Variant position in UCSC-based canonical isoform OTR Exon Intron 5' UTR 3' UTR 1 14000 2 12226 12000 9373 10000 8000 3 6000 5 4 4000 1308 2000 192 324 291 322 291 110 22 88 22 92 187 74 32 12 44 0 v5-smentonyunber vs. sanetonnumber vs diffsontumber , diffinition winder VS. Intron 45.5UTR vs.sameotR 15.5 UTP 3 UTR vs. Exon 3 UTR 15. OTR 15. OTR 45.0TR VS. SUTR 25UTR 45. Etor vs. Intron 15. JUTP 45. FUT

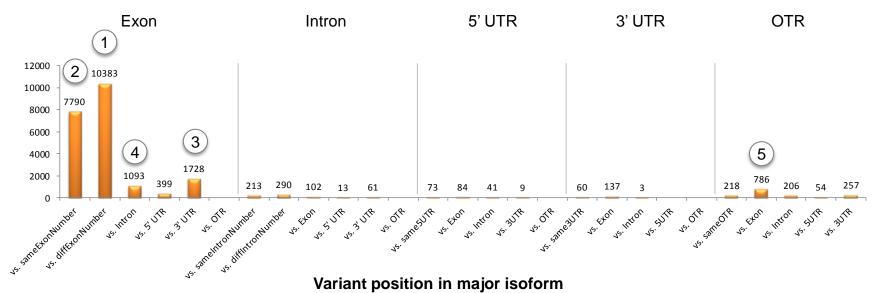
Variant position in major isoform

Total variants examined = 147,540 Total mismatched cases between major and canonical isoforms = 29,669 Mismatched cases that canonical isoform is available = 27,655

Rank 1 = $12,226 / 147,540 * 100 = 8.3\% \rightarrow$ Annotations were changed in 743 cases (0.5%). Rank 2 = 9,373 / 147,540 * 100 = 6.4% Rank 3 = 2,145 / 147,540 * 100 = 1.5% Rank 4 = 1,308 / 147,540 * 100 = 0.9% Rank 5 = 492 / 147,540 * 100 = 0.3%

Fig. S2

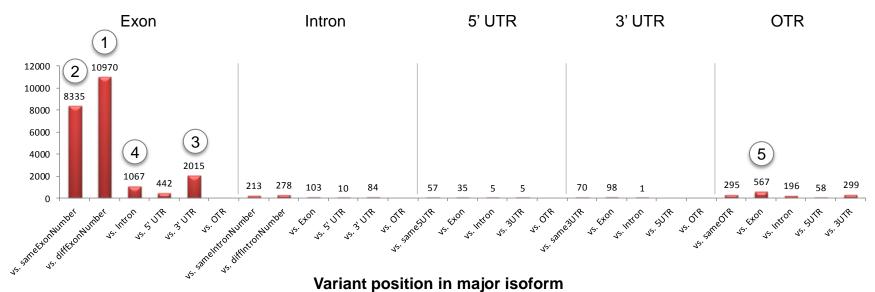
Variant position in UniProt-based canonical isoform



Total variants examined = 147,540 Total mismatched cases between major and canonical isoforms = 29,669 Mismatched cases that canonical isoform is available = 24,000

Rank 1 = 10,383 / 147,540 * 100 = 7.0% → Annotations were changed in 110 cases (0.07%). Rank 2 = 7,790 / 147,540 * 100 = 5.3% Rank 3 = 1,728 / 147,540 * 100 = 1.2% Rank 4 = 1,093 / 147,540 * 100 = 0.7% Rank 5 = 786 / 147,540 * 100 = 0.5%

Variant position in APPRIS principal isoform



Total variants examined = 147,540Total mismatched cases between major and principal isoforms = 29,669Mismatched cases that canonical isoform is available = 25,203

Rank 1 = 10,970 / 147,540 * 100 = 7.4% → Annotations were changed in 467 cases (0.3%). Rank 2 = 8,335 / 147,540 * 100 = 5.6%Rank 3 = 2,015 / 147,540 * 100 = 1.4%Rank 4 = 1,067 / 147,540 * 100 = 0.7%Rank 5 = 567 / 147,540 * 100 = 0.4%