

Differential Equations

$$\begin{aligned}
 \frac{d[\text{EGF}]}{dt} &= \text{cell} \cdot \text{kruEGF} \cdot [\text{boundEGFReceptor}] \\
 &\quad - \text{cell} \cdot \text{krbEGF} \cdot [\text{EGF}] \cdot [\text{freeEGFReceptor}] \\
 \\
 \frac{d[\text{NGF}]}{dt} &= \text{kruNGF} \cdot [\text{boundNGFReceptor}] \cdot \text{cell} \\
 &\quad - \text{krbNGF} \cdot [\text{NGF}] \cdot [\text{freeNGFReceptor}] \cdot \text{cell} \\
 \\
 \frac{d[\text{freeEGFReceptor}]}{dt} &= \text{cell} \cdot \text{kruEGF} \cdot [\text{boundEGFReceptor}] \\
 &\quad - \text{cell} \cdot \text{krbEGF} \cdot [\text{EGF}] \cdot [\text{freeEGFReceptor}] \\
 \\
 \frac{d[\text{boundEGFReceptor}]}{dt} &= \text{cell} \cdot \text{krbEGF} \cdot [\text{EGF}] \cdot [\text{freeEGFReceptor}] \\
 &\quad - \text{cell} \cdot \text{kruEGF} \cdot [\text{boundEGFReceptor}] \\
 \\
 \frac{d[\text{freeNGFReceptor}]}{dt} &= \text{kruNGF} \cdot [\text{boundNGFReceptor}] \cdot \text{cell} \\
 &\quad - \text{krbNGF} \cdot [\text{NGF}] \cdot [\text{freeNGFReceptor}] \cdot \text{cell} \\
 \\
 \frac{d[\text{boundNGFReceptor}]}{dt} &= \text{krbNGF} \cdot [\text{NGF}] \cdot [\text{freeNGFReceptor}] \cdot \text{cell} \\
 &\quad - \text{kruNGF} \cdot [\text{boundNGFReceptor}] \cdot \text{cell} \\
 \\
 \frac{d[\text{SosInactive}]}{dt} &= \frac{\text{cell} \cdot \text{kdSos} \cdot [\text{P90RskActive}] \cdot [\text{SosActive}]}{([\text{SosActive}] + \text{KmdSos})} \\
 &\quad - \frac{\text{cell} \cdot \text{kEGF} \cdot [\text{boundEGFReceptor}] \cdot [\text{SosInactive}]}{([\text{SosInactive}] + \text{KmEGF})} \\
 &\quad - \frac{\text{cell} \cdot \text{kNGF} \cdot [\text{boundNGFReceptor}] \cdot [\text{SosInactive}]}{([\text{SosInactive}] + \text{KmNGF})} \\
 \\
 \frac{d[\text{SosActive}]}{dt} &= \frac{\text{cell} \cdot \text{kEGF} \cdot [\text{boundEGFReceptor}] \cdot [\text{SosInactive}]}{([\text{SosInactive}] + \text{KmEGF})} \\
 &\quad + \frac{\text{cell} \cdot \text{kNGF} \cdot [\text{boundNGFReceptor}] \cdot [\text{SosInactive}]}{([\text{SosInactive}] + \text{KmNGF})} \\
 &\quad - \frac{\text{cell} \cdot \text{kdSos} \cdot [\text{P90RskActive}] \cdot [\text{SosActive}]}{([\text{SosActive}] + \text{KmdSos})} \\
 \\
 \frac{d[\text{P90RskInactive}]}{dt} &= - \left(\frac{\text{cell} \cdot \text{kpP90Rsk} \cdot [\text{ErkActive}] \cdot [\text{P90RskInactive}]}{([\text{P90RskInactive}] + \text{KmpP90Rsk})} \right) \\
 \\
 \frac{d[\text{P90RskActive}]}{dt} &= \frac{\text{cell} \cdot \text{kpP90Rsk} \cdot [\text{ErkActive}] \cdot [\text{P90RskInactive}]}{([\text{P90RskInactive}] + \text{KmpP90Rsk})} \\
 \\
 \frac{d[\text{RasInactive}]}{dt} &= \frac{\text{cell} \cdot \text{kRasGap} \cdot [\text{RasGapActive}] \cdot [\text{RasActive}]}{([\text{RasActive}] + \text{KmRasGap})} \\
 &\quad - \frac{\text{cell} \cdot \text{kSos} \cdot [\text{SosActive}] \cdot [\text{RasInactive}]}{([\text{RasInactive}] + \text{KmSos})} \\
 \\
 \frac{d[\text{RasActive}]}{dt} &= \frac{\text{cell} \cdot \text{kSos} \cdot [\text{SosActive}] \cdot [\text{RasInactive}]}{([\text{RasInactive}] + \text{KmSos})} \\
 &\quad - \frac{\text{cell} \cdot \text{kRasGap} \cdot [\text{RasGapActive}] \cdot [\text{RasActive}]}{([\text{RasActive}] + \text{KmRasGap})}
 \end{aligned}$$

