

Table S1: Parameter values of the full and optimal models in the last iteration. Here the average deviation at each point between the optimal and the full model is less than 25%. Model 1 refers to EGFR model without constraint to prevent the pathway to Ras protein whereas Model 2 refers to EGFR model with the constraint.

Parameter	Model 1		Model 2	
	$k_f^6$	$k_r^6$	$k_f^7$	$k_r^7$
$k_1$	$3.1 \times 10^{-3}$	$2.8 \times 10^{-3}$	$3 \times 10^{-3}$	$2.7 \times 10^{-3}$
$k_{-1}$	$5.77 \times 10^{-2}$	$5.31 \times 10^{-2}$	$5.84 \times 10^{-2}$	$5.09 \times 10^{-2}$
$k_2$	$9.5 \times 10^{-3}$	$9.9 \times 10^{-3}$	$9.6 \times 10^{-3}$	$9.8 \times 10^{-3}$
$k_{-2}$	0.1336	$9.24 \times 10^{-2}$	$8.22 \times 10^{-2}$	0.1087
$k_3$	1.0679	0.79	0.9481	1.0264
$k_{-3}$	0.0194	0	0.016	0
$V_4$	450.0001	477.8907	440.9235	437.1619
$K_4$	49.9990	48.2002	51.0252	46.1166
$k_5$	$6.57 \times 10^{-2}$	$5.95 \times 10^{-2}$	$6.4 \times 10^{-2}$	$5.1 \times 10^{-2}$
$k_{-5}$	0.1917	0	0.3116	0
$k_6$	0.9992	1.0237	0.9264	0.8975
$k_{-6}$	$4.41 \times 10^{-2}$	0	$4.04 \times 10^{-2}$	0
$k_7$	0.3079	0.2803	0.3053	0.2852
$k_{-7}$	$4 \times 10^{-3}$	$7.4 \times 10^{-3}$	$5 \times 10^{-3}$	$4.6 \times 10^{-3}$
$V_8$	0.9894	0	0.7757	0
$K_8$	100	112.7967	101.4729	97.2425
$k_9$	$4.2 \times 10^{-3}$	$4.4 \times 10^{-3}$	$2.8 \times 10^{-3}$	$2.7 \times 10^{-3}$
$k_{-9}$	$7.95 \times 10^{-2}$	$4.03 \times 10^{-2}$	$4.45 \times 10^{-2}$	$3.19 \times 10^{-2}$
$k_{10}$	$5.8 \times 10^{-3}$	0	$4.9 \times 10^{-3}$	0
$k_{-10}$	$5.26 \times 10^{-2}$	0	$4.05 \times 10^{-2}$	0
$k_{11}$	$7.68 \times 10^{-2}$	0	$9.18 \times 10^{-2}$	0
$k_{-11}$	$1.66 \times 10^{-2}$	0	$1.64 \times 10^{-2}$	0
$k_{12}$	$1.73 \times 10^{-2}$	0	$7.5 \times 10^{-3}$	0
$k_{-12}$	$1.073 \times 10^{-4}$	0	$5.07 \times 10^{-5}$	0
$k_{13}$	$9.48 \times 10^{-2}$	0.1036	$8.49 \times 10^{-2}$	$8.39 \times 10^{-2}$
$k_{-13}$	0.6009	0	0.6059	0
$k_{14}$	5.9984	$1 \times 10^4$	5.3676	$1 \times 10^4$
$k_{-14}$	$5.63 \times 10^{-2}$	0	$9.71 \times 10^{-2}$	0
$k_{15}$	0.3346	0.2904	0.3024	0.2903
$k_{-15}$	$1 \times 10^{-3}$	$2.6 \times 10^{-3}$	$1.6 \times 10^{-3}$	$1.1 \times 10^{-3}$
$K_{16}$	1.6962	1.9833	1.7624	1.6462
$V_{16}$	340	330.3266	324.2768	327.9718
$k_{17}$	$2.9 \times 10^{-3}$	$2.8 \times 10^{-3}$	$3.4 \times 10^{-3}$	$2.7 \times 10^{-3}$
$k_{-17}$	0.1043	0	$8.53 \times 10^{-2}$	0
$k_{18}$	0.2973	0.2272	0.2962	0.2016
$k_{-18}$	$4 \times 10^{-4}$	$1.6 \times 10^{-3}$	$2.54 \times 10^{-5}$	0
$k_{19}$	$1.83 \times 10^{-2}$	0	$1.65 \times 10^{-2}$	0
$k_{-19}$	$2.25 \times 10^{-2}$	0	0.1027	0
$k_{20}$	0.1235	0	0.1159	0.1928
$k_{-20}$	$8.496 \times 10^{-4}$	0	$2.596 \times 10^{-4}$	$6.813 \times 10^{-4}$
$k_{21}$	$2.7 \times 10^{-3}$	$3.1 \times 10^{-3}$	$2.8 \times 10^{-3}$	$2.4 \times 10^{-3}$
$k_{-21}$	$9.84 \times 10^{-2}$	$3.54 \times 10^{-2}$	$9.72 \times 10^{-2}$	$7.51 \times 10^{-2}$
$k_{22}$	$2.95 \times 10^{-2}$	0	$7.48 \times 10^{-2}$	$9.6 \times 10^{-3}$
$k_{-22}$	$5.35 \times 10^{-2}$	0	$1.38 \times 10^{-2}$	0.1298
$k_{23}$	$9.66 \times 10^{-2}$	0	0.1307	0
$k_{-23}$	$2.99 \times 10^{-2}$	0	$4.13 \times 10^{-2}$	0
$k_{24}$	$1.24 \times 10^{-2}$	0	$1.41 \times 10^{-2}$	0
$k_{-24}$	$4.86 \times 10^{-2}$	0	$8.14 \times 10^{-2}$	0

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Table S1 – *Continued from previous page*

Parameter	Model 1		Model 2	
	$\mathbf{k}_f^6$	$\mathbf{k}_r^6$	$\mathbf{k}_f^\tau$	$\mathbf{k}_r^\tau$
$k_{25}$	1.0068	1.1123	0.9916	1.0597
$k_{-25}$	$3.36 \times 10^{-2}$	$3.41 \times 10^{-2}$	$3.12 \times 10^{-2}$	$3.43 \times 10^{-2}$