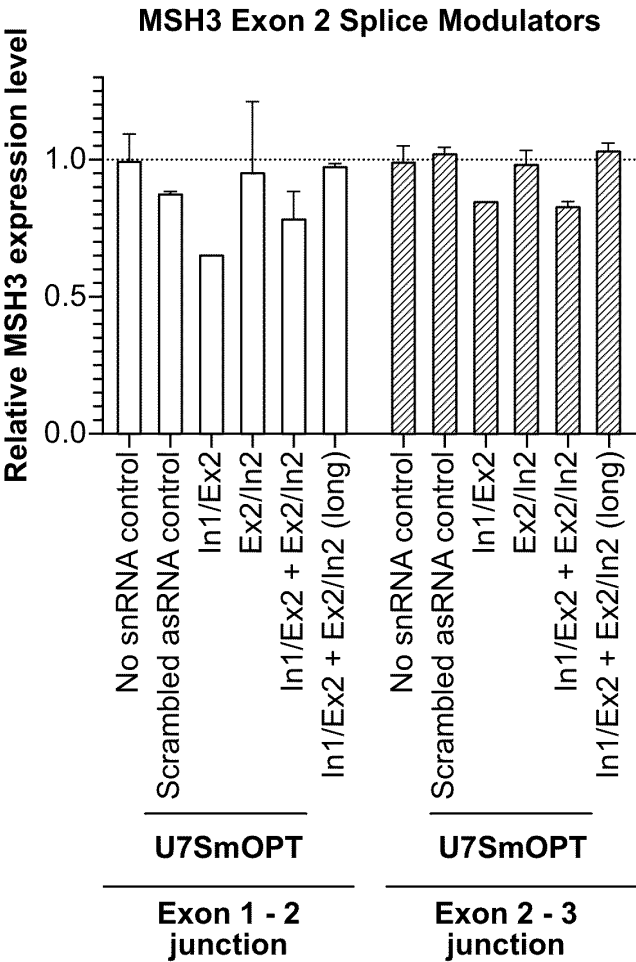
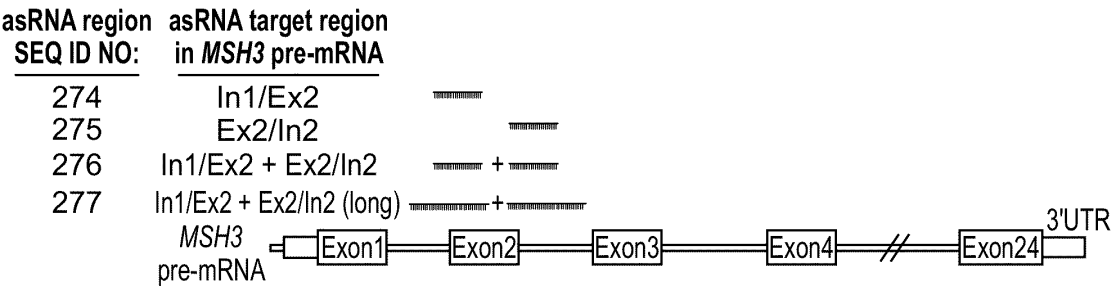


FIG. 37

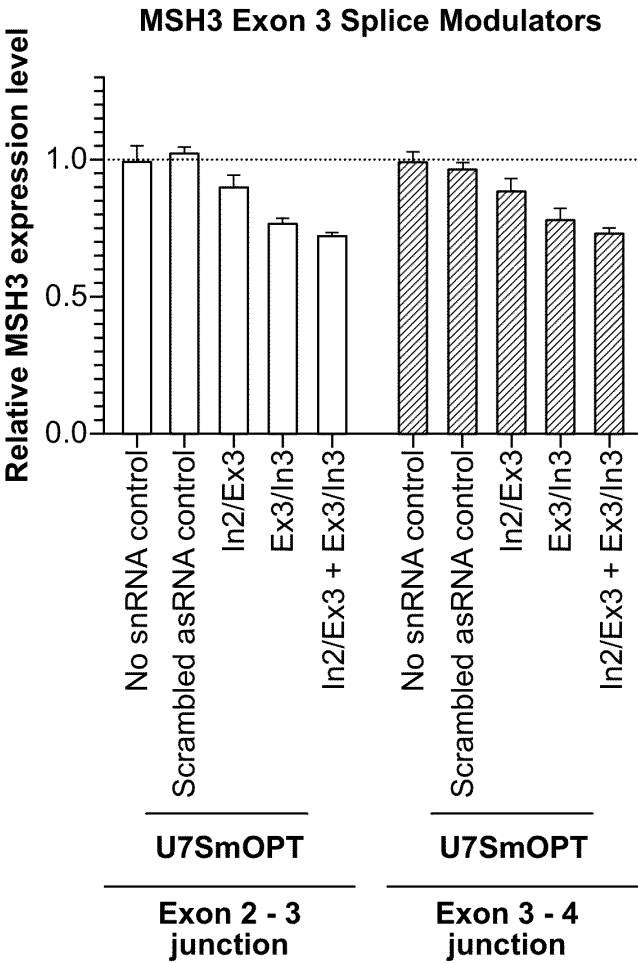
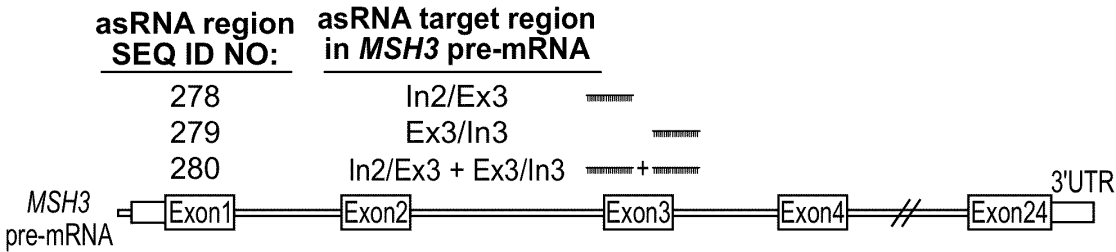


FIG. 38

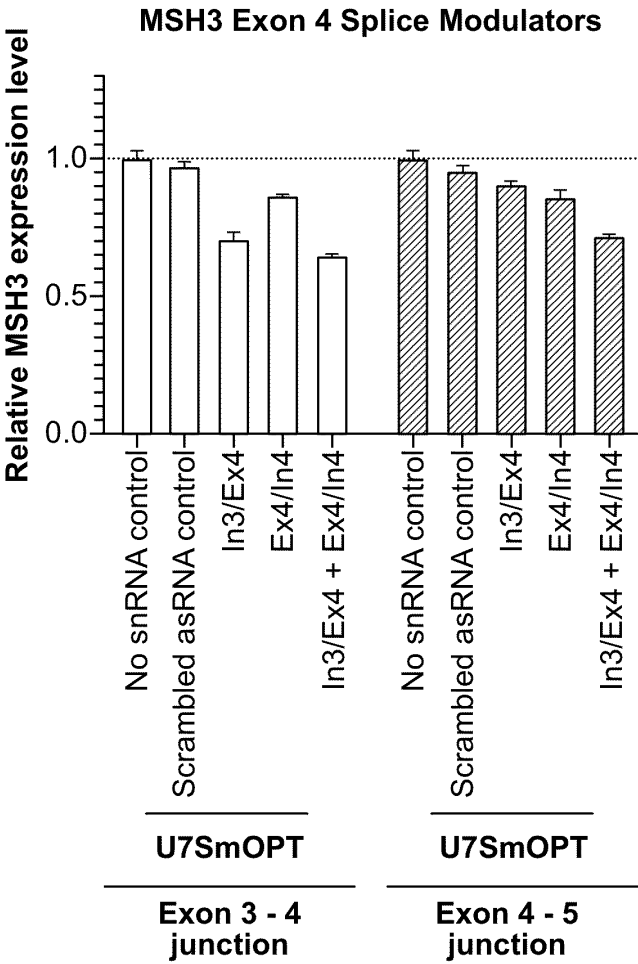
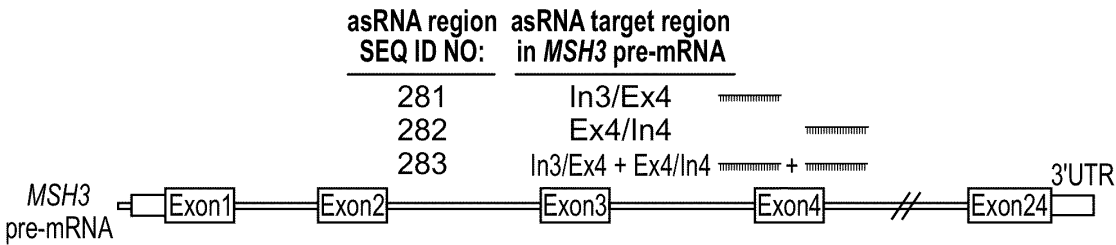


