

**Table S1** Kinetic (un)folding parameters of the Crkl SH2 domain calculated at different Na<sub>2</sub>SO<sub>4</sub> concentrations.

[Na <sub>2</sub> SO <sub>4</sub> ] (M)	$k_{\text{IN}}$ (s <sup>-1</sup> )	$k_{\text{NI}}$ (s <sup>-1</sup> )	$K_{\text{IU}}$
0	26 ± 2	0.0084 ± 0.0007	0.0052 ± 0.0011
0.15	124 ± 9	0.0030 ± 0.0003	0.0013 ± 0.0003
0.3	150 ± 12	0.0015 ± 0.0001	0.0003 ± 0.0001

Note: The entire data set was globally fitted with Eq.2, sharing  $m_{\text{I-N}}$ ,  $m_{\text{N-I}}$  and  $m_{\text{I-D}}$  values:  $m_{\text{I-N}} = 0.28 \pm 0.05$  Kcal mol<sup>-1</sup> M<sup>-1</sup>,  $m_{\text{N-I}} = 1.00 \pm 0.01$  Kcal mol<sup>-1</sup> M<sup>-1</sup>,  $m_{\text{I-D}} = 2.60 \pm 0.05$  Kcal mol<sup>-1</sup> M<sup>-1</sup>. The global  $m_{\text{D-N}}$  value was  $3.9 \pm 0.7$  Kcal mol<sup>-1</sup> M<sup>-1</sup> calculated as the sum of  $m_{\text{I-N}}$ ,  $m_{\text{N-I}}$ , and  $m_{\text{I-D}}$ .

**Table S2.** Kinetic (un)folding parameters of the Crkl SH2 domain calculated at different pH conditions in presence of 0.15 M Na<sub>2</sub>SO<sub>4</sub>.

pH	$k_{IN}$ (s <sup>-1</sup> )	$k_{NI}$ (s <sup>-1</sup> )	$K_{ID}$	$m_{N-I}$ (Kcal mol <sup>-1</sup> M <sup>-1</sup> )	$m_{D-N}$ (Kcal mol <sup>-1</sup> M <sup>-1</sup> )
4.0	9 ± 5	1.1490 ± 0.1543	0.2776 ± 0.2969	1.35 ± 0.04	4.13 ± 0.36
4.5	15 ± 1	0.0927 ± 0.0106	0.0428 ± 0.0077	1.36 ± 0.03	4.14 ± 0.33
5.0	35 ± 5	0.0192 ± 0.0019	0.0409 ± 0.0090	1.22 ± 0.02	4.00 ± 0.31
5.5	126 ± 17	0.0049 ± 0.0004	0.0485 ± 0.0107	1.12 ± 0.01	3.91 ± 0.30
6.7	122 ± 11	0.0040 ± 0.0005	0.0158 ± 0.0029	0.98 ± 0.02	3.76 ± 0.29
7.2	93 ± 8	0.0036 ± 0.0005	0.0014 ± 0.0002	0.98 ± 0.02	3.76 ± 0.29
8.0	55 ± 5	0.0034 ± 0.0005	0.0009 ± 0.0002	1.02 ± 0.02	3.80 ± 0.30
8.5	57 ± 5	0.0059 ± 0.0009	0.0010 ± 0.0002	0.97 ± 0.02	3.75 ± 0.29
9.0	63 ± 6	0.0110 ± 0.0016	0.0013 ± 0.0002	0.92 ± 0.02	3.70 ± 0.29

Note: The entire data set at different pH was globally fitted with Eq.2, sharing  $m_{I-N}$  and  $m_{I-D}$  values;  $m_{I-N}$  = 0.06 ± 0.06 Kcal mol<sup>-1</sup> M<sup>-1</sup>;  $m_{I-D}$  = 2.72 ± 0.05 Kcal mol<sup>-1</sup> M<sup>-1</sup>;  $m_{D-N}$  value =  $m_{I-N}$  +  $m_{N-I}$  +  $m_{I-D}$ .

**Table S3** Kinetics parameters obtained from pseudo-first order binding reaction between the wild-type Crkl SH2 domain and Pax<sub>112-123</sub> peptide, at different ionic strengths and 283K.

[NaCl] (M)	$k_{\text{on}}$ ( $\mu\text{M}^{-1} \text{s}^{-1}$ )	$k_{\text{off}}$ ( $\text{s}^{-1}$ )	$K_{\text{D}}$ ( $\mu\text{M}$ )
0.15	$45.9 \pm 2.2$	$20.4 \pm 1.5$	$0.4 \pm 0.3$
0.3	$24.7 \pm 2.8$	$21.8 \pm 1.1$	$0.9 \pm 0.7$
0.5	$10.4 \pm 1.4$	$28.6 \pm 0.5$	$2.8 \pm 0.4$
1	$6.9 \pm 0.6$	$39.6 \pm 4.0$	$5.8 \pm 0.7$