

Combining Rules

Based on Tavtigian et al, 2018 in conjunction with forthcoming points-based guidance.

Points per Criterion		Classification of Variant		
Criterion	Points	Total Points	Classification	Posterior Probability
P – Very Strong	8	≥ 10	Pathogenic	≥ 0.99
P - Strong	4	6-9	Likely Pathogenic	≥ 0.9 to < 0.99
P - Moderate	2	0-5	VUS	≥ 0.10 to < 0.9
P - Supporting	1	-6 to -1	Likely Benign	≥0.001, < 0.1
B - Supporting	-1	-7 and below	Benign	< 0.001
B - Strong	-4			

Additional Caveats:

- PP3+PM1 combined can reach no higher than Strong
- When PM5_Strong is reached, do not combine PM1 to PM5