

Full Model Code

The model used in this paper was created using a software package called Berkeley MadonnaTM. (Berkeley MadonnaTM can be downloaded as a free trial version at <http://www.berkeleymadonna.com/download.html>)

Below is the full Berkeley MadonnaTM code for the model.

In order to run the model, the following steps should be followed:

- 1) Replace the terms β , γ , δ , μ , π , ρ , σ and τ with their names e.g. beta, gamma, etc.
- 2) Cut and paste the entire code into the Equations window of Berkeley MadonnaTM
- 3) Run the code with T_{BI} (total number of blood stage infections), N_{ra} (number of infections resistant to artesunate) and $perc_{ra}$ (percentage of infections resistant to artesunate) displayed in the Graph window
- 4) Use the parameter sliders to alter parameter values to explore their effects on the results

STARTTIME = 1960

STOPTIME=2050

DT = 0.0027

;Model to examine the effect of elimination interventions on artesunate resistance in Western Cambodia

;Dr Richard Maude 27/11/08

;All times are in years

;Artesunate monotherapy begins in 1975

;Artesunate resistance first arises in 1980

;Interventions begin in 2009

;NOTATION

;N = total population size ($N = S+L+B+I$)

;t = time for drug to clear sensitive infection

;c = clearance rate of infection by drug according to drug sensitivity

;S = susceptible people

;L = liver stage infections

;B = noninfectious blood stage infections

;T = total

;I = infectious blood stage infections

;r = resistant to...

;d = drug given

;o = no drug

;a = artesunate

;ai = artesunate as part of ACT

;b = piperaquine

;ab = ACT (dihydroartemisinin plus piperaquine)

;p = primaquine

;e.g.1 t_{Bda} = time for artesunate to clear noninfectious blood stage infection

;e.g.2 c_{Bradab} = clearance rate (c) of noninfectious blood stage infections (B) resistant to artesunate (ra) by ACT (dab)

;e.g.3 c_{Irod_p} = clearance rate (c) of infectious blood stage infections (I) resistant to no drugs i.e. sensitive to all (ro) by primaquine (dp)

;i₀ = no intervention

;i₁ = mass drug administration (MDA) (Intervention 1)

;i_{1a} = treatment of symptomatic patients who have already received MDA

;i₂ = masss screen and treat (MSAT) with atovaquone/proguanil (Intervention 2)

;i₃ = MSAT with ACT (Intervention 3)

;e.g.4 d_{ai1a} = treatment of symptomatic patients with artesunate monotherapy (d_a) who have already received MDA

;e.g.5 d_{abi1a} = treatment of symptomatic patients with ACT who have already received MDA

;X = effective duration of drug action

;e.g.6 X_{ai} = effective duration of artesunate as part of ACT

;e = relative effectiveness of drug against drug resistant infections vs drug sensitive infections

;bn = bed nets

;dur = duration

;PARAMETERS

;Population parameters

$\mu = 0.015$; i.e. birth rate & death rate

$N_0 = 3200000$; total population at time=0

$I_0 = N_0 * p_{inf}$; total infected at time=0

;Natural history of infection

$\gamma = 365/5$; rate of liver stage becoming noninfectious blood stage

$\sigma = 365/15$; rate of developing gametocytes

$\delta = 365/60$; natural recovery rate of infectious

;Drug action

;Time for drug to clear drug sensitive infection in days

$t_{Bda} = 7$; time for artesunate to clear B

$t_{Ida} = 4$; time for artesunate to clear I

$t_{Bdb} = 3$; time for clearance of noninfectious blood stage by piperaquine

$t_{Idb} = 21$; time for clearance of infectious blood stage by piperaquine

$t_{Bdab} = 3$; time for ACT to clear B = 3 days if synergy or 7 days if none

$t_{Idab} = 3$; time for ACT to clear I = 3 days if synergy or 4 days if none

$t_{Ldvg} = 3$; time for clearance of liver stage by atovaquone/proguanil

$t_{Bdvg} = 3$; time for clearance of noninfectious blood stage by atovaquone/proguanil

$t_{Idvg} = 4.5$; time for clearance of infectious blood stage by atovaquone/proguanil

$t_{Ldv} = 6$; time for clearance of liver stage by atovaquone

$t_{Bdv} = 3$; time for clearance of noninfectious blood stage by atovaquone

$t_{Idv} = 4.5$; time for clearance of infectious blood stage by atovaquone

$t_{Ldp} = 7$; time for clearance of liver stage by primaquine

$t_{Idp} = 1$; time for clearance of infectious blood stage by primaquine

;rate of clearance of infection by drug according to drug sensitivity

$c_{Brodo} = 0$; rate of clearance of sensitive noninfectious blood stage by no drug