FIG. 39

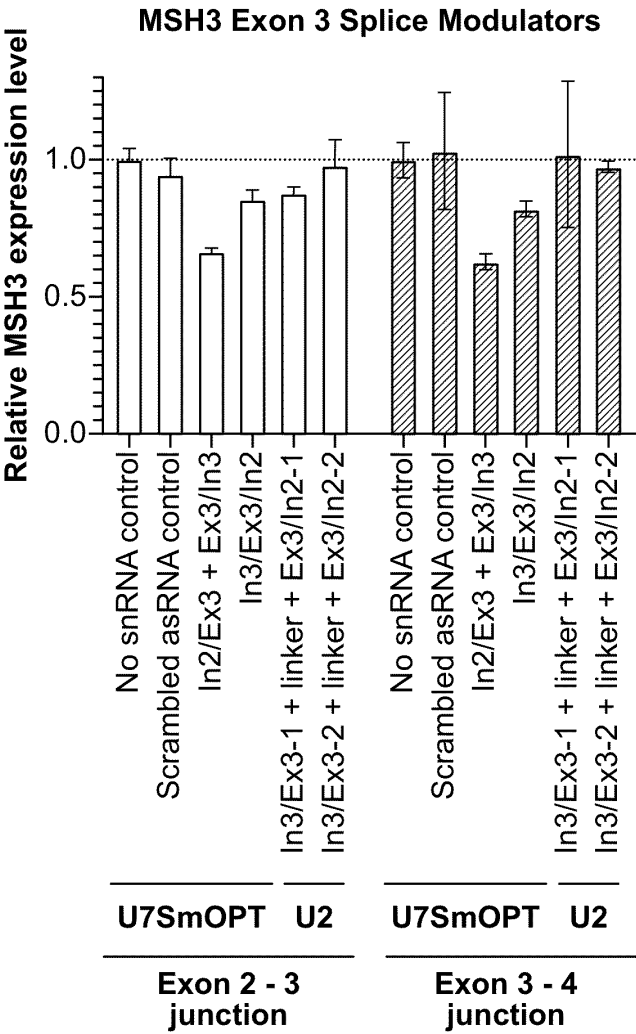
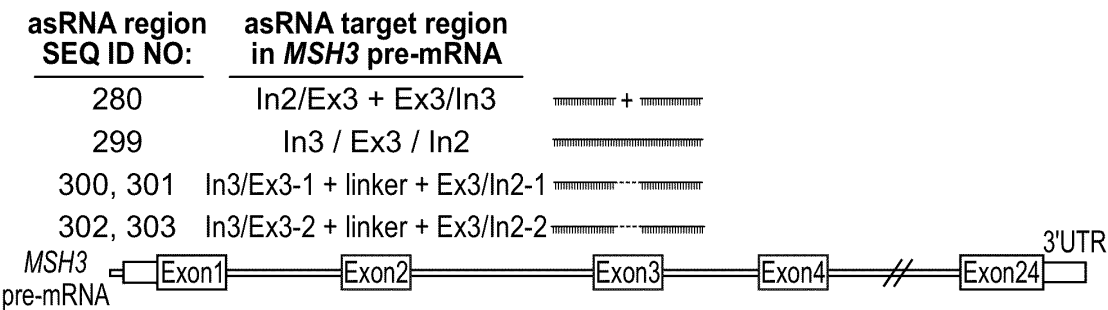
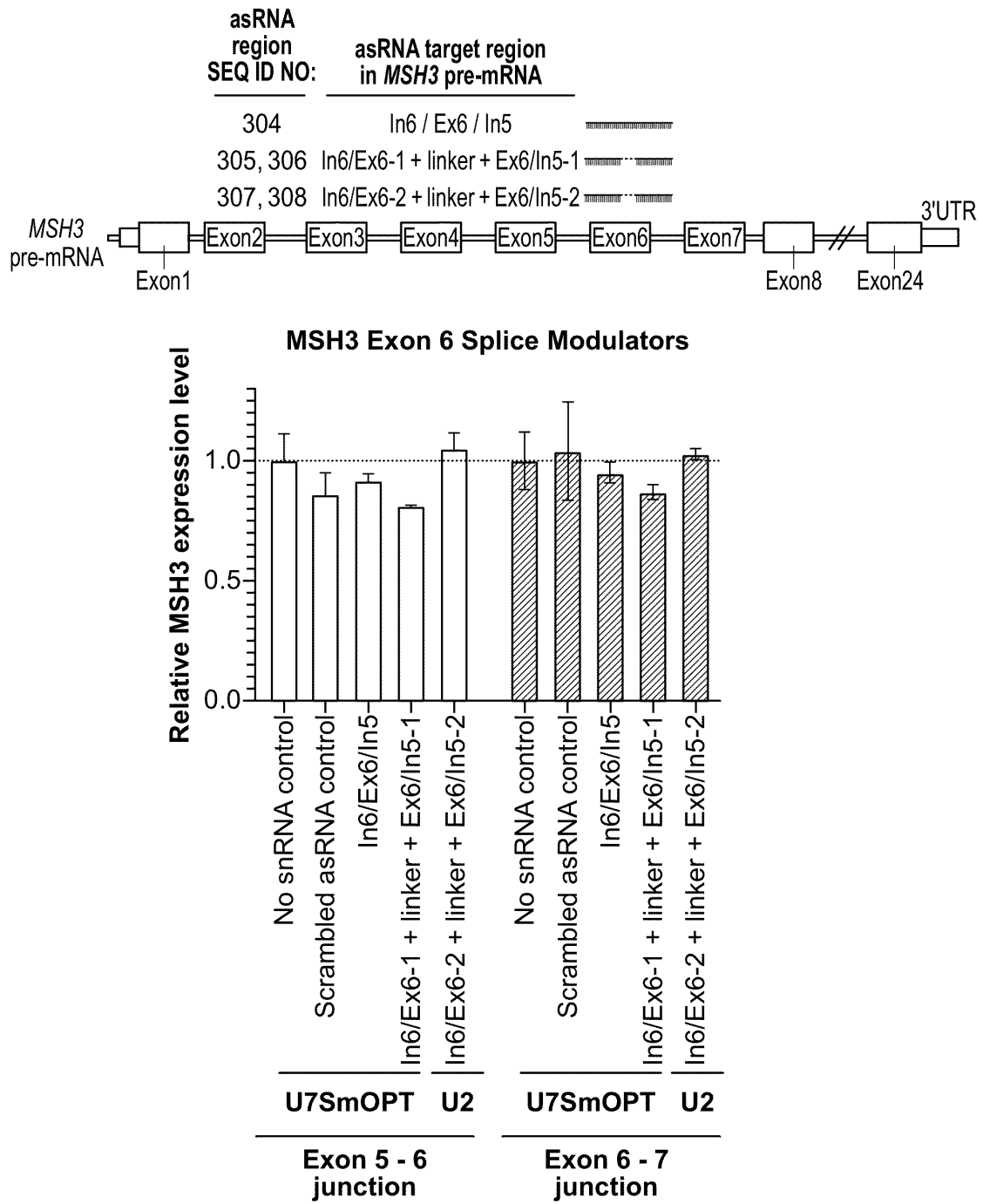
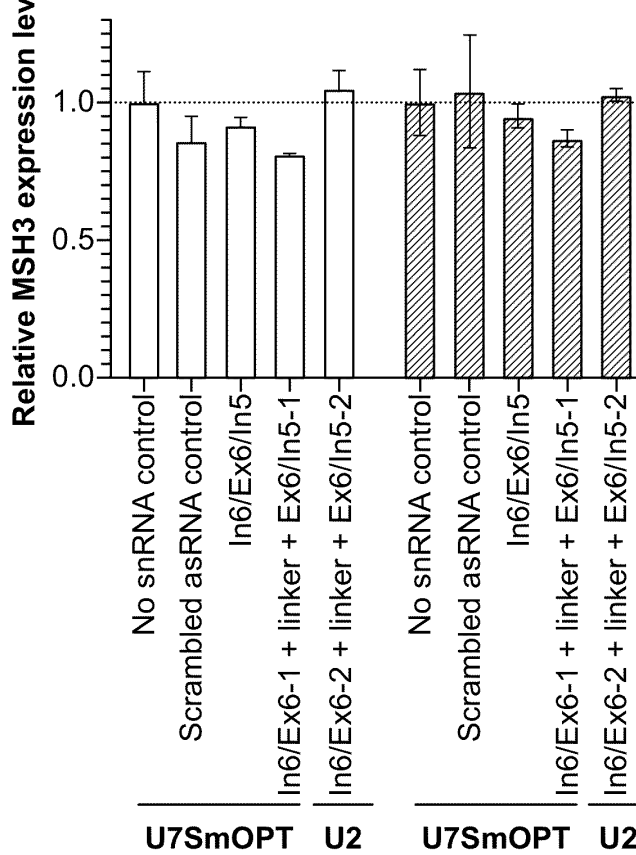


FIG. 40



Relative *MSH3* expression level



U7SmOPT

U2

U7SmOPT

U2

Exon 5 - 6
junction

Exon 6 - 7
junction

FIG. 41

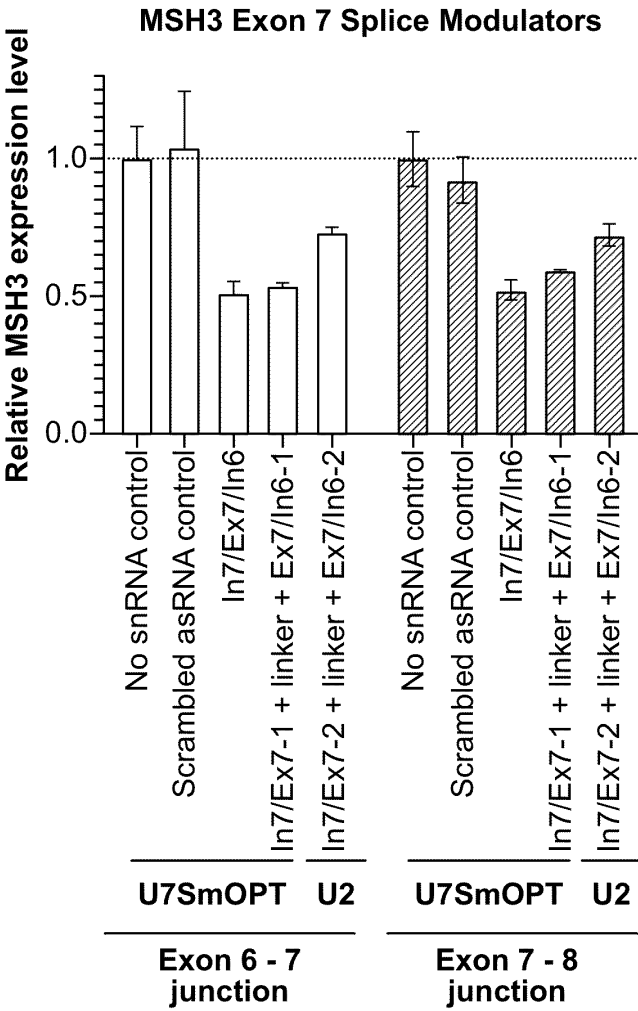
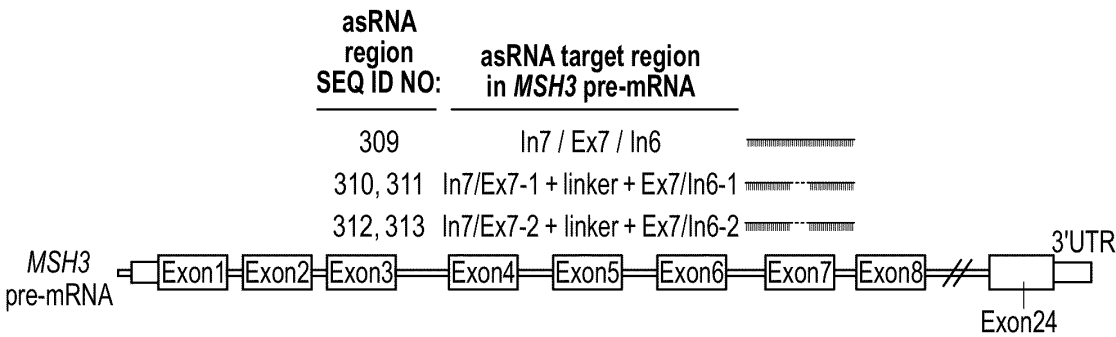


