

RYR1 PS4 specification

PP4 Features to be considered for PS4 point application. PP4 should not be applied separately for AD variants because PS4 specifications factor in phenotypic specificity when determining PS4 strength.

To meet PP4, a congenital myopathy testing panel should be performed without identification of other causative variants and AT LEAST TWO of these features should be present

- **Presence on Muscle Biopsy of:** mini cores or central cores (histology or electron microscopy)
- Exercise, heat, or anesthetic induced rhabdomyolysis
- Characteristic muscle imaging (see Figure 8, Saade et al 2019 PMID: 31060725)

Proband phenotype	Specificity	Points per proband
PP4 Met	Specific	0.5 points
One PP4 feature	Suggestive	0.25 points

PS4_Supporting	PS4_Moderate	PS4_Strong
0.25 points	0.5 points	1 points