$c_{Brado} = 0$; rate of clearance of artesunate resistant noninfectious blood stage by no drug

$c_{Brbdo} = 0$; rate of clearance of piperaquine resistant noninfectious blood stage by no drug

$c_{Irodo} = 0$; rate of clearance of sensitive infectious blood stage by no drug

$c_{Irado} = 0$; rate of clearance of artesunate resistant infectious blood stage by no drug

$c_{Irpdo} = 0$; rate of clearance of piperaquine resistant infectious blood stage by no drug

$c_{Broda} = 365/t_{Bda}$; rate of clearance of blood stage resistant to none treated with artesunate

$c_{Brada} = e_{rada} * c_{Broda}$; rate of clearance of blood stage resistant to artesunate treated with artesunate

$c_{Brbda} = 365/t_{Bdb}$; rate of clearance of blood stage resistant to piperaquine treated with artesunate

$c_{Iroda} = 365/t_{Ida}$; rate of clearance of I resistant to none treated with artesunate

$c_{Irada} = e_{rada} * c_{Iroda}$; rate of clearance of I resistant to artesunate treated with artesunate

$c_{Irnda} = 365/t_{Idb}$; rate of clearance of I resistant to piperaquine treated with artesunate

$c_{Brodab} = 365/t_{Bdab}$; rate of clearance of B resistant to none treated by ACT

$c_{Bradab} = e_{rada} * 365/t_{Bdab} + (1-e_{rada}) * 365/t_{Bdb}$; rate of clearance of artesunate resistant noninfectious blood stage by ACT

$c_{Brbdab} = e_{rbdb} * 365/t_{Bdab} + (1-e_{rbdb}) * 365/t_{Bda}$; rate of clearance of piperaquine resistant noninfectious blood stage by ACT

$c_{Irodab} = 365/t_{Idab}$; rate of clearance of sensitive infectious blood stage by ACT

$c_{Irtradab} = e_{rada} * 365/t_{Idab} + (1-e_{rada}) * 365/t_{Idb}$; rate of clearance of artesunate resistant infectious blood stage by ACT

$c_{Irbdab} = e_{rbdb} * 365/t_{Idab} + (1-e_{rbdb}) * 365/t_{Ida}$; rate of clearance of piperaquine resistant infectious blood stage by ACT

$c_{Brodb} = 365/t_{Bdb}$; rate of clearance of sensitive noninfectious blood stage treated with piperaquine (after artesunate i.e. minus 3 days)

$c_{Bradb} = 365/t_{Bdb}$; rate of clearance of artesunate resistant noninfectious blood stage treated with piperaquine

$c_{Brbdb} = e_{rbdb} * 365/t_{Bdb}$; rate of clearance of piperaquine resistant noninfectious blood stage treated with piperaquine

$c_{Irodb} = 365/t_{Idb}$; rate of clearance of sensitive infectious blood stage treated with piperaquine

$c_{Iradb} = 365/t_{Idb}$; rate of clearance of artesunate resistant infectious blood stage treated with piperaquine

$c_{Irbdb} = e_{rbdb} * 365/t_{Idb}$; rate of clearance of piperaquine resistant infectious blood stage treated with piperaquine

$c_{Lp} = p_p * 365/t_{Ldp}$; rate of clearance of liver stage treated with primaquine

$c_{Ip} = p_p * 365/t_{Idp}$; rate of clearance of infectious blood stage treated with primaquine

$c_{Ldvg} = 365/t_{Ldvg}$; rate of clearance of liver stage treated with atovaquone/proguanil

$c_{Bdvg} = 365/t_{Bdvg}$; rate of clearance of noninfectious blood stage treated with atovaquone/proguanil

$c_{Idvg} = 365/t_{Idvg}$; rate of clearance of infectious blood stage treated with atovaquone/proguanil

$c_{Ldv} = 365/t_{Ldv}$; rate of clearance of liver stage treated with atovaquone after atovaquone/proguanil

$c_{Bdv} = 365/t_{Bdv}$; rate of clearance of noninfectious blood stage treated with atovaquone after atovaquone/proguanil

$c_{Idv} = 365/t_{Idv}$; rate of clearance of infectious blood stage treated with atovaquone after atovaquone/proguanil