$$\begin{aligned}
\frac{d[\text{Raf1Inactive}]}{dt} &= \frac{\text{cell} \cdot \text{kRaf1} \cdot [\text{Raf1PPtase}] \cdot [\text{Raf1Active}]}{([\text{Raf1Active}] + \text{KmdRaf1})} \\
&+ \frac{\text{cell} \cdot \text{kRaf1ByAkt} \cdot [\text{AktActive}] \cdot [\text{Raf1Active}]}{([\text{Raf1Active}] + \text{KmRaf1ByAkt})} \\
&- \frac{\text{cell} \cdot \text{kRasToRaf1} \cdot [\text{RasActive}] \cdot [\text{Raf1Inactive}]}{([\text{Raf1Inactive}] + \text{KmRasToRaf1})} \\
\frac{d[\text{Raf1Active}]}{dt} &= \frac{\text{cell} \cdot \text{kRasToRaf1} \cdot [\text{RasActive}] \cdot [\text{Raf1Inactive}]}{([\text{Raf1Inactive}] + \text{KmRasToRaf1})} \\
&- \frac{\text{cell} \cdot \text{kRaf1} \cdot [\text{Raf1PPtase}] \cdot [\text{Raf1Active}]}{([\text{Raf1Active}] + \text{KmdRaf1})} \\
&- \frac{\text{cell} \cdot \text{kRaf1ByAkt} \cdot [\text{AktActive}] \cdot [\text{Raf1Active}]}{([\text{Raf1Active}] + \text{KmRaf1ByAkt})} \\
\frac{d[\text{BRafInactive}]}{dt} &= \frac{\text{cell} \cdot \text{kBRaf} \cdot [\text{Raf1PPtase}] \cdot [\text{BRafActive}]}{([\text{BRafActive}] + \text{KmdBRaf})} \\
&- \frac{\text{cell} \cdot \text{kRap1ToBRaf} \cdot [\text{Rap1Active}] \cdot [\text{BRafInactive}]}{([\text{BRafInactive}] + \text{KmRap1ToBRaf})} \\
\frac{d[\text{BRafActive}]}{dt} &= \frac{\text{cell} \cdot \text{kRap1ToBRaf} \cdot [\text{Rap1Active}] \cdot [\text{BRafInactive}]}{([\text{BRafInactive}] + \text{KmRap1ToBRaf})} \\
&- \frac{\text{cell} \cdot \text{kBRaf} \cdot [\text{Raf1PPtase}] \cdot [\text{BRafActive}]}{([\text{BRafActive}] + \text{KmdBRaf})} \\
\frac{d[\text{MekInactive}]}{dt} &= \frac{\text{cell} \cdot \text{kMek} \cdot [\text{PP2AActive}] \cdot [\text{MekActive}]}{([\text{MekActive}] + \text{KmdMek})} \\
&- \frac{\text{cell} \cdot \text{kpRaf1} \cdot [\text{Raf1Active}] \cdot [\text{MekInactive}]}{([\text{MekInactive}] + \text{KmpRaf1})} \\
&- \frac{\text{cell} \cdot \text{kpBRaf} \cdot [\text{BRafActive}] \cdot [\text{MekInactive}]}{([\text{MekInactive}] + \text{KmpBRaf})} \\
\frac{d[\text{MekActive}]}{dt} &= \frac{\text{cell} \cdot \text{kpRaf1} \cdot [\text{Raf1Active}] \cdot [\text{MekInactive}]}{([\text{MekInactive}] + \text{KmpRaf1})} \\
&+ \frac{\text{cell} \cdot \text{kpBRaf} \cdot [\text{BRafActive}] \cdot [\text{MekInactive}]}{([\text{MekInactive}] + \text{KmpBRaf})} \\
&- \frac{\text{cell} \cdot \text{kMek} \cdot [\text{PP2AActive}] \cdot [\text{MekActive}]}{([\text{MekActive}] + \text{KmdMek})} \\
\frac{d[\text{ErkInactive}]}{dt} &= \frac{\text{cell} \cdot \text{kErk} \cdot [\text{PP2AActive}] \cdot [\text{ErkActive}]}{([\text{ErkActive}] + \text{KmdErk})} \\
&- \frac{\text{cell} \cdot \text{kpMekCytoplasmic} \cdot [\text{MekActive}] \cdot [\text{ErkInactive}]}{([\text{ErkInactive}] + \text{KmpMekCytoplasmic})} \\
\frac{d[\text{ErkActive}]}{dt} &= \frac{\text{cell} \cdot \text{kpMekCytoplasmic} \cdot [\text{MekActive}] \cdot [\text{ErkInactive}]}{([\text{ErkInactive}] + \text{KmpMekCytoplasmic})} \\
&- \frac{\text{cell} \cdot \text{kErk} \cdot [\text{PP2AActive}] \cdot [\text{ErkActive}]}{([\text{ErkActive}] + \text{KmdErk})} \\
\frac{d[\text{PI3KInactive}]}{dt} &= - \left(\frac{\text{cell} \cdot \text{kPI3K} \cdot [\text{boundEGFReceptor}] \cdot [\text{PI3KInactive}]}{([\text{PI3KInactive}] + \text{KmPI3K})} \right) \\
&- \frac{\text{cell} \cdot \text{kPI3KRas} \cdot [\text{RasActive}] \cdot [\text{PI3KInactive}]}{([\text{PI3KInactive}] + \text{KmPI3KRas})} \\
\frac{d[\text{PI3KActive}]}{dt} &= \frac{\text{cell} \cdot \text{kPI3K} \cdot [\text{boundEGFReceptor}] \cdot [\text{PI3KInactive}]}{([\text{PI3KInactive}] + \text{KmPI3K})} \\
&+ \frac{\text{cell} \cdot \text{kPI3KRas} \cdot [\text{RasActive}] \cdot [\text{PI3KInactive}]}{([\text{PI3KInactive}] + \text{KmPI3KRas})}
\end{aligned}$$

