Protein folding results for VKGL missense variants in the MEFV gene Based on VKGL public consensus release April 2023, FoldX 5.0, and AlphaFold2 human proteome v4 At a  $\Delta\Delta G$  threshold of 0.66 the PPV is 70%, the sensitivity is 48%, the NPV is 56% and the specificity is 76% NA766H 20 Difference in Gibbs free energy change ( $\Delta\Delta G$ ) between GRCh37 reference and variant protein PA754R **VKGL** classification 10 -LB/B LP/P VA659F MA6931 VUS GA236V FA480C DA637G TA<sub>12</sub>I 200 400 600 800 Amino acid position within the protein alpha chain of MEFV