**SUPPLEMENTARY FILE S4**

**Measures used for performance evaluations.**

To assess the performance of the three different predictors’ score values, the following parameters were used where TP= true positive (LEANING PATHOGENIC variants with score above cutoff); TN= true negative (LEANING BENIGN variants with score below cutoffs); FP= false positive (LEANING BENIGN variants with score above cutoffs); FN= false negative (LEANING PATHOGENIC variants with score below cutoff).

**Sensitivity**

TPTP + FN

**Specificity**

TNTN + FP

**Accuracy**

TP + TN TP + FP + TN + FN

**Matthews correlation coefficient (MCC)**

TP × TN − FP × FN

**Youden's J index**

Sensitivity + Specificity – 1