Protein folding results for VKGL missense variants in the PLCG2 gene Based on VKGL public consensus release April 2023, FoldX 5.0, and AlphaFold2 human proteome v4 At a  $\Delta\Delta G$  threshold of 13.28 the PPV is 100%, the sensitivity is 100%, the NPV is 100% and the specificity is 100% SA707Y (DVV) and variant protein change energy **VKGL** classification Difference in Gibbs free ene between GRCh37 reference LB/B YA647S LP/P VUS YA482H LA218I GA366R NA798S RA956C AA133V SA198R YA780C IA671V NA946S IA301V AA11V IA141V AA308V PA767L RA1001H TA632K LA848F VA985I SA1042T SA11Z2G VA4241 YA13D TA620M RA700W 300 600 900 1,200 0 Amino acid position within the protein alpha chain of PLCG2