$$\begin{aligned}
\frac{d[\text{AktInactive}]}{dt} &= - \left(\frac{\text{cell} \cdot k_{\text{Akt}} \cdot [\text{PI3KActive}] \cdot [\text{AktInactive}]}{([\text{AktInactive}] + K_{m\text{Akt}})} \right) \\
\frac{d[\text{AktActive}]}{dt} &= \frac{\text{cell} \cdot k_{\text{Akt}} \cdot [\text{PI3KActive}] \cdot [\text{AktInactive}]}{([\text{AktInactive}] + K_{m\text{Akt}})} \\
\frac{d[\text{C3GInactive}]}{dt} &= - \left(\frac{\text{cell} \cdot k_{\text{C3GNGF}} \cdot [\text{boundNGFReceptor}] \cdot [\text{C3GInactive}]}{([\text{C3GInactive}] + K_{m\text{C3GNGF}})} \right) \\
\frac{d[\text{C3GActive}]}{dt} &= \frac{\text{cell} \cdot k_{\text{C3GNGF}} \cdot [\text{boundNGFReceptor}] \cdot [\text{C3GInactive}]}{([\text{C3GInactive}] + K_{m\text{C3GNGF}})} \\
\frac{d[\text{Rap1Inactive}]}{dt} &= \frac{\text{cell} \cdot k_{\text{RapGap}} \cdot [\text{RapGapActive}] \cdot [\text{Rap1Active}]}{([\text{Rap1Active}] + K_{m\text{RapGap}})} \\
&\quad - \frac{\text{cell} \cdot k_{\text{C3G}} \cdot [\text{C3GActive}] \cdot [\text{Rap1Inactive}]}{([\text{Rap1Inactive}] + K_{m\text{C3G}})} \\
\frac{d[\text{Rap1Active}]}{dt} &= \frac{\text{cell} \cdot k_{\text{C3G}} \cdot [\text{C3GActive}] \cdot [\text{Rap1Inactive}]}{([\text{Rap1Inactive}] + K_{m\text{C3G}})} \\
&\quad - \frac{\text{cell} \cdot k_{\text{RapGap}} \cdot [\text{RapGapActive}] \cdot [\text{Rap1Active}]}{([\text{Rap1Active}] + K_{m\text{RapGap}})}
\end{aligned}$$

Optimizable Parameters

krbEGF	2.18503e - 05
kruEGF	0.0121008
krbNGF	1.38209e - 07
kruNGF	0.00723811
kEGF	694.731
KmEGF	6086070.0
kNGF	389.428
KmNGF	2112.66
kdSos	1611.97
KmdSos	896896.0
kSos	32.344
KmSos	35954.3
kRasGap	1509.36
KmRasGap	1432410.0
kRasToRaf1	0.884096
KmRasToRaf1	62464.6
kpRaf1	185.759
KmpRaf1	4768350.0
kpBRaf	125.089
KmpBRaf	157948.0
kdMek	2.83243
KmdMek	518753.0
kpMekCytoplasmic	9.85367
KmpMekCytoplasmic	1007340.0
kdErk	8.8912
KmdErk	3496490.0
kpP90Rsk	0.0213697

KmpP90Rsk	763523.0
kPI3K	10.6737
KmPI3K	184912.0
kPI3KRas	0.0771067
KmPI3KRas	272056.0
kAkt	0.0566279
KmAkt	653951.0
kdRaf1ByAkt	15.1212
KmRaf1ByAkt	119355.0
kC3GNGF	146.912
KmC3GNGF	12876.2
kC3G	1.40145
KmC3G	10965.6
kRapGap	27.265
KmRapGap	295990.0
kRap1ToBRaf	2.20995
KmRap1ToBRaf	1025460.0
kdRaf1	0.126329
KmdRaf1	1061.71
kdBRaf	441.287
KmdBRaf	10879500.0