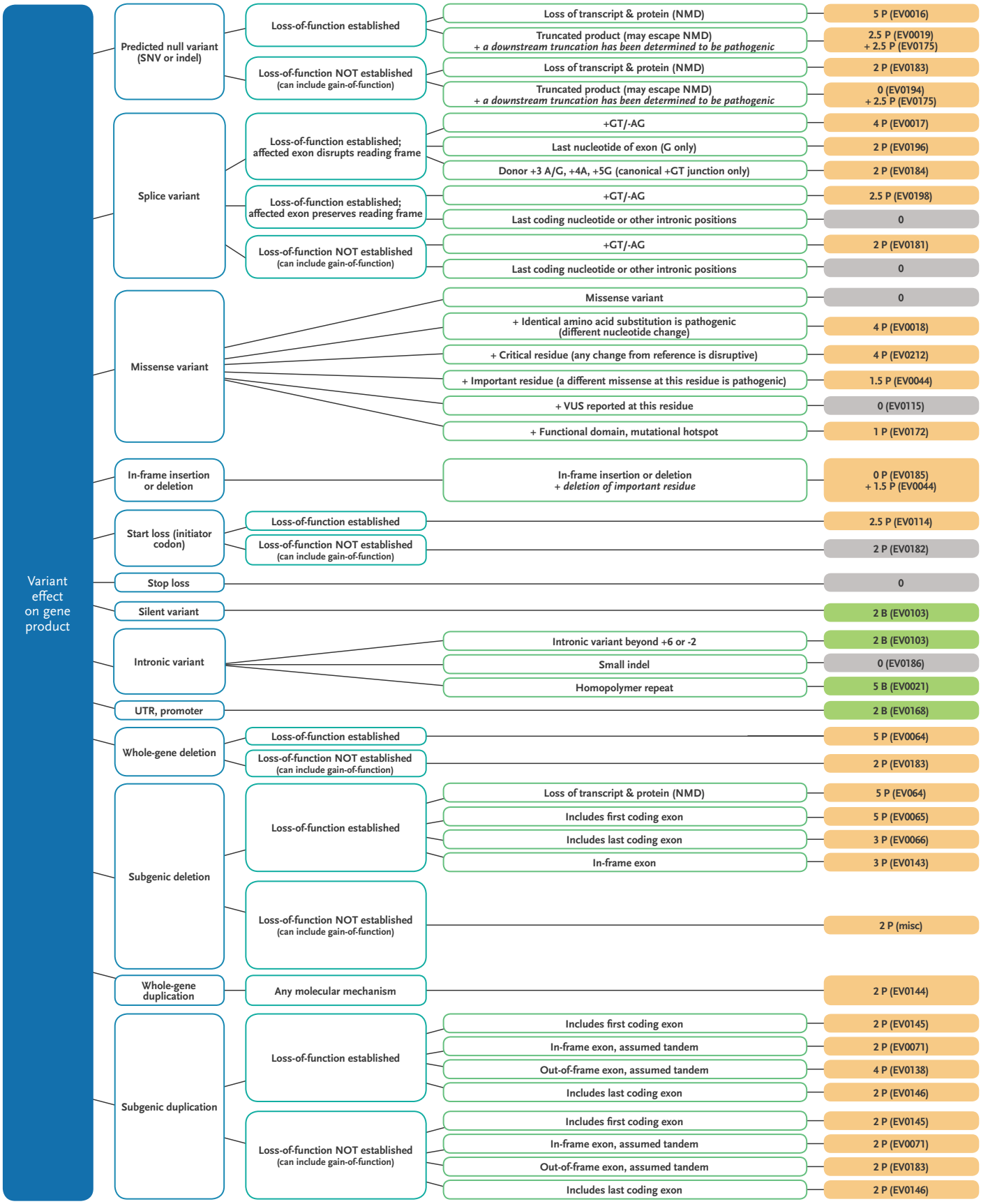
Variant effect on gene product
Predicted null variant (SNV or indel)
Loss-of-function established
Loss of transcript & protein (NMD)
5 P (EV0016)
Truncated product (may escape NMD)
+ a downstream truncation has been determined to be pathogenic
2.5 P (EV0019)
+ 2.5 P (EV0175)
Loss-of-function NOT established
(can include gain-of-function)
Loss of transcript & protein (NMD)
2 P (EV0183)
Truncated product (may escape NMD)
+ a downstream truncation has been determined to be pathogenic
0 (EV0194)
+ 2.5 P (EV0175)
Splice variant
Loss-of-function established; affected exon disrupts reading frame
+GT/-AG
4 P (EV0017)
Last nucleotide of exon (G only)
2 P (EV0196)
Donor +3 A/G, +4A, +5G (canonical +GT junction only)
2 P (EV0184)
Loss-of-function established; affected exon preserves reading frame
+GT/-AG
2.5 P (EV0198)
Last coding nucleotide or other intronic positions
0
Loss-of-function NOT established
(can include gain-of-function)
+GT/-AG
2 P (EV0181)
Last coding nucleotide or other intronic positions
0
Missense variant
Missense variant
0
+ Identical amino acid substitution is pathogenic (different nucleotide change)
4 P (EV0018)
+ Critical residue (any change from reference is disruptive)
4 P (EV0212)
+ Important residue (a different missense at this residue is pathogenic)
1.5 P (EV0044)
+ VUS reported at this residue
0 (EV0115)
+ Functional domain, mutational hotspot
1 P (EV0172)
In-frame insertion or deletion
In-frame insertion or deletion
+ deletion of important residue
0 P (EV0185)
+ 1.5 P (EV0044)
Start loss (initiator codon)
Loss-of-function established
2.5 P (EV0114)
Loss-of-function NOT established
(can include gain-of-function)
2 P (EV0182)
Stop loss
0
Silent variant
2 B (EV0103)
Intronic variant
Intronic variant beyond +6 or -2
2 B (EV0103)
Small indel
0 (EV0186)
Homopolymer repeat
5 B (EV0021)
UTR, promoter
2 B (EV0168)
Whole-gene deletion
Loss-of-function established
5 P (EV0064)
Loss-of-function NOT established
(can include gain-of-function)
2 P (EV0183)
Subgenic deletion
Loss-of-function established
Loss of transcript & protein (NMD)
5 P (EV064)
Includes first coding exon
5 P (EV0065)
Includes last coding exon
3 P (EV0066)
In-frame exon
3 P (EV0143)
Loss-of-function NOT established
(can include gain-of-function)
2 P (misc)
Whole-gene duplication
Any molecular mechanism
2 P (EV0144)
Subgenic duplication
Loss-of-function established
Includes first coding exon
2 P (EV0145)
In-frame exon, assumed tandem
2 P (EV0071)
Out-of-frame exon, assumed tandem
4 P (EV0138)
Includes last coding exon
2 P (EV0146)
Loss-of-function NOT established
(can include gain-of-function)
Includes first coding exon
2 P (EV0145)
In-frame exon, assumed tandem
2 P (EV0071)
Out-of-frame exon, assumed tandem
2 P (EV0183)
Includes last coding exon
2 P (EV0146)