

# 1 Equations

$$\begin{aligned}
\frac{dLeish_{promas}(t)}{dt} &= \frac{-\log(0.01)}{24} \text{Leish\_promas}(t) \\
\frac{dTLR4(t)}{dt} &= c_{TLR4} + p64 \text{STAT3}(t) - \text{TLR4}(t) \max(p64 + 0.1c_{TLR4}, 0.0001) \\
\frac{dTLR4a(t)}{dt} &= c_{TLR4a} - \text{TLR4a}(t) \max(0.1c_{TLR4a}, 0.0001) + \frac{100p2\text{Leish\_promas}(t) \text{TLR4}(t)}{kntr + \text{Leish\_promas}(t)} \\
\frac{dNOTCH3(t)}{dt} &= c_{NOTCH3} + p3\text{TLR4a}(t) - \text{NOTCH3}(t) \max(p3 + 0.1c_{NOTCH3}, 0.0001) \\
\frac{dTRAF6(t)}{dt} &= c_{TRAF6} + (-0.1c_{TRAF6} + 0.001p5 + 10p7) \text{TLR4a}(t) \\
&\quad - \text{TRAF6}(t) \max(0.001p5, 0.0001) - p7\text{NLRP12}(t) \text{TRAF6}(t) \\
\frac{dMAP4K3(t)}{dt} &= c_{MAP4K3} + p9\text{TRAF6}(t) - \text{MAP4K3}(t) \max(p9 + 0.1c_{MAP4K3}, 0.0001) \\
\frac{dNFkB(t)}{dt} &= c_{NFkB} + p10\text{MAP4K3}(t) - \text{NFkB}(t) \max(p10 + 0.1c_{NFkB}, 0.0001) \\
&\quad - p51\text{Leish\_amas}(t) \text{NFkB}(t) \\
\frac{dIL1B(t)}{dt} &= c_{IL1B} + p11\text{NFkB}(t) + p12\text{P2RX7a}(t) - \text{IL1B}(t) \max(p11 + p12 + 0.1c_{IL1B}, 0.0001) \\
\frac{dTnFA(t)}{dt} &= c_{TNFA} + p13\text{NFkB}(t) - \text{TNFA}(t) \max(p13 + 0.1c_{TNFA}, 0.0001) \\
\frac{dCREB(t)}{dt} &= c_{CREB} + p15\text{MAP4K3}(t) + p16\text{ADORA2Ba}(t) \\
&\quad - \text{CREB}(t) \max(p15 + p16 + 0.1c_{CREB}, 0.0001) \\
\frac{dIL10(t)}{dt} &= c_{IL10} + p17\text{CREB}(t) - \text{IL10}(t) \max(p17 + 0.1c_{IL10}, 0.0001) \\
\frac{dSTAT3(t)}{dt} &= c_{STAT3} + p19\text{IL10}(t) - \text{STAT3}(t) \max(p19 + 0.1c_{STAT3}, 0.0001) \\
&\quad - lpst\text{Leish\_promas}(t) \text{STAT3}(t) \\
\frac{dNLRP12(t)}{dt} &= c_{NLRP12} + p20\text{STAT3}(t) - \text{NLRP12}(t) \max(p20 + 0.1c_{NLRP12}, 0.0001) \\
\frac{dCCL24(t)}{dt} &= c_{CCL24} + p22\text{STAT3}(t) - \text{CCL24}(t) \max(p22 + 0.1c_{CCL24}, 0.0001) \\
\frac{dTIGAR(t)}{dt} &= c_{TIGAR} + p24\text{CREB}(t) - \text{TIGAR}(t) \max(p24 + 0.1c_{TIGAR}, 0.0001) \\
\frac{dFOS(t)}{dt} &= c_{FOS} + p26\text{MAP4K3}(t) + p65\text{STAT3}(t) - \text{FOS}(t) \max(p26 + p65 + 0.1c_{FOS}, 0.0001) \\
\frac{dNADP_{oxi}(t)}{dt} &= c_{NADP_{oxi}} + \text{FOS}(t) (p27 + 10p28) - \text{NADP\_oxi}(t) \max(p27 + 0.1c_{NADP_{oxi}}, 0.0001) \\
&\quad - p28\text{NADP\_oxi}(t) \text{NOTCH3}(t)
\end{aligned}$$