FIG. 42

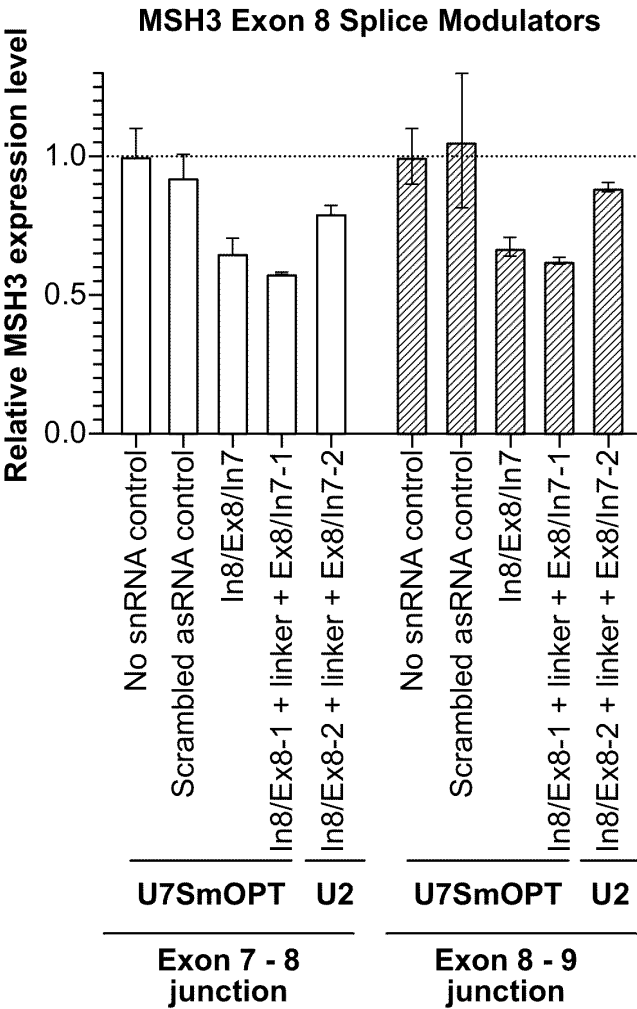
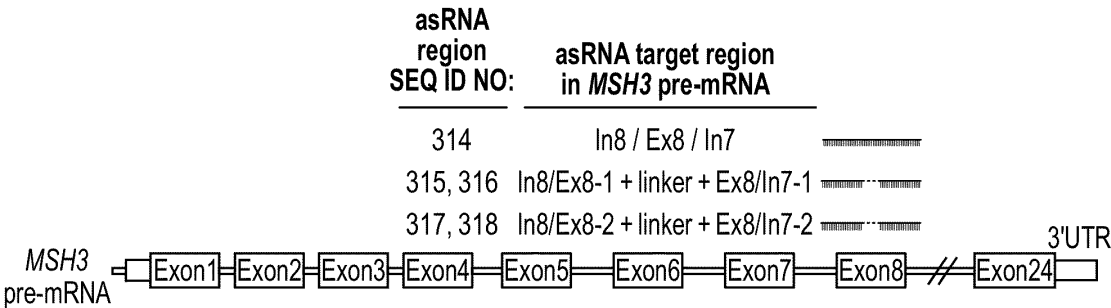


FIG. 43

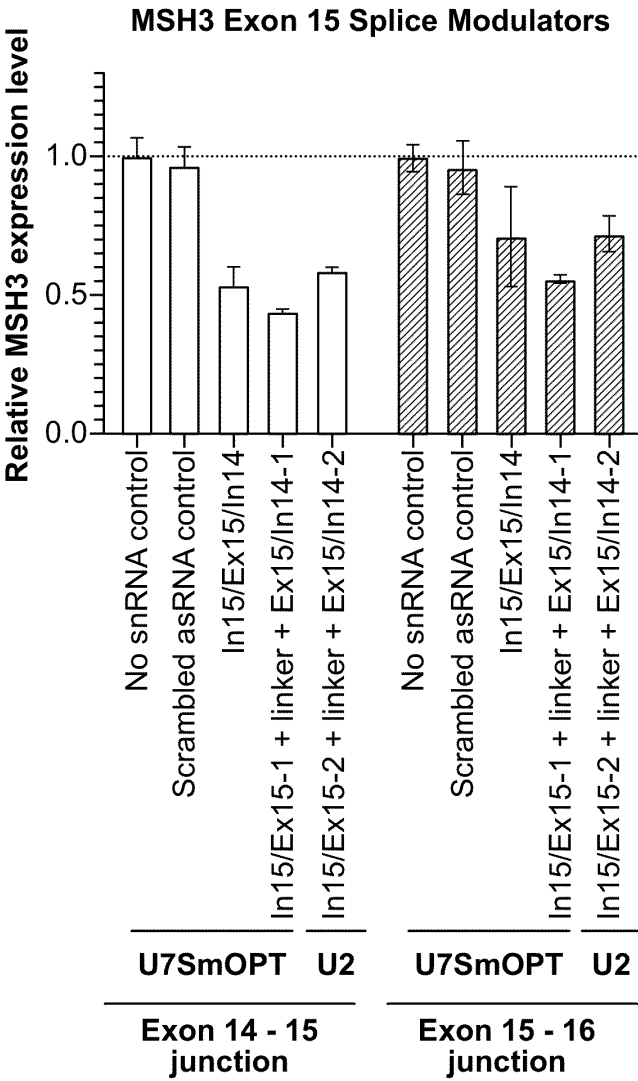
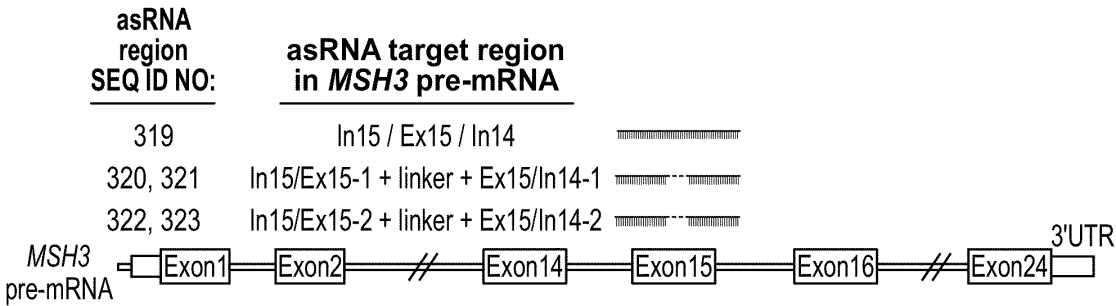


