

Table S6 - The “neighborhood features” model

| Feature | Estimate |
|---------------------|----------|
| (Intercept) | -9.486 |
| AA20D_A | 0.041 |
| AA20D_C | 0.149 |
| AA20D_D | -0.130 |
| AA20D_F | -0.002 |
| AA20D_G | -0.101 |
| AA20D_H | -0.125 |
| AA20D_I | 0.140 |
| AA20D_K | -0.004 |
| AA20D_L | -0.150 |
| AA20D_M | -0.001 |
| AA20D_P | -0.339 |
| AA20D_Q | -0.062 |
| AA20D_S | -0.214 |
| AA20D_T | -0.188 |
| AA20D_V | -0.078 |
| AA20D_W | 0.242 |
| AA20D_Y | 0.008 |
| RelEntropySubAA | 2.672 |
| RelEntropySuperAA | -1.400 |
| HydroAvgWT | -0.003 |
| HydroAvgDiff | 0.057 |
| PolarAA | -0.015 |
| NonpolarAA | -0.051 |
| ChargedAA | -0.021 |
| NegAA | 0.070 |
| AA2Ligand | -0.032 |
| AA2FTLigand | 0.039 |
| HydroMomentMut | -0.005 |
| HydroMomentDiff | -0.005 |
| SolvAccessAA | -0.006 |
| RelSolvAccessAA | 1.995 |
| sBfactorAA | 0.013 |
| snormBfactorAA | 0.013 |
| Eucl20D_ALA | 0.086 |
| Eucl20D_ARG | 0.023 |
| Eucl20D ASN | -0.142 |
| Eucl20D ASP | 0.054 |
| Eucl20D CYS | -0.169 |
| Eucl20D GLN | 0.050 |
| Eucl20D GLU | -0.123 |
| Eucl20D GLY | 0.114 |
| Eucl20D ILE | 0.218 |
| Eucl20D LEU | -0.205 |
| Eucl20D LYS | 0.113 |
| Eucl20D MET | 0.237 |
| Eucl20D PHE | 0.091 |
| Eucl20D PRO | 0.124 |
| Eucl20D THR | 0.360 |
| Eucl20D TRP | -0.307 |
| Eucl20D TYR | -0.154 |
| Eucl20D VAL | -0.057 |
| RelEntropySubEucl | 4.342 |
| RelEntropySuperEucl | -1.481 |
| HydroAvgEucl | -0.008 |
| HydroWToverAvgEucl | 0.573 |
| HydroMutoverAvgEucl | -3.332 |

| | |
|--------------------|--------|
| PosEucl | 0.094 |
| SolvAccessEucl | -0.013 |
| RelSolvAccessEucl | 8.019 |
| BfactorEucl | 0.005 |
| Eucl2FT | 0.050 |
| Eucl2Ligand | -0.035 |
| Eucl2FTLigand | 0.020 |
| Hbond_6A | -0.043 |
| SaltBridge_6A | 0.120 |
| Hbond_2layers | 0.007 |
| SaltBridge_2layers | -0.256 |
| DT20D_A | -0.069 |
| DT20D_C | -0.843 |
| DT20D_D | 0.192 |
| DT20D_F | -0.061 |
| DT20D_G | 0.043 |
| DT20D_H | -0.005 |
| DT20D_I | 0.083 |
| DT20D_K | 0.421 |
| DT20D_L | -0.070 |
| DT20D_M | -0.060 |
| DT20D_N | -0.149 |
| DT20D_P | 0.191 |
| DT20D_Q | 0.318 |
| DT20D_R | 0.140 |
| DT20D_S | 0.147 |
| DT20D_T | 0.065 |
| DT20D_V | -0.295 |
| DT20D_Y | -0.005 |
| EntropySuperDT | -5.188 |
| RelEntropySubDT | 2.862 |
| RelEntropySuperDT | -8.278 |
| HydroAvgDT | 0.147 |
| HydroMutoverAvgDT | 2.195 |
| NonPolarDT | -0.029 |
| ChargedDT | 0.226 |
| SolvAccessDT | -0.058 |
| RelSolvAccessDT | 6.982 |
| BfactorDT | 0.004 |
| DTcountType0 | 0.073 |
| DTcountType1 | 0.120 |
| DTcountType2 | 0.148 |
| DTcountType3 | -0.006 |