$$\begin{aligned}
\frac{dROS_{cytos}(t)}{dt} &= nftf3Leish\_promas(t) NADP\_oxi(t) + nftf4Leish\_amas(t) NADP\_oxi(t) \\
&\quad - p57Glutathione(t) ROS\_cytos(t) \\
\frac{dLeish_{amas}(t)}{dt} &= 0.096Leish\_promas(t) \\
\frac{dP2RX7(t)}{dt} &= c_{P2RX7} + p32STAT3(t) - P2RX7(t) \max(p32 + 0.1c_{P2RX7}, 0.0001) \\
\frac{dP2RX7a(t)}{dt} &= c_{P2RX7a} - P2RX7a(t) \max\left(0.1c_{P2RX7a} + \frac{10.0p33}{10.0 + kilp2r}, 0.0001\right) \\
&\quad + \frac{p33ATP\_ext(t) P2RX7(t)}{kilp2r + ATP\_ext(t)} \\
\frac{dNFE2L2(t)}{dt} &= c_{NFE2L2} + p34NFkB(t) + p35ROS\_Mito(t) - NFE2L2(t) \max(p34 + 0.1c_{NFE2L2}, 0.0001) \\
\frac{dGlutathione(t)}{dt} &= c_{Glutathione} + p36NFE2L2(t) - Glutathione(t) \max(p36 + 0.1c_{Glutathione}, 0.0001) \\
\frac{dPANX(t)}{dt} &= c_{PANX} + pcrebtrCREB(t) - PANX(t) \max(pcrebtr + 0.1c_{PANX} - 10.0n_{ext\_tr}, 0.0001) \\
&\quad - n_{ext\_tr}ATP\_ext(t) PANX(t) \\
\frac{dPANXa(t)}{dt} &= c_{PANXa} - PANXa(t) \max(0.1c_{PANXa}, 0.0001) + \frac{k_{PANXa}ATP\_cytos(t) Leish\_promas(t) PANX(t)}{katppnx + ATP\_cytos(t)} \\
\frac{dATP_{ext}(t)}{dt} &= p61PANXa(t) - ATP\_ext(t) \max(p61, 0.0001) \\
\frac{dNT5E(t)}{dt} &= c_{NT5E} + p66STAT3(t) + p67HIF1A(t) + p68NFkB(t) - NT5E(t) \max(p66 + p67 + p68 + 0.1c_{NT5E}, 0.0001) \\
\frac{dAdenosine(t)}{dt} &= -Adenosine(t) \max\left(\frac{10.0v38}{10.0 + k38}, 0.0001\right) + \frac{v38ATP\_ext(t) NT5E(t)}{k38 + ATP\_ext(t)} \\
\frac{dADORA2B(t)}{dt} &= c_{ADORA2B} + p68HIF1A(t) - ADORA2B(t) \max(p68 + 0.1c_{ADORA2B}, 0.0001) \\
\frac{dADORA2Ba(t)}{dt} &= c_{ADORA2Ba} - ADORA2Ba(t) \max\left(0.1c_{ADORA2Ba} + \frac{10.0i_{ADAd}}{10.0 + kadocreb}, 0.0001\right) + \\
&\quad \frac{i_{ADAd}ADORA2B(t) Adenosine(t)}{kadocreb + Adenosine(t)} \\
\frac{dATP_{cytos}(t)}{dt} &= p18Glycolysis(t) - ATP\_cytos(t) \max(p18, 0.0001) - p37ATP\_cytos(t) Leish\_promas(t) PANX(t) \\
\frac{dGlycolysis(t)}{dt} &= c_{Glycolysis} + (p43 - 0.1c_{Glycolysis} + 10p41) HIF1A(t) - Glycolysis(t) \max(p43, 0.0001) \\
&\quad - p41Glycolysis(t) TIGAR(t) \\
\frac{dPyruvate(t)}{dt} &= Glycolysis(t) (p44 + p45) - p44Pyruvate(t) - p45Pyruvate(t) \\
\frac{dSuccinate(t)}{dt} &= c_{Succinate} + p45_{prop}Pyruvate(t) - p60Succinate(t) - Succinate(t) \max(p45_{prop} + 0.1c_{Succinate} - p60, 0.0001)
\end{aligned}$$

$$\begin{aligned}
\frac{dTCA(t)}{dt} &= p60_{prop} \text{Succinate}(t) - TCA(t) \max(p60_{prop}, 0.0001) - p55 \text{ROS\_Mito}(t) TCA(t) \\
\frac{dQH2(t)}{dt} &= c_{QH2} + p46 \text{Succinate}(t) - QH2(t) \max(p46 + 0.1c_{QH2}, 0.0001) \\
\frac{dATP_{Mito}(t)}{dt} &= o_{ATP\_Mito} + (p47 - 0.1o_{ATP\_Mito} + 10p53) QH2(t) - ATP\_Mito(t) \max(p47, 0.0001) \\
&\quad - p48 ATP\_Mito(t) \text{ROS\_Mito}(t) - p53 ATP\_Mito(t) \text{HIF1A}(t) \\
\frac{dROS_{Mito}(t)}{dt} &= -d_{ROS\_Mito} \text{ROS\_Mito}(t) + p21 \text{Glycolysis}(t) \text{Leish\_promas}(t) \\
\frac{dHIF1A(t)}{dt} &= c_{HIF1A} + p49 \text{NFkB}(t) + p62 \text{Leish\_promas}(t) \text{Succinate}(t) - \text{HIF1A}(t) \max(p49 + 0.1c_{HIF1A}, 0.0001) \\
\frac{dLactate(t)}{dt} &= c_{Lactate} + p44_{prop} \text{Pyruvate}(t) + p50 \text{HIF1A}(t) - \text{Lactate}(t) \max(p44_{prop} + p50 + 0.1c_{Lactate}, 0.0001)
\end{aligned}$$