FIG. 44

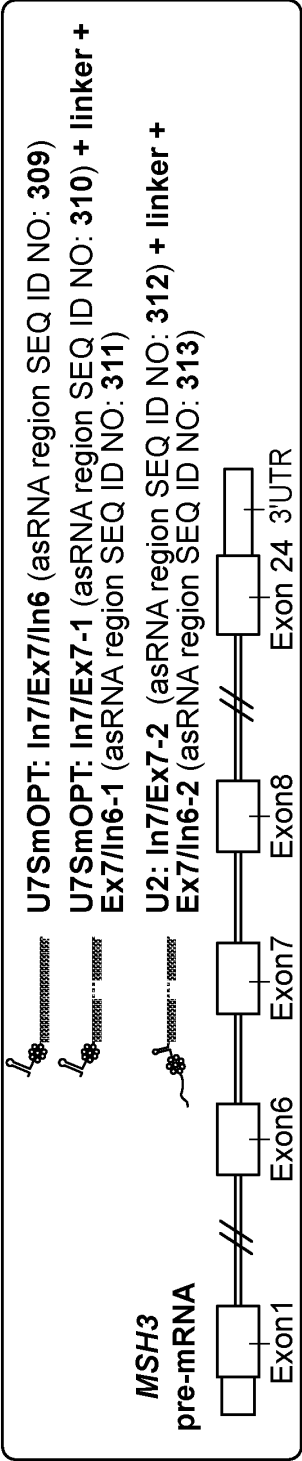


FIG. 45A

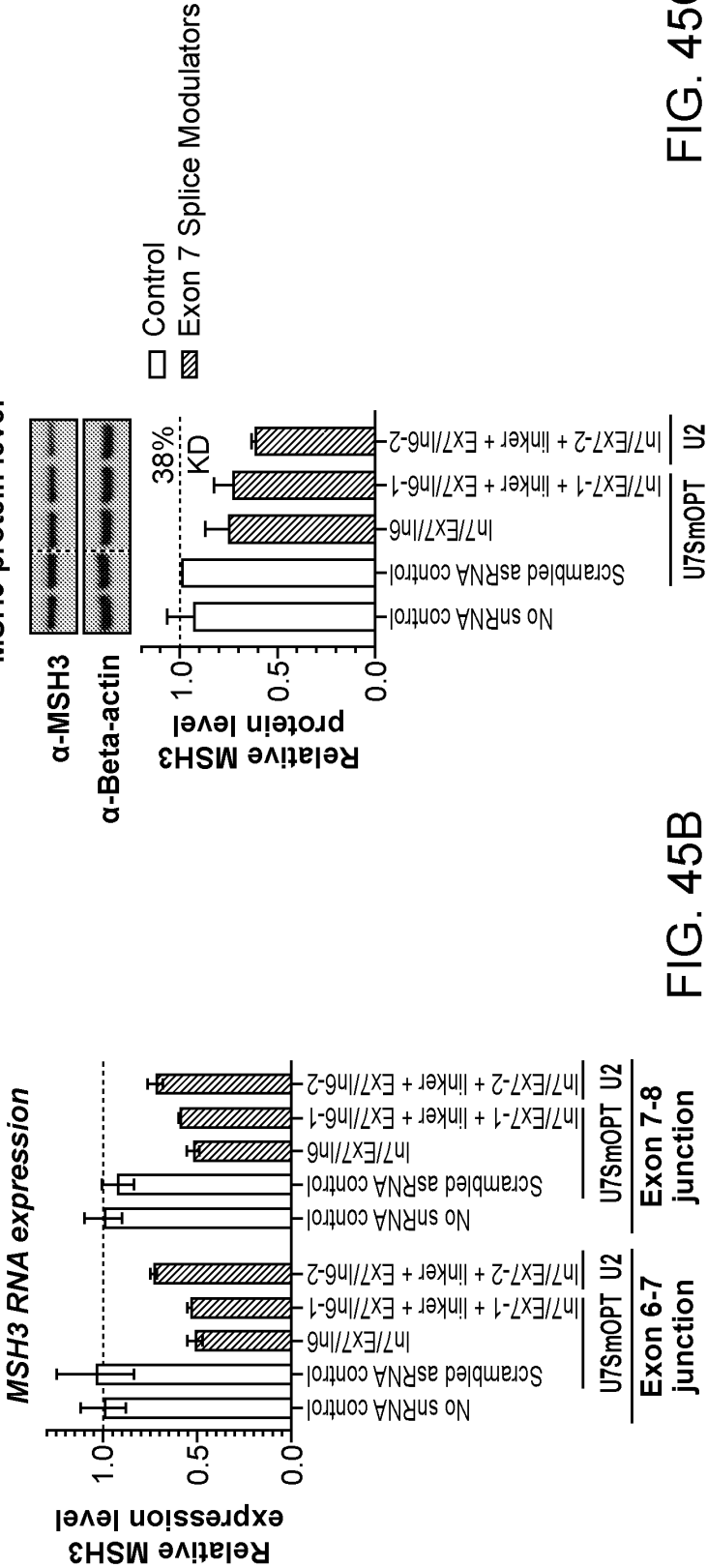


FIG. 45C

FIG. 45B

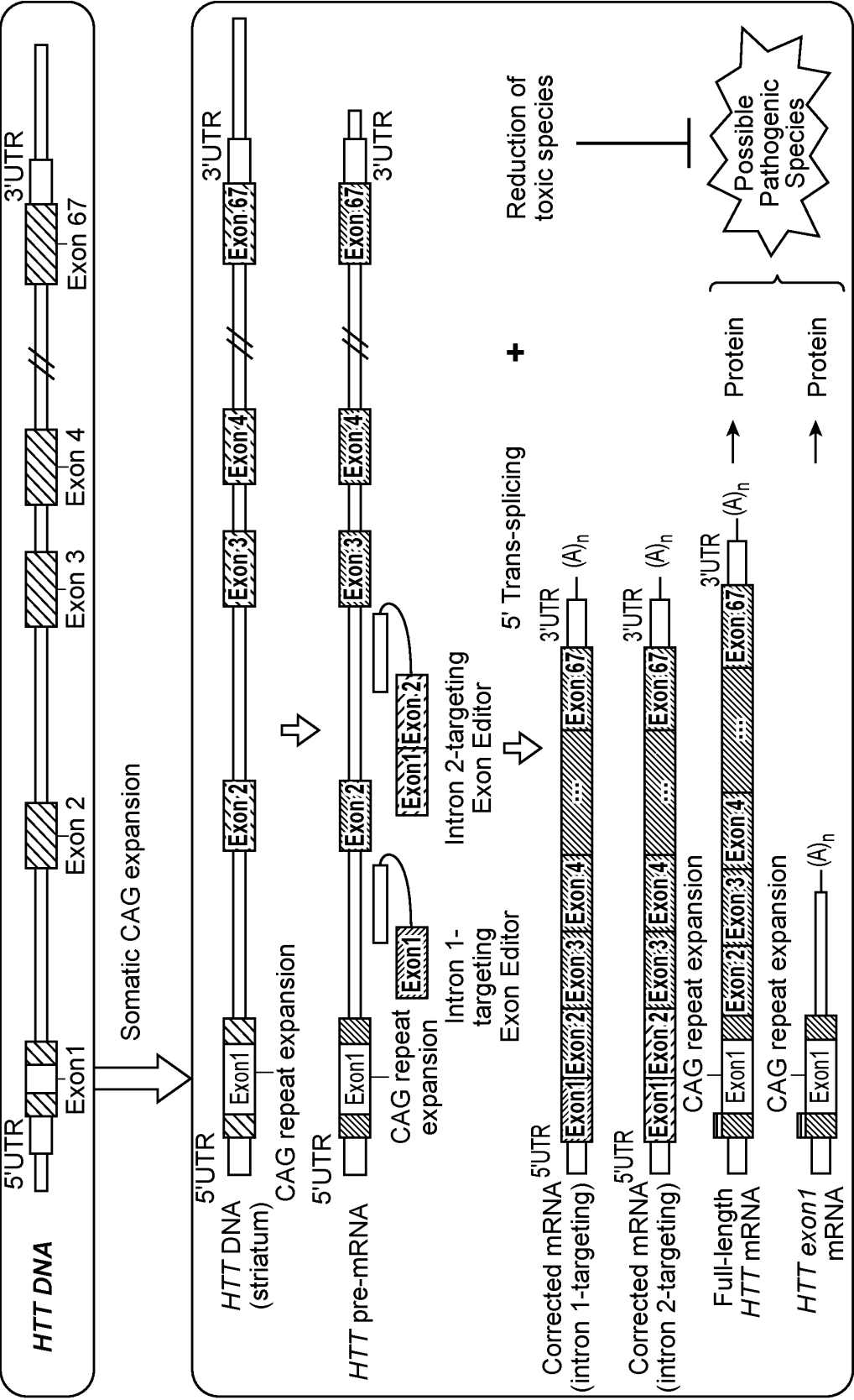


FIG. 46

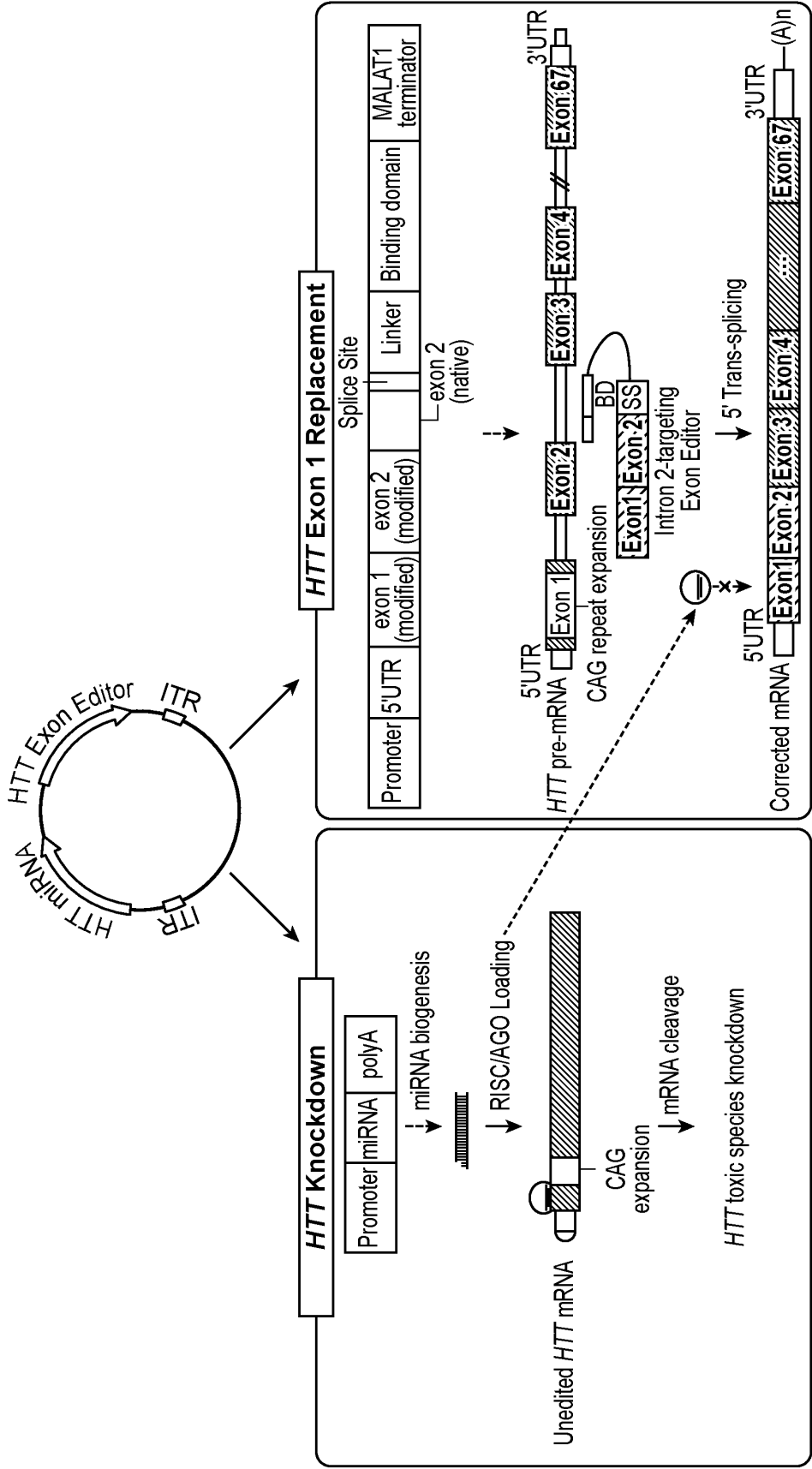


