

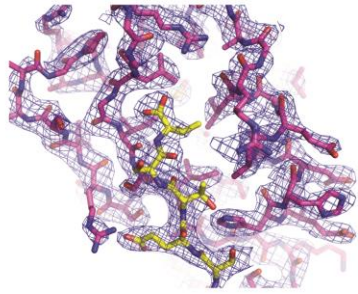
Expanded view figure legends

FIGURE EV2: 2Fo-Fc electron density maps of PDZ1, PDZ2 and PDZ3 complexes with Vangl2 as well as PDZ2 on its own. **A)** Electron density map encompassing the binding groove of Scribble PDZ1 (magenta sticks) in complex with Vangl2 peptide (yellow sticks). The electron density map is shown as a blue mesh contoured at 1.5σ . **B)** Electron density map encompassing the binding groove of Scribble PDZ2 (green sticks) in complex with Vangl2 (yellow sticks). The electron density map is shown as a blue mesh contoured at 1.5σ . **C)** Electron density map encompassing the binding groove of Scribble PDZ3 (orange sticks) in complex with Vangl2 peptide (yellow sticks). The electron density map is shown as a blue mesh contoured at 1.5σ .

FIGURE EV3: Circular dichroism spectroscopy of wild type and mutant Scribble PDZ1, PDZ2, PDZ3 and PDZ4 domains.

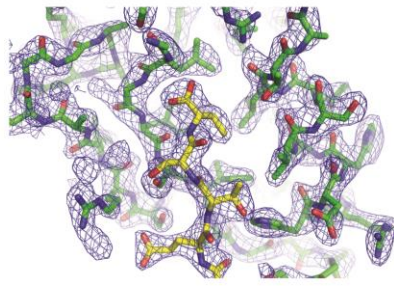
Circular dichroism spectra recorded for wild type and mutant Scribble PDZ1,2,3 and 4 domains indicated that there were no major spectral differences between the proteins, suggesting that they were similarly folded, with mutations not leading to unfolding of the PDZ domains.

A



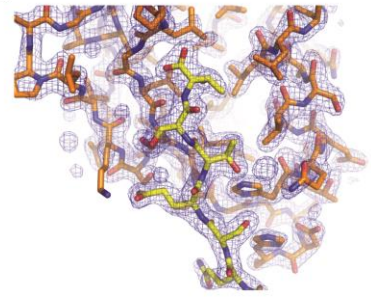
SCRIB PDZ1:Vangl2

B

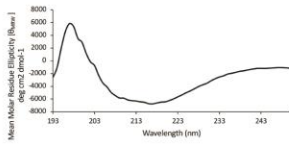


SCRIB PDZ2:Vangl2

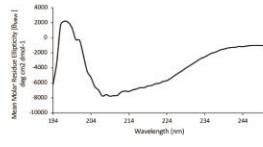
C



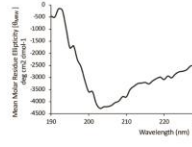
SCRIB PDZ3:Vangl2



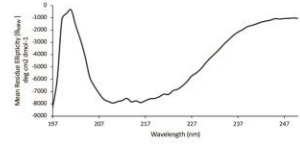
PDZ1 WT



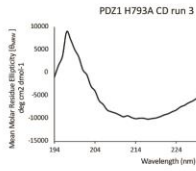
PDZ2 WT



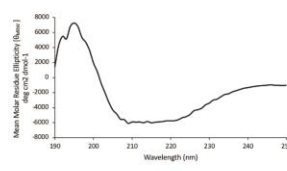
PDZ3 WT



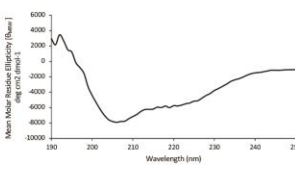
PDZ4 WT



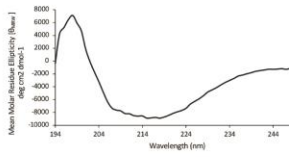
PDZ1 H793A



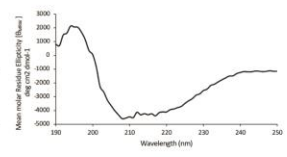
PDZ2 R896A



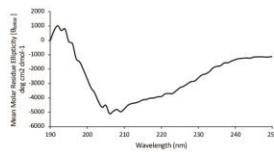
PDZ3 P1043L



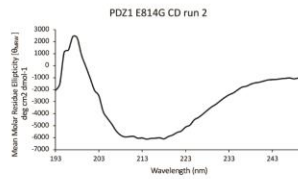
PDZ1 Q808H



PDZ2 H928A



PDZ3 R1044Q



PDZ1 E814G

Table EV1: X-ray crystallographic data collection and refinement statistics.

	PDZ1:Vangl2	PDZ2 apo	PDZ2:Vangl2	PDZ3:Vangl2
Data collection				
Space group	C 1 2 1	P 1	I 2 ₁ 2 ₁ 2 ₁	P 4 ₁ 2 ₁ 2
No of molecules in AU	4+4	4	4+4	2+2
Cell dimensions				
<i>a</i> , <i>b</i> , <i>c</i> (Å)	219.37, 53.09, 53.52	37.89, 41.69, 49.02	78.05, 60.77, 96.02	54.71, 54.71, 129.98
α , β , γ (°)	90.00, 104.12, 90.00	90.27, 91.96, 105.84	90.00, 106.48, 90.00	90.00, 90.00, 90.00
Wavelength (Å)	0.9537	0.9537	0.9537	0.9537
Resolution (Å)*	51.51-2.20 (2.28-2.20)	31.60-2.44 (2.53-2.44)	47.17-2.50 (2.59-2.50)	37.08-1.95 (2.02-1.95)
<i>R</i> _{sym} Or <i>R</i> _{merge} *	0.066 (0.620)	0.121 0.650)	0.076 (0.425)	0.060 (0.763)
<i>I</i> / σ <i>I</i> *	8.3 (1.6)	10.5 (2.1)	16.2 (4.5)	38.4 (5.5)
CC(1/2)	0.999 (0.743)	0.994 (0.792)	0.999 (0.856)	1.000 (0.952)
Completeness (%)*	99.9 (100.0)	98.3 98.0)	91.5 (93.2)	99.9 (99.93)
Redundancy*	5.0 (5.1)	3.9 (3.9)	5.7 (5.7)	27.6 (28.6)
Wilson B-factor	49.57	28.24	30.84	34.1
Refinement				
Resolution (Å)	51.51-2.20	31.60-2.44	47.17-2.50	37.08-1.95
No. reflections	30485	10507	13813	15101
<i>R</i> _{work} / <i>R</i> _{free}	0.260/0.288	0.225/0.269	0.236/0.270	0.223/0.262
No. non-hydrogen atoms				
Protein	3238	2699	2777	1328
Ligand/ion	111	8	32	-
Water	46	81	98	81
<i>B</i> -factors				
Protein	68.66	32.90	35.60	46.36
Ligand/ion	71.34	37.21	42.39	-
Water	59.55	31.71	31.62	45.70
R.m.s. deviations				
Bond lengths (Å)	0.006	0.004	0.003	0.004
Bond angle (°)	1.09	1.01	0.68	1.03
Ramachandran plot (%)				
Favored	97.8	97.4	98.1	99.4
Allowed	2.2	2.6	1.9	0.6
Disallowed	0	0	0	0