## 2 Parameters

$p64 = 3.21$   
 $c_{TLR4} = 0.0246$   
 $kntr = 1.64$   
 $p2 = 17.8$   
 $c_{TLR4a} = 42.4$   
 $p3 = 0.0386$   
 $c_{NOTCH3} = 0.00401$   
 $c_{TRAF6} = 1.22$   
 $p7 = 0.00158$   
 $p5 = 951.0$   
 $c_{MAP4K3} = 0.000455$   
 $p9 = 378.0$   
 $c_{NFkB} = 0.0291$   
 $p10 = 1.81$   
 $p51 = 0.184$   
 $p12 = 4.02e - 5$   
 $p11 = 0.987$   
 $c_{IL1B} = 0.0669$   
 $c_{TNFA} = 0.298$   
 $p13 = 1.19$   
 $p15 = 0.0514$   
 $p16 = 0.319$   
 $c_{CREB} = 0.000878$   
 $p17 = 43.7$   
 $c_{IL10} = 0.31$   
 $p19 = 0.159$   
 $lpst = 13.1$   
 $c_{STAT3} = 0.000127$   
 $c_{NLRP12} = 3.05e - 5$   
 $p20 = 0.136$   
 $c_{CCL24} = 0.00737$   
 $p22 = 0.153$   
 $c_{TIGAR} = 0.0177$

$p24 = 0.724$   
 $p26 = 0.252$   
 $p65 = 0.0339$   
 $c_{FOS} = 0.0139$   
 $c_{NADP_{oxi}} = 1.45$   
 $p27 = 0.102$   
 $p28 = 2.01e - 5$   
 $n_{ftf3} = 0.258$   
 $n_{ftf4} = 2.22$   
 $p57 = 0.0397$   
 $p32 = 0.33$   
 $c_{P2RX7} = 2.26$   
 $p33 = 455.0$   
 $k_{ilp2r} = 1.61$   
 $c_{P2RX7a} = 0.978$   
 $p34 = 1.22$   
 $p35 = 3.43$   
 $c_{NFE2L2} = 0.92$   
 $c_{Glutathione} = 20.6$   
 $p36 = 0.159$   
 $n_{ext-tr} = 0.0169$   
 $p_{crebtr} = 0.191$   
 $c_{PANX} = 0.0815$   
 $k_{atppnx} = 0.242$   
 $k_{PANXa} = 0.314$   
 $c_{PANXa} = 0.827$   
 $p61 = 0.377$   
 $p66 = 0.0336$   
 $p68 = 0.0139$   
 $c_{NT5E} = 0.0055$   
 $p67 = 0.388$   
 $v38 = 3.18$   
 $k38 = 0.00369$   
 $c_{ADORA2B} = 1.65$   
 $i_{ADAd} = 1.32$   
 $k_{adocreb} = 22.1$   
 $c_{ADORA2Ba} = 0.878$   
 $p18 = 1.35$   
 $p37 = 1.11$   
 $p41 = 1.8$

$c_{Glycolysis} = 0.186$   
 $p43 = 0.178$   
 $p44 = 0.656$   
 $p45 = 0.596$   
 $p60 = 0.515$   
 $c_{Succinate} = 0.194$   
 $p45_{prop} = 13.8$   
 $p55 = 0.0178$   
 $p60_{prop} = 3.8$   
 $p46 = 0.00841$   
 $c_{QH2} = 0.00685$   
 $o_{ATP\_Mito} = 0.635$   
 $p47 = 49.8$   
 $p53 = 0.334$   
 $p48 = 0.00279$   
 $p21 = 0.012$   
 $d_{ROS\_Mito} = 0.385$   
 $p49 = 0.47$   
 $p62 = 0.885$   
 $c_{HIF1A} = 1.67e - 5$   
 $p50 = 0.000463$   
 $c_{Lactate} = 0.321$   
 $p44_{prop} = 0.0341$