FIG. 47

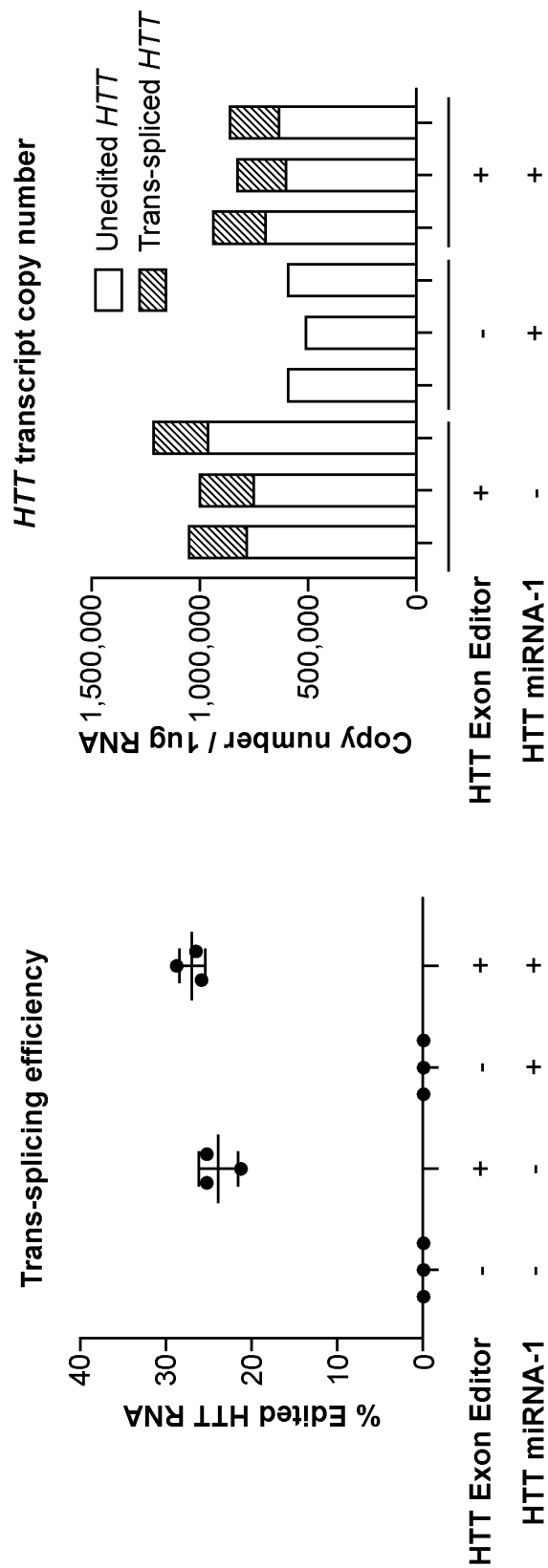


FIG. 48A

FIG. 48B

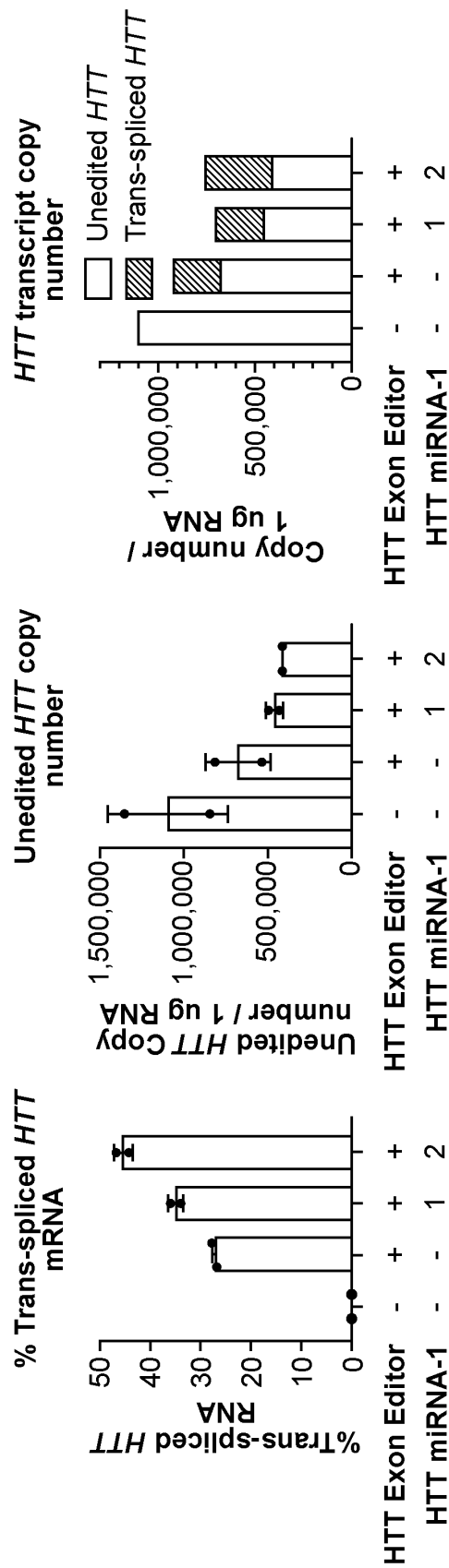


FIG. 49A

FIG. 49B

FIG. 49C

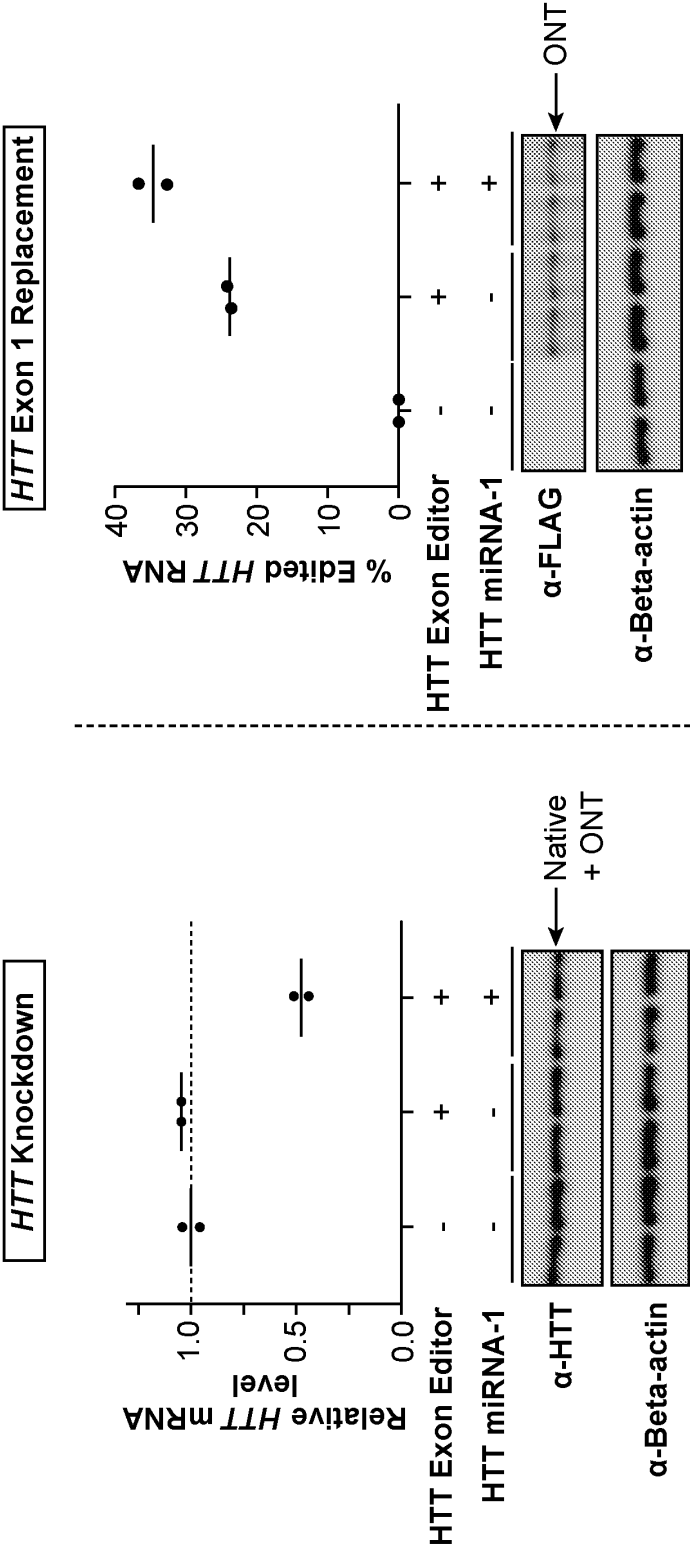


FIG. 50

