

**Table S12 - The “structure features” model**

<b>Feature</b>	<b>Estimate</b>
(Intercept)	-3.952
SolvAccessWT	0.008
SolvAccessMut	0.004
RelSolvAccessWT	-4.968
RelSolvAccessMut	-0.675
BuryWT	1.131
BuryMut	0.301
IsLigand	0.318
InStruct	1.492
HelixBreaker	-1.198
Bfactor	0.090
sBfactor	-0.058
snormBfactor	-0.008
ddGPoPMuSiC	0.384
ddGratioFoldX	1.362
AA2FT	0.002
AA2Ligand	-0.033
AA2FTLigand	0.050
HydroMomentMut	-0.006
HydroMomentDiff	-0.002
SolvAccessAA	0.012
RelSolvAccessAA	-2.700
sBfactorAA	0.005
snormBfactorAA	0.019
Eucl20D_ALA	0.133
Eucl20D_ASN	-0.067
Eucl20D_ASP	0.033
Eucl20D_CYS	-0.058
Eucl20D_GLN	0.041
Eucl20D_GLU	-0.070
Eucl20D_GLY	0.144
Eucl20D_HIS	0.082
Eucl20D_ILE	0.202
Eucl20D_LEU	-0.249
Eucl20D_LYS	0.108
Eucl20D_MET	0.195
Eucl20D_PHE	0.099
Eucl20D_PRO	0.075
Eucl20D_SER	-0.036
Eucl20D_THR	0.319
Eucl20D_TRP	-0.188
Eucl20D_TYR	-0.265
Eucl20D_VAL	-0.071
EntropySubEucl	13.860
EntropySuperEucl	-22.087
RelEntropySubEucl	22.839
RelEntropySuperEucl	-25.571
HydroAvgEucl	0.030
HydroWToverAvgEucl	1.570
HydroMutoverAvgEucl	-2.907
PosEucl	0.123
SolvAccessEucl	0.021
RelSolvAccessEucl	-3.037
BfactorEucl	0.122
normBfactorEucl	0.001
sBfactorEucl	-0.065
snormBfactorEucl	-0.029

Eucl2FT	0.050
Eucl2Ligand	-0.066
Eucl2FTLigand	0.071
Hbond_6A	-0.006
SaltBridge_6A	0.180
Hbond_2layers	-0.007
SaltBridge_2layers	-0.643
DT20D_A	-0.047
DT20D_C	-0.859
DT20D_D	0.181
DT20D_F	-0.001
DT20D_G	0.089
DT20D_H	-0.276
DT20D_I	0.151
DT20D_K	0.463
DT20D_L	-0.084
DT20D_M	-0.056
DT20D_N	-0.120
DT20D_P	0.104
DT20D_Q	0.148
DT20D_S	0.197
DT20D_T	0.013
DT20D_V	-0.360
DT20D_W	0.030
DT20D_Y	0.095
EntropySubDT	-0.174
EntropySuperDT	-3.959
RelEntropySubDT	2.254
RelEntropySuperDT	-6.224
HydroAvgDT	0.111
HydroWToverAvgDT	-1.747
HydroMutoverAvgDT	2.459
ChargedDT	0.149
NegDT	-0.082
SolvAccessDT	-0.018
RelSolvAccessDT	8.095
sBfactorDT	-0.063
snormBfactorDT	-0.008
DTcountType1	0.037
DTcountType2	0.105
DTcountType3	-0.116
DTcountType4	-0.118

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