

Fig.S6**A**

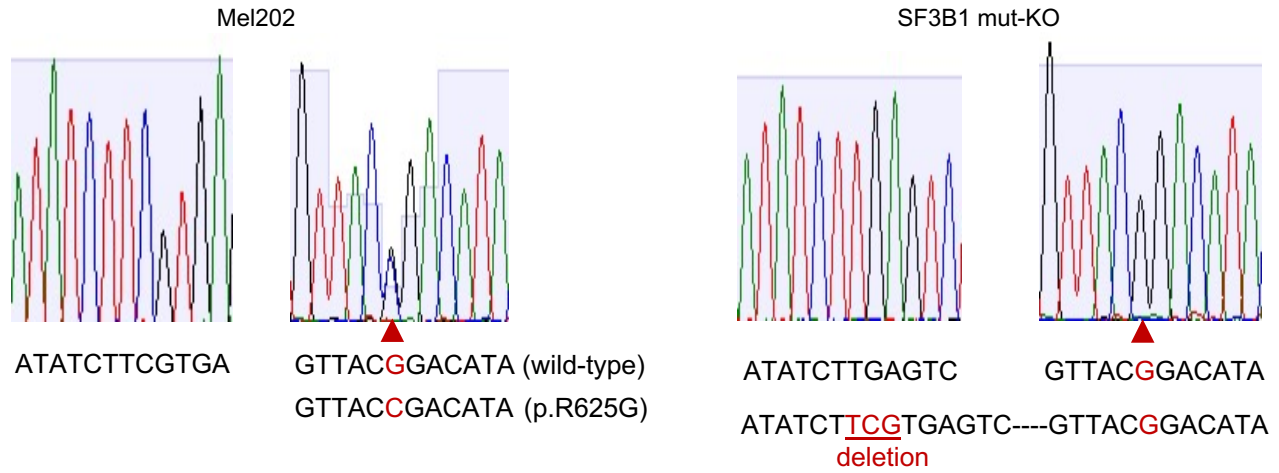
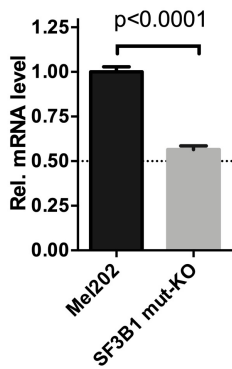
Guide RNA (SF3B1 g1) 5'-AAGATCGCCAAGACTCACGA-3'
 Genomic DNA ATGGCGAAGATCGCCAAGACTCACGAAGGTAA
 PAM

Mel202 SF3B1 mut-KO
 Mutant allele ATGGCGAAGATCGCCAAGACTC-1 bp-CGAAGGTAA
 Indel

Wild-type allele ATGGCGAAGATCGCCAAGACTCA-3 bp-AGGTAA
 Indel
 p.His8_Glu9delinsGln

C

Mel202 KO Clone	SF3B1 WT allele	SF3B1 Mutant allele
SF3B1 mut-KO	3 bp indel	1 bp indel
#1	6 bp indel	1 bp indel
#2	Missense mutation (H8P)	1 bp indel
#3	6 bp indel	1 bp indel
#4	3 bp indel	1 bp indel
#5	6 bp indel	6 bp indel

B**D****E**