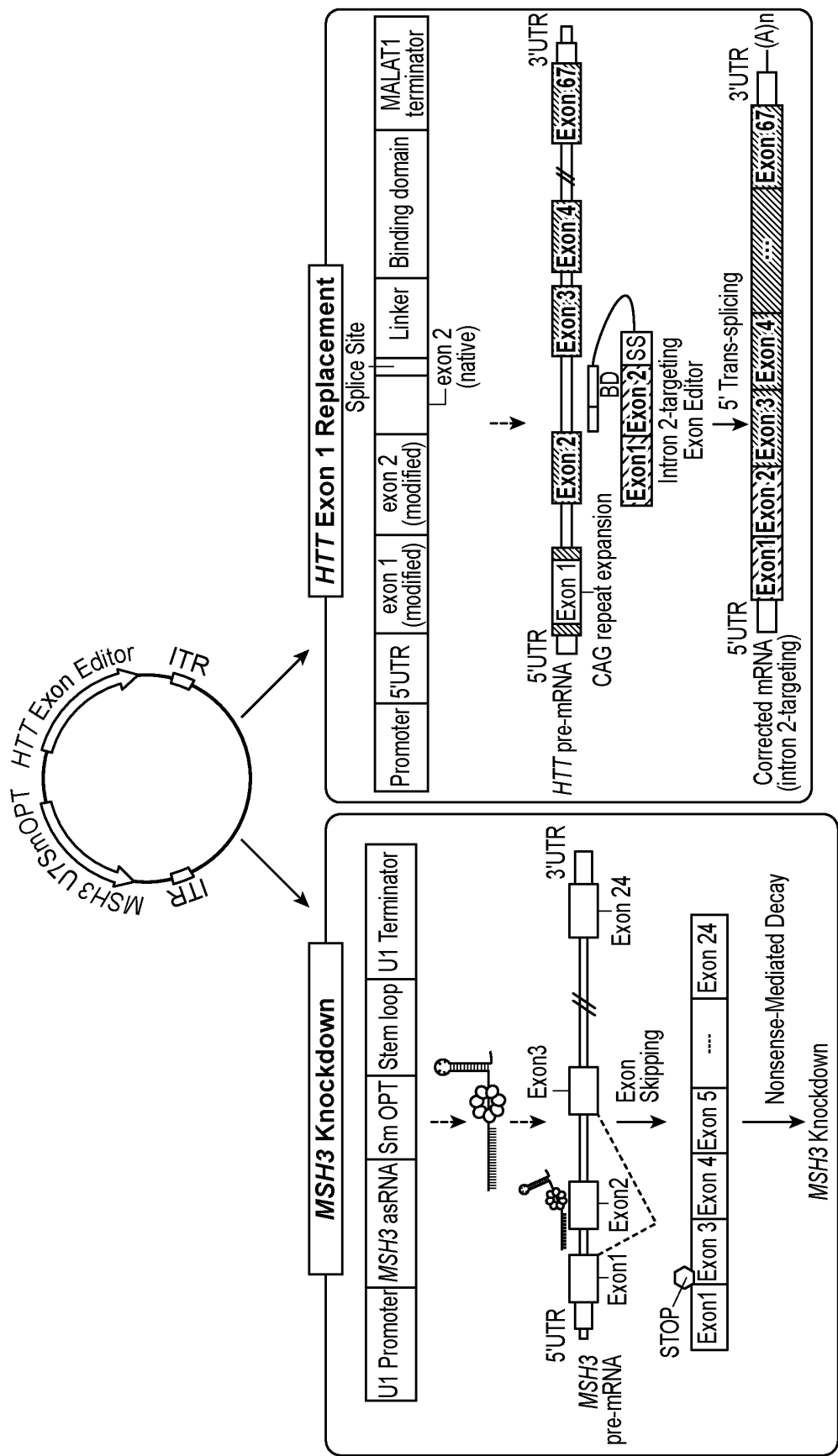


FIG. 51

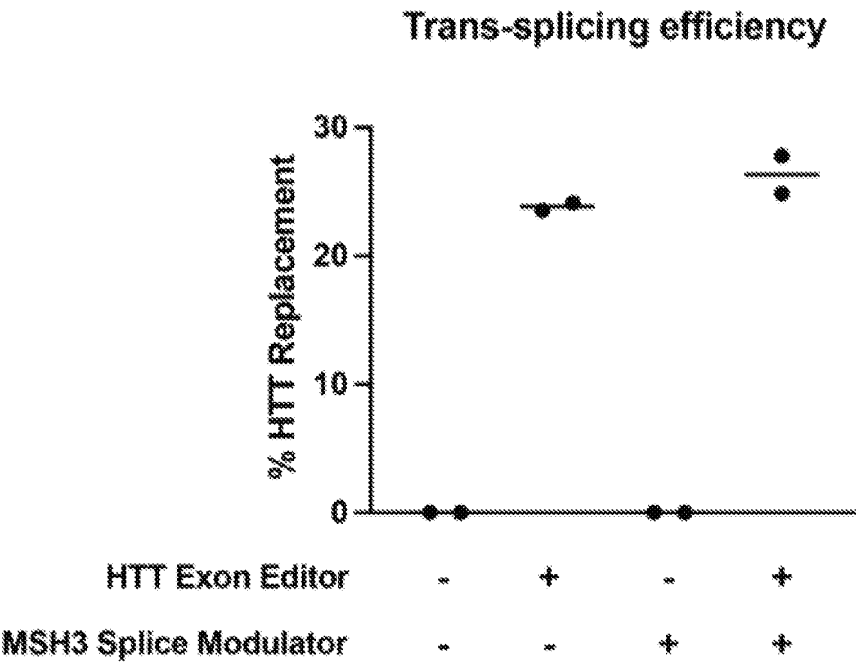


FIG. 52A

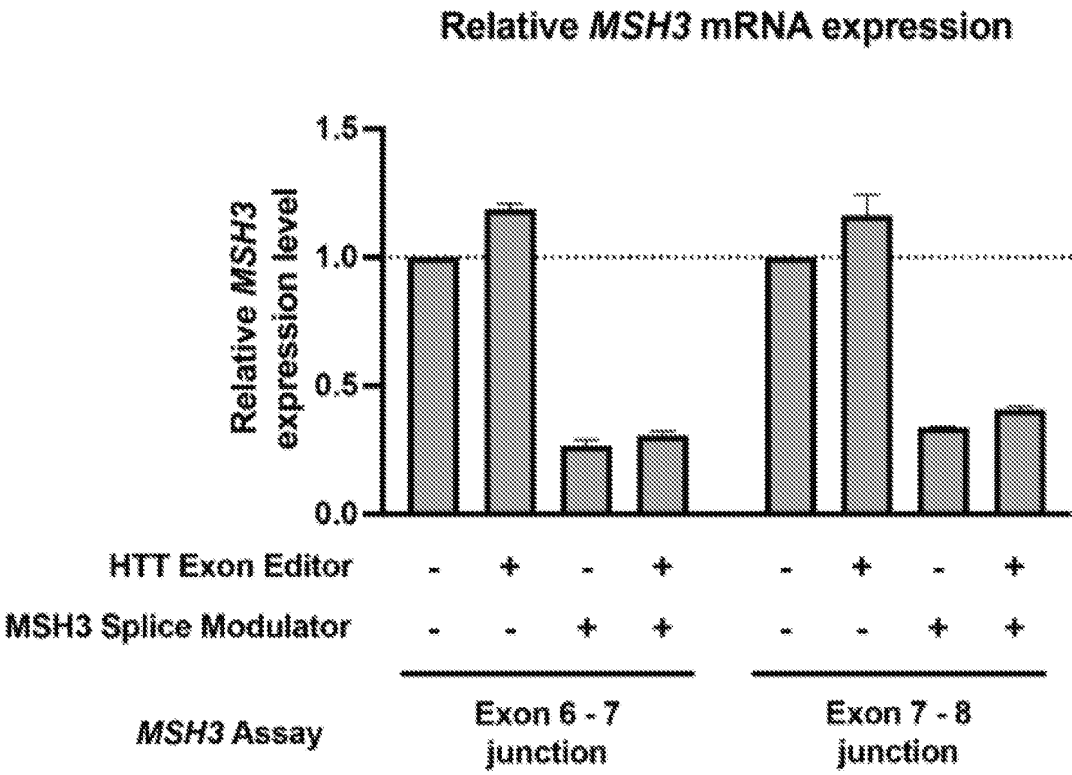
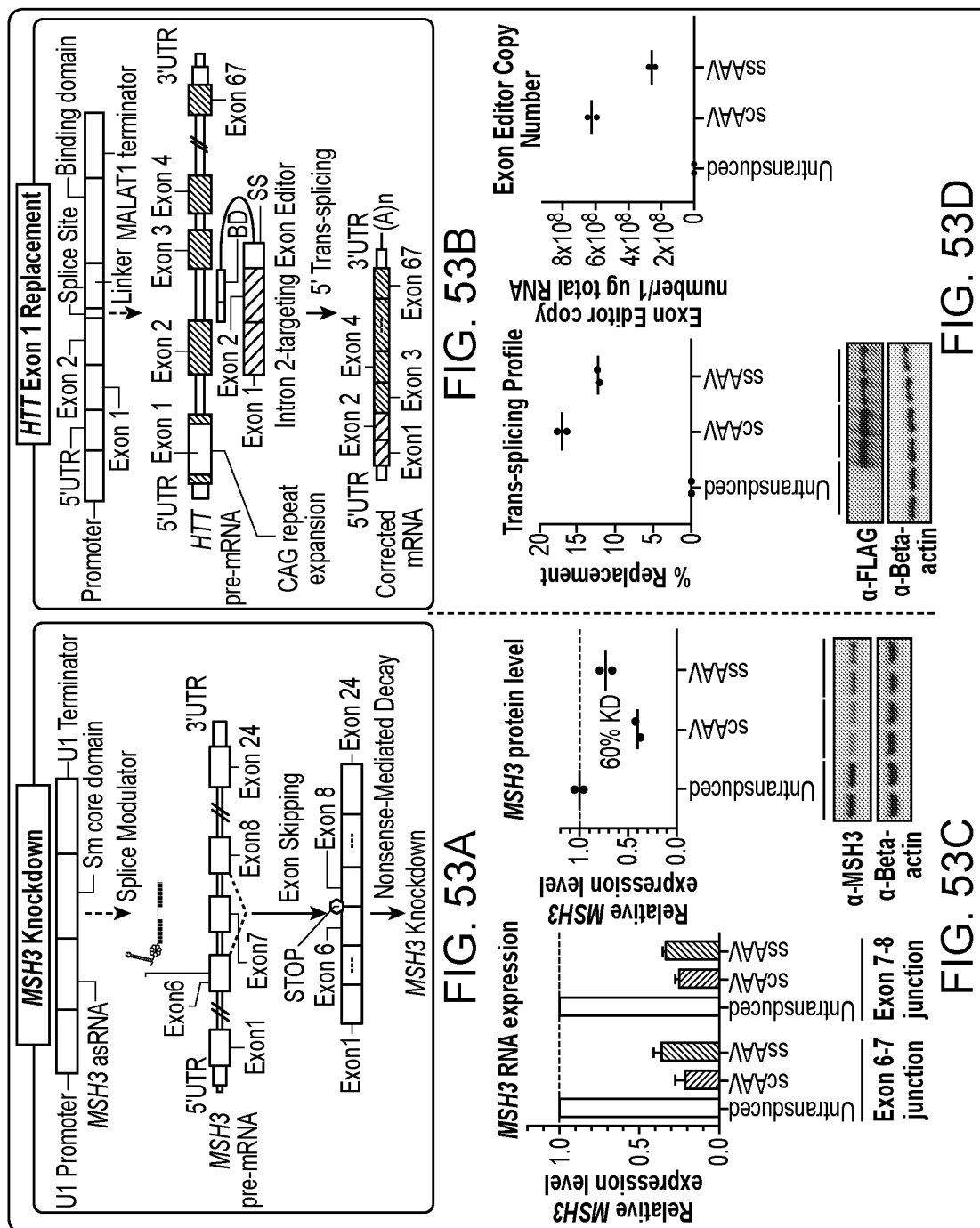
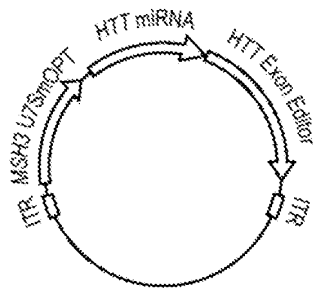


