

Supplemental; Small compounds modulate and bind MscL similarly

S3 Table. Hotspot residue identification using MM-GBSA binding free energy decomposition analysis for Eco-MscL/K05. A hotspot residue is recognized when its interaction energy with the ligand is better than -1.0 kcal/mol. Those cells without numbers have neglectable interaction energies.

Hotspot Residue	Scenario 3 Channel-Close	Scenario 7 Channel-Open
PHE7	-6.58	
PHE10	-5.12	-0.11
ALA11	-0.27	-1.25
ARG13		-3.93
GLY14		-3.98
ASN15		-4.01
VAL17		-2.18
ASP18		-4.08
PRO115		-1.01
ILE161/25	-2.99	>-0.1
ASN236		-3.09
ASN239		-1.93
ARG240		-6.84
PRO249		-2.88
PRO251		-3.51
PHE357/85	-2.44	
PHE358/86	-2.19	
PHE361/89	-2.00	
PHE365/93	-2.46	
PHE554		-1.03
ARG557		-2.36