

Table S5: Table of Estimated Parameters in Models 1-8 and Models S2-S3.

	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6	Model 7	Model 8	Model S2	Model S3
$kLRC_1$ ($nM^{-1}min^{-1}$)	213.3	79.09	1036	219.9	0.7669	2388	34.11	46.32	0.9068	2443
$kLRC_2$ ($nM^{-1}min^{-1}$)	3459	22.56	7900	62.49	4.677	0.1182	1025	0.04041	0.04956	0.0066
$kfSmad2$ ($nM^{-1}min^{-1}$)	0.03958	0.02861	8.385e-4	0.1599	0.04441	0.6500	0.01185	1.252	0.9100	6.930
$krSmad2$ ($nM^{-1}min^{-1}$)	2.443e5	123.6	8598	1332	113.7	2.036e5	99.44	4546	8183	1.851e4
$kfSmadsComplex$ ($nM^{-1}min^{-1}$)	0.4670	0.01039	0.02771	2.586e-5	0.03195	3.122e-4	0.01329	1.452	1.074	2.080
$krSmadsComplex$ ($nM^{-1}min^{-1}$)	0.2858	0.3054	4.024	2.297e-3	1.220	0.05235	0.1560	6.740	4.194	30.59
$kdeph_{pSmad2}$ (min^{-1})	999.6	1.142	0	0	0.6757	11.06	9.676	0	0	0
$kdeg_{pSmad2}$ (min^{-1})	0	0	0	0.6789	0.07010	0.9444	3.163	0.5199	0	0
v_{Smad2} ($nMmin^{-1}$)	0	0	0	0	0	0	<u>5.714</u>	<u>5.714</u>	0	0
$kdeg_{Smad2}$ (min^{-1})	0	0	0	0	0	0	<u>0.01000</u>	<u>0.01000</u>	0	0
$klid$ ($nM^{-1}min^{-1}$)	0	6.779e-3	313.2	0	8.579	0	0	0	0	0
v_{PPM1A} ($nMmin^{-1}$)	0	0	0	0	0	0	0	<u>7.000</u>	<u>0.1000</u>	<u>7.000</u>
$kdeg_{PPM1A}$ (min^{-1})	0	0	0	0	0	0	0	<u>0.07000</u>	<u>0.01000</u>	<u>0.07000</u>
$kdeph_{PPM1A}$ (min^{-1})	0	0	0	0	0	0	0	9.356	0.3980	10.22
$kPPM1A$ (min^{-1})	0	0	0	0	0	0	0	0	1.178e-3	0
$kfPPM1A$ ($nM^{-1}min^{-1}$)	0	0	0	0	0	0	0	0.1819	45.63	0.3583
$kbPPM1A$ (min^{-1})	0	0	0	0	0	0	0	41.98	7.354	282.7
$krPPM1A$ (min^{-1})	0	0	0	0	0	0	0	1.604e-10	0	3.95e-7
$kfPP$ ($nM^{-1}min^{-1}$)	0	0	0	0	0	0	0	82.26	0	1355
$kbPP$ (min^{-1})	0	0	0	0	0	0	0	7.366e-11	0	5.252e-4
$krPP$ (min^{-1})	0	0	0	0	0	0	0	0.8848	0	0.3323
$kfPTEN$ ($nM^{-1}min^{-1}$)	0	0	0	0	0	0	0	6.312	0	4380
$kbPTEN$ (min^{-1})	0	0	0	0	0	0	0	6.788e-10	0	3.182e-4
$kimp_{PPM1A}$ (min^{-1})	0	0	0	0	0	0	0	<u>0.07</u>	0	<u>0.07</u>
$kimp_{PP}$ (min^{-1})	0	0	0	0	0	0	0	2.378e-3	0	0.0023
$kexp_{PP}$ (min^{-1})	0	0	0	0	0	0	0	1.145e-10	0	9.471e-7
$kexp_{PTEN}$ (min^{-1})	0	0	0	0	0	0	0	3.268e-05	0	26.44
$PTEN_{Cyt}$ (nM)	0	0	0	0	0	0	0	176.3	0	176.4

Model S1 is excluded from this table because multiple fitted models (with different parameters) were selected from the structure of Model S1. The underlined values have been manually tuned. All the others were estimated numerically with MATLAB fmincon optimization. The last parameter (for PTEN) is an initial concentration. All the other parameters are rate constants.