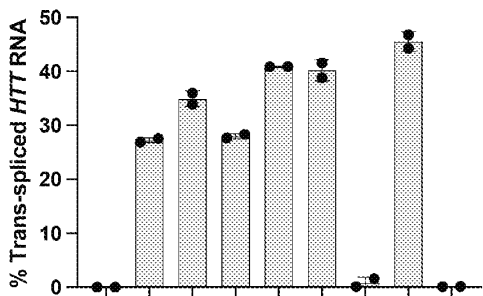
FIG. 52B



50 / 56

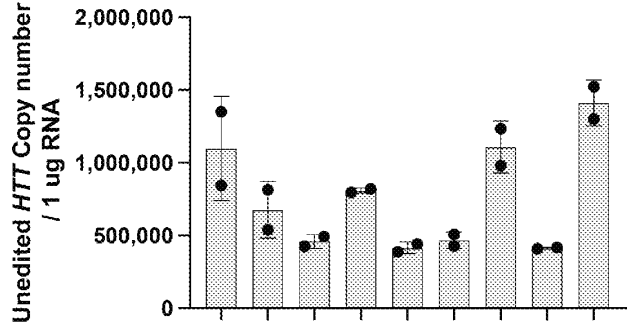


% Trans-spliced *HTT* mRNA



HTT Exon Editor	-	+	+	+	+	+	SM	+	SM
HTT miRNA	-	-	1	-	1	2	Scr	2	Scr
MSH3 Splice Modulator	-	-	-	+	+	+	Scr	-	-

Unedited *HTT* copy number



HTT Exon Editor	-	+	+	+	+	+	SM	+	SM
HTT miRNA	-	-	1	-	1	2	Scr	2	Scr
MSH3 Splice Modulator	-	-	-	+	+	+	Scr	-	-

FIG. 54

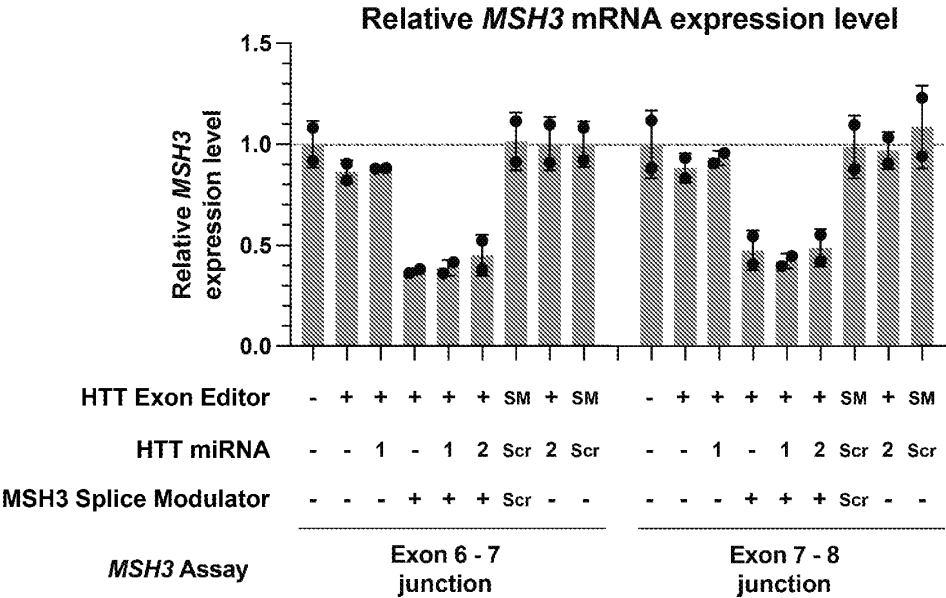


FIG. 54 (Cont.)

