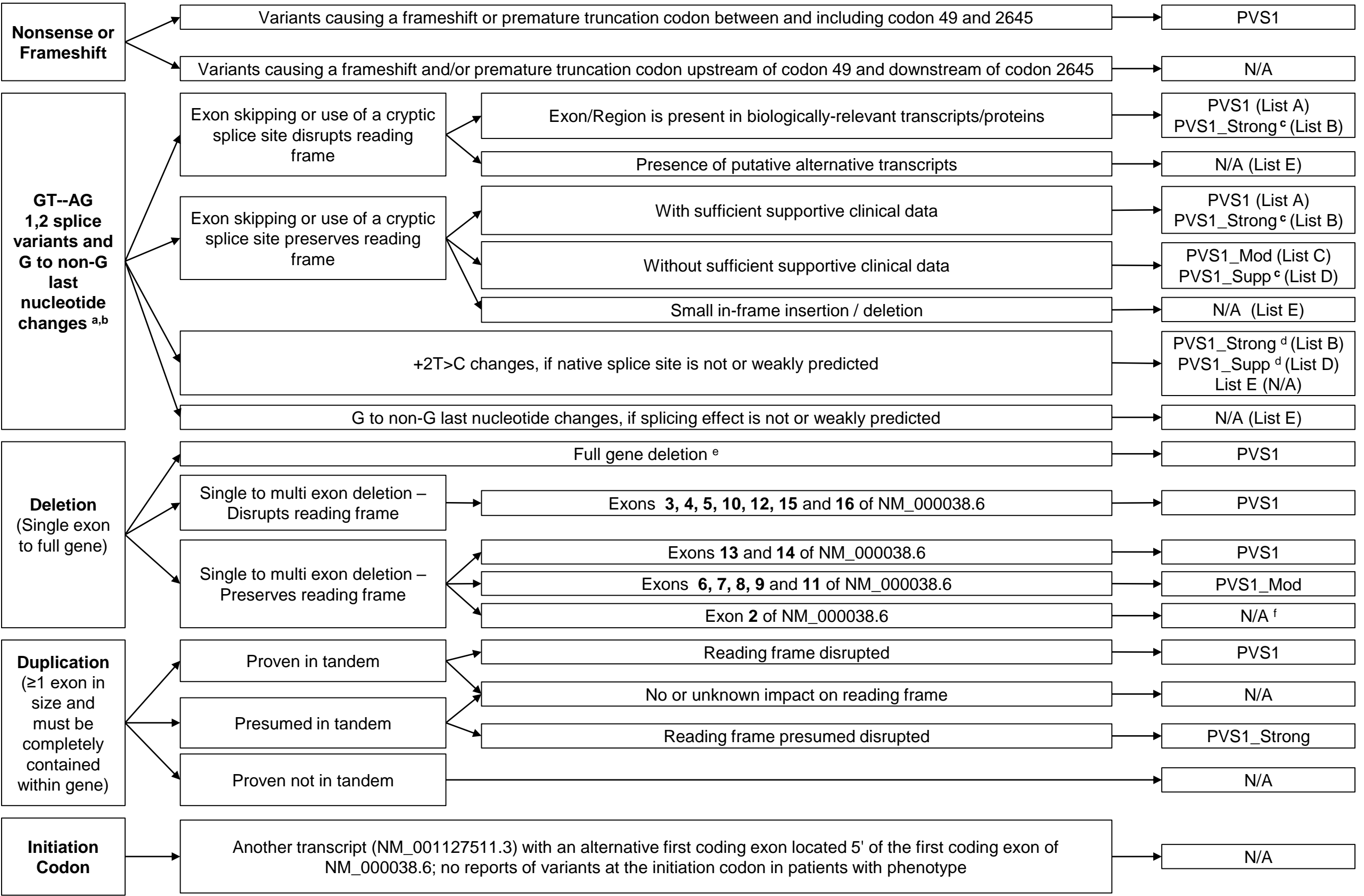


Figure 1(A) APC specific modification of the PVS1 decision tree by Tayoun et al. 2018



^a Splice variants must have no detectable nearby (+/- 20 nt) strong consensus splice sequence that may reconstitute in-frame splicing.

^b For details refer to Fig 1(B). PVS1_variable is only applicable to listed variants.

^c For G to non-G last NT changes, evidence strengths are downgraded by one level.

^d For +2T>C changes for which native splice site is not or weakly predicted, strengths are one level down from the other GT--AG 1,2 splice variants at the same site Fig 1(B).

^e For full gene deletions of a known haploinsufficient gene, a pathogenic classification is warranted in the absence of conflicting evidence with PVS1 alone.

^f Not applicable if promoter 1 A and 1B are also deleted.

Abbreviations: NT – nucleotide, Mod – moderate, Supp – supporting, N/A – not applicable.

Figure 1(B) APC GT--AG 1,2 splice variants and G to non-G last nucleotide changes PVS1 strength specifications

List A (PVS1)			List B (PVS1_Strong)	List C (PVS1_Moderate)	List D (PVS1_Supporting)	List E (N/A)	
c.136-1G>A,C,T	c.646-1G>A,C,T	c.1549-1G>A,C,T	c.220G>A,C,T	c.645+1G>A,C,T	c.729+2T>C	c.-18-1G>A,C,T	c.934-1G>A,C,T
c.136-2A>C,G,T	c.646-2A>C,G,T	c.1549-2A>C,G,T	c.422G>A,C,T	c.645+2T>A,G	c.933G>A,C,T	c.-18-2A>C,G,T	c.934-2A>C,G,T
c.220+1G>A,C,T	c.730-1G>A,C,T	c.1626+1G>A,C,T	c.834G>A,C,T	c.729+1G>A,C,T		c.135G>A,C,T	c.1313-1G>A,C,T
c.220+2T>A,C,G	c.834+1G>A,C,T	c.1626+2T>A,C,G	c.1548G>A,C,T	c.729+2T>A,G		c.135+1G>A,C,T	c.1313-2A>C,G,T
c.221-1G>A,C,T	c.834+2T>A,C,G	c.1627-1G>A,C,T	c.1548+2T>C	c.730-2A>C,G,T		c.135+2T>A,C,G	c.1408G>A,C,T
c.221-2A>C,G,T	c.835-1G>A	c.1627-2A>C,G,T	c.1626G>A,C,T	c.835-1G>C,T		c.645G>A,T,C	c.1959-1G>C,T
c.422+1G>A,C,T	c.933+1G>A,C,T	c.1743+1G>A,C,T	c.1743G>A,C,T	c.835-2A>C,G,T		c.645+2T>C	c.1959-2A>C,G,T
c.422+2T>A,C,G	c.933+2T>A,C,G	c.1743+2T>A,C,G	c.1958G>A,C,T	c.1408+1G>A,C,T		c.729G>A,T,C	
c.423-1G>A,C,T	c.1312+1G>A,C,T	c.1744-1G>A,C,T		c.1408+2T>A,C,G			
c.423-2A>C,G,T	c.1312+2T>A,C,G	c.1744-2A>C,G,T					
c.531+1G>A,C,T	c.1409-1G>A,C,T	c.1958+1G>A,C,T					
c.531+2T>A,C,G	c.1409-2A>C,G,T	c.1958+2T>A,C,G					
c.532-1G>A,C,T	c.1548+1G>A,C,T	c.1959-1G>A					
c.532-2A>C,G,T	c.1548+2T>A,G						