

Algorithm	Test set	All mutations			$\Delta\Delta G_{exp} \geq 0$		$\Delta\Delta G_{exp} < 0$	
		R	RMSE	Slope	R	RMSE	R	RMSE
RF	S5296	0.73	1.23	1.04	0.54	1.20	0.50	1.25
	S921	0.78	1.48	1.52	0.72	1.54	0.60	1.33
SVM	S5296	0.70*	1.27	0.97	0.52*	1.26	0.49	1.29
	S921	0.73*	1.53	1.34	0.67*	1.59	0.55*	1.39
XGBoost	S5296	0.71*	1.25	0.99	0.53*	1.22	0.50	1.28
	S921	0.77	1.45	1.40	0.71	1.51	0.59	1.32

R: Pearson correlation coefficient between experimental and predicted $\Delta\Delta G$ values. RMSE (kcal mol⁻¹): root-mean-square error. Slope: the slope of the regression line between experimental and predicted $\Delta\Delta G$ values. All presented values of correlation coefficients are statistically significantly different from zero (p-value \ll 0.01, t-test). *p-value < 0.01 compared to Random Forest (Hittner2003 test).