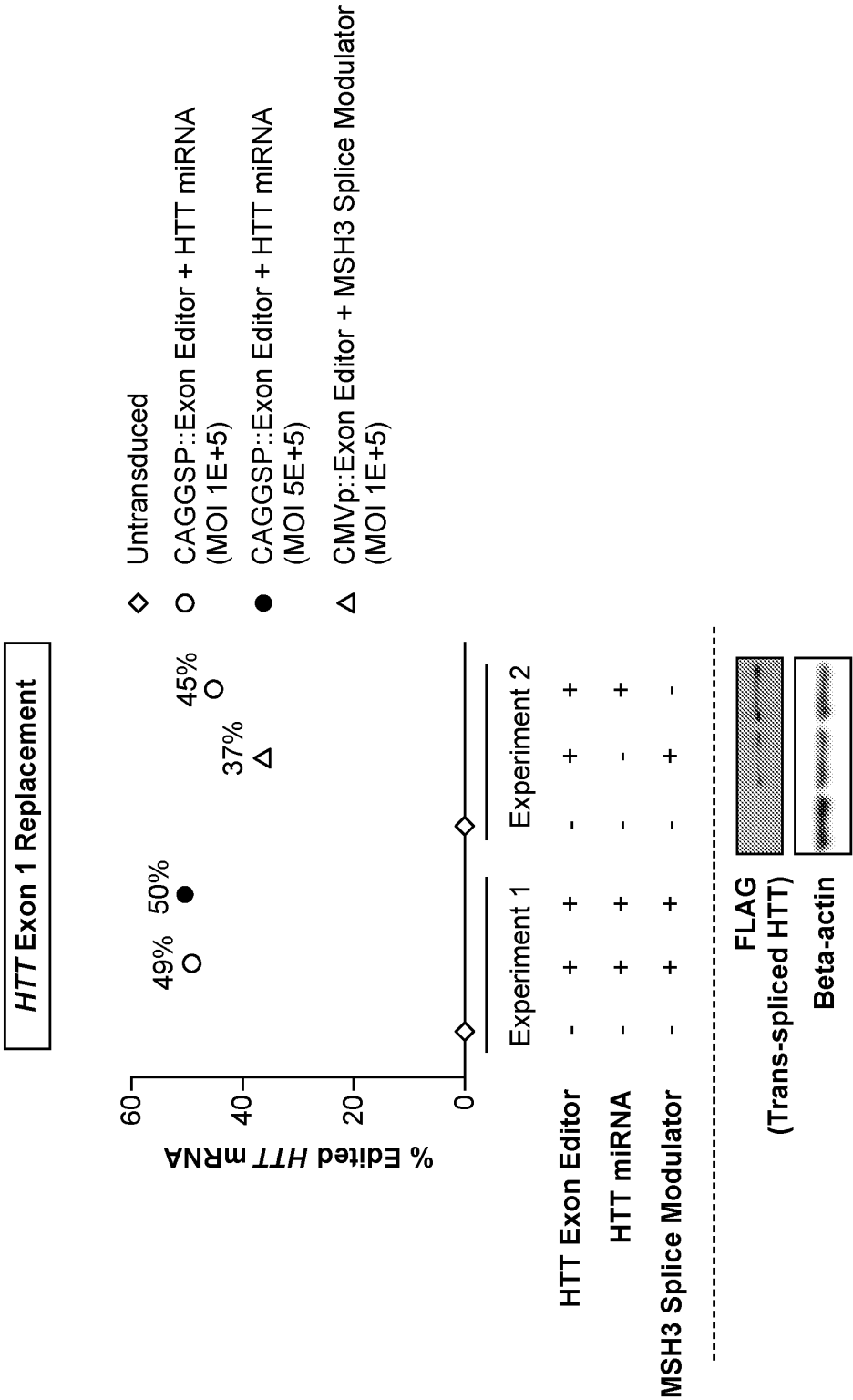


FIG. 55

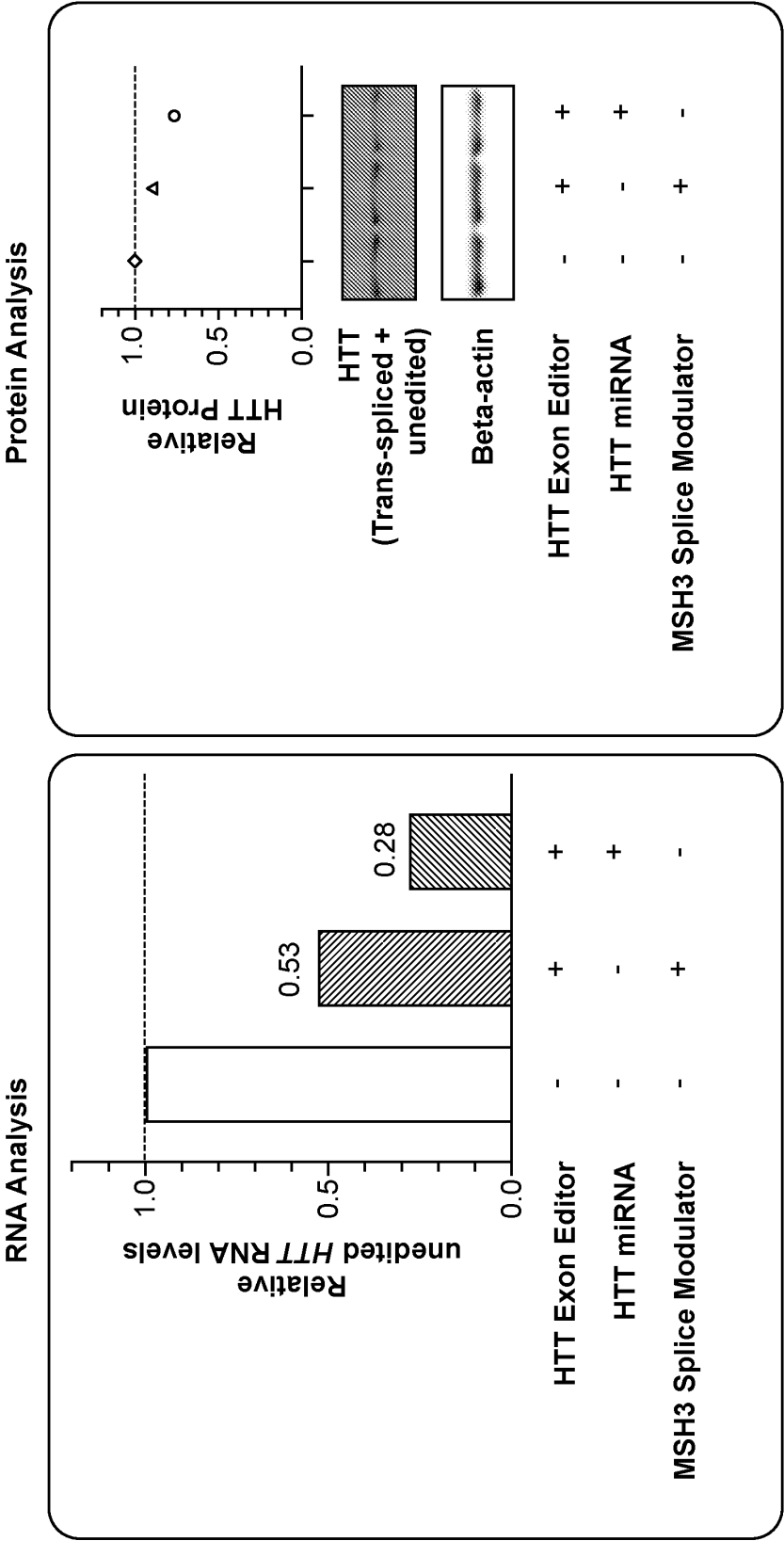


FIG. 56

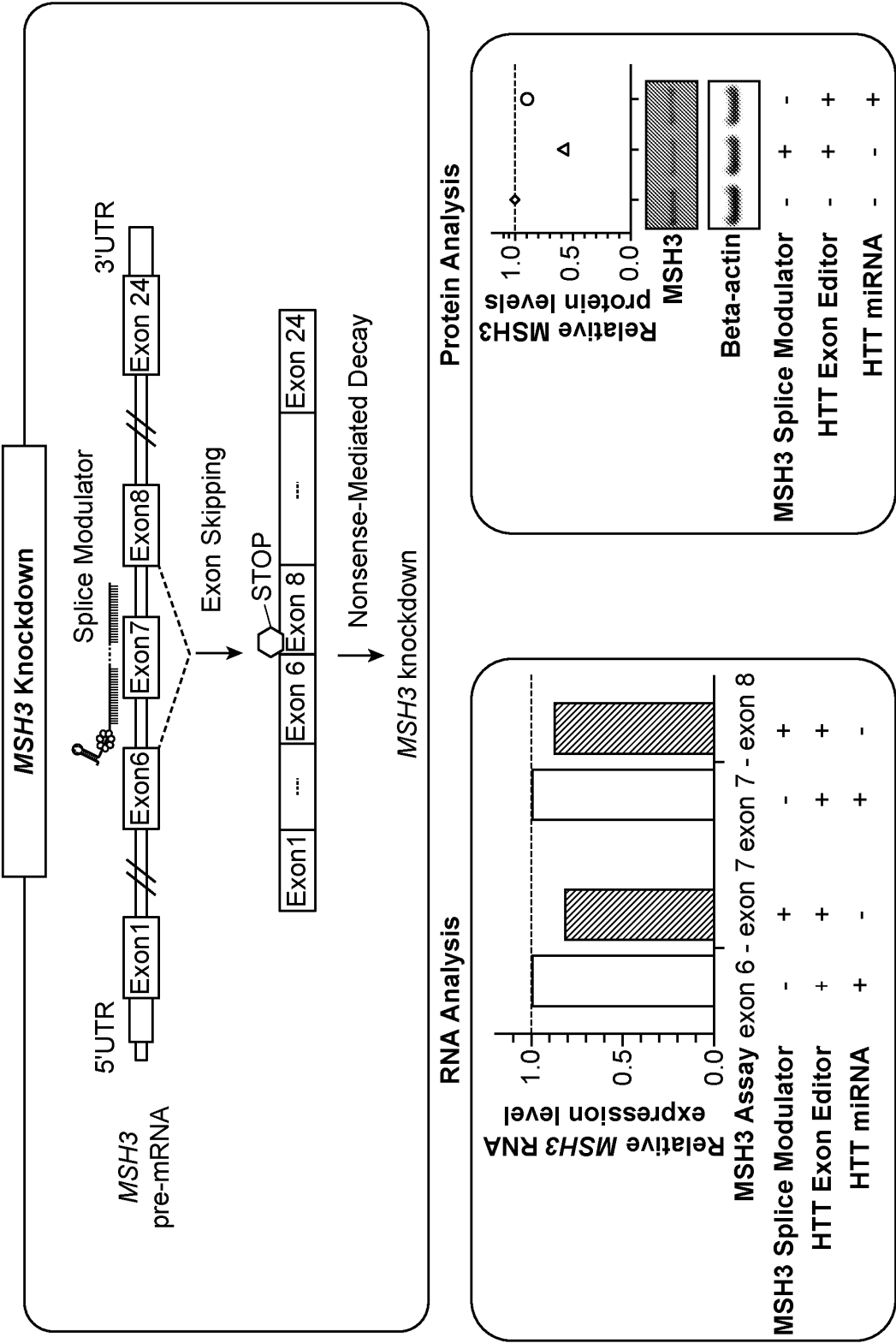


FIG. 57

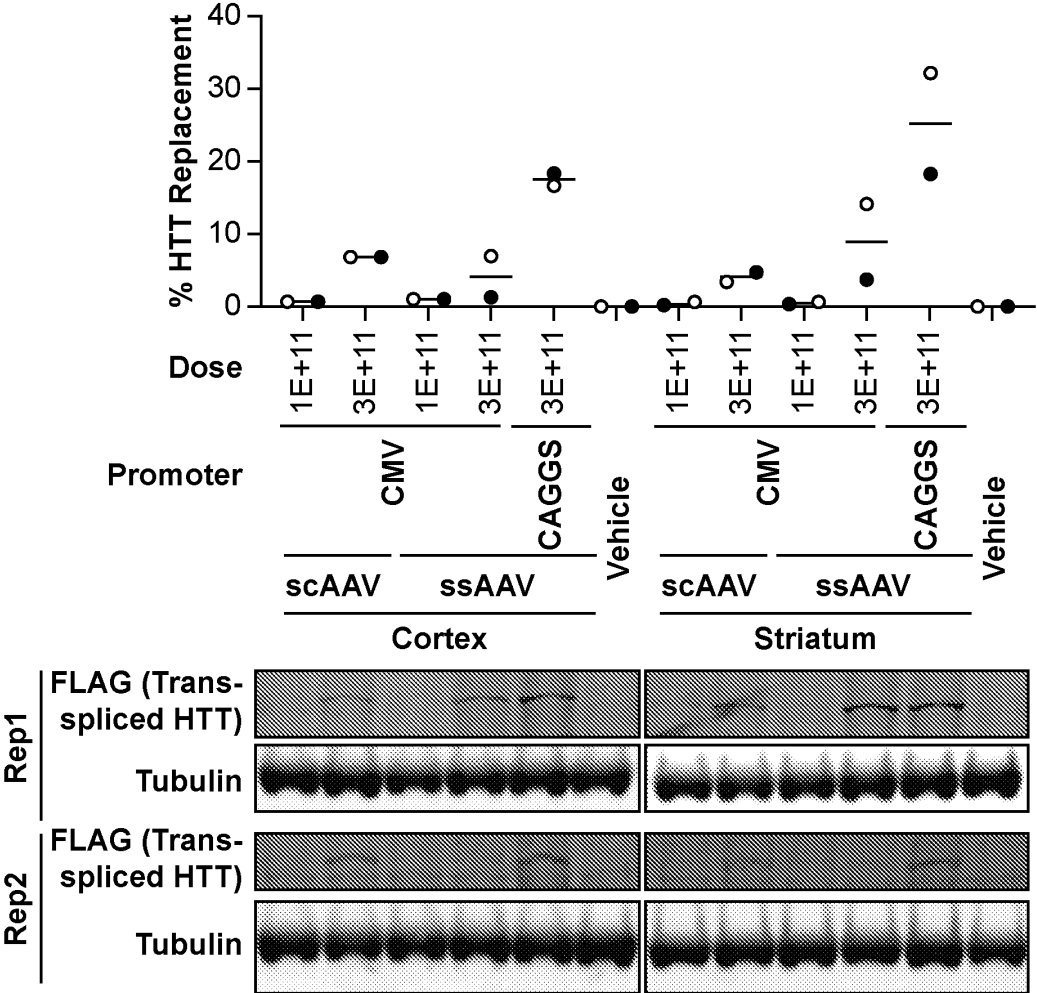


FIG. 58

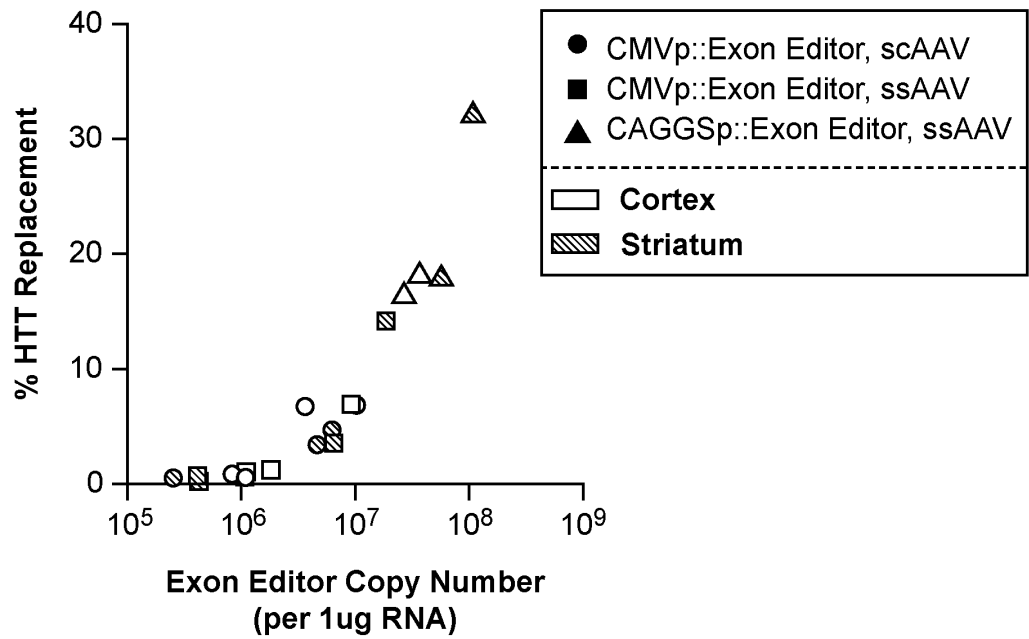


FIG. 59