;Effective duration of drug action

$X_{ao} = 7/365$; effective duration of artesunate monotherapy

$X_{ai} = 3/365$; effective duration of artesunate as part of ACT

$X_b = 20/365$; effective duration of piperaquine

$X_g = 4/365$; effective duration of proguanil as part of atovaquone/proguanil

$X_v = 15/365$; effective duration of atovaquone

$X_p = 1/365$; effective duration of primaquine

;Drug resistance

$e_{rada} = (1 - prec_{ra}) * pct_{roda}/pct_{rada}$; relative effectiveness of artesunate against artesunate resistant infections (0-1)

$pct_{roda} = 30$; parasite clearance time for artesunate vs sensitive infections

$pct_{rada} = 83$; parasite clearance time for artesunate vs resistant infections

$prec_{ra} = 0.35$; proportion recrudescences in resistant infections

$e_{rbdb} = 0.8$; relative effectiveness of piperaquine in piperaquine resistant infections (0-1)

;Rates of receiving treatments

;Artesunate monotherapy

$\tau = 365/16 * propRx_a * \text{SQUAREPULSE}(start_a, dur_a)$; rate of starting artesunate in infected patients

$dur_a = stop_a - start_a$; duration availability of artesunate

$start_a = 1975$; time that artesunate first introduced

$stop_a = 2009$; time from when artesunate no longer available

;Treat symptomatic infection with ACT

$\tau_{ab} = \tau_{mag_{ab}} * propRx_{ab} * \text{SQUAREPULSE}(start_{ab}, dur_{ab})$

$\tau_{mag_{ab}} = 365/16$

$\text{dur}_{ab} = 41$; 'long-term'
 $\text{start}_{ab} = 2009$;

;Intervention 1: MDA with ACT
 $\tau_{pulse_1} = \text{if mod}(\text{time}-\text{durt}_1, 1) \geq 1 - \text{durt}_1 \text{ then propRx}_{i1} * \tau_{mag_{i1}} \text{ else 0}$
 $\tau_1 = \tau_{pulse_1} * \text{SQUAREPULSE}(2009, \text{dur}_{i1})$
 $\tau_{mag_{i1}} = 4$; rate of starting intervention 1 in 1/years
 $\text{dur}_{i1} = 0$; total duration of intervention 1 in years
 $\text{durt}_1 = 0.25$; duration of each round of intervention 1 in years

;Intervention 2: MSAT with atovaquone/proguanil
 $\tau_{pulse_2} = \text{if mod}(\text{time}-\text{durt}_2, \text{fr}_{i2}) \geq \text{fr}_{i2} - \text{durt}_2 \text{ then propRx}_{i2} * \tau_{mag_{i2}} \text{ else 0}$
 $\tau_2 = \tau_{pulse_2} * \text{SQUAREPULSE}(2009, \text{dur}_{i2})$
 $\tau_{mag_{i2}} = 4$; rate of starting intervention 2 in 1/years
 $\text{dur}_{i2} = 0$; total duration of intervention 2 in years
 $\text{durt}_2 = 0.25$; duration of each round of intervention 2 in years
 $\text{fr}_{i2} = 1/n_{i2}$; 1/frequency of intervention 2 per year
 $n_{i2} = 1$; frequency of intervention 2 per year

;Intervention 3: MSAT with ACT.
;this uses the same section of code as switching treatment to ACT
; to make this run, do the following:
;1) add a ';' to the beginning of the first line of the 'treat symptomatic infection with ACT' section (above) to give "; $\tau_{ab} = \tau_{mag_{ab}} * \text{propRx}_{ab} * \text{SQUAREPULSE}(\text{start}_{ab}, \text{dur}_{ab})$ "
 $\tau_{pulse_3} = \text{if mod}(\text{time}-\text{durt}_3, 1) \geq 1 - \text{durt}_3 \text{ then propRx}_{i3} * \tau_{mag_{i3}} \text{ else 0}$
;2) remove the ';' from the beginning of the line "; $\tau_{ab} = \tau_{pulse_3} * \text{SQUAREPULSE}(2009, \text{dur}_{i3})$ " (below)
 $\tau_{pulse_3} = \text{if mod}(\text{time}-\text{durt}_3, 1) \geq 1 - \text{durt}_3 \text{ then propRx}_{i3} * \tau_{mag_{i3}} \text{ else 0}$
; $\tau_{ab} = \tau_{pulse_3} * \text{SQUAREPULSE}(2009, \text{dur}_{i3})$
 $\tau_{mag_{i3}} = 4$; rate of starting intervention 3 in 1/years
 $\text{dur}_{i3} = 0$; total duration of intervention 3 in years
 $\text{durt}_3 = 0.25$; duration of each round of intervention 3 in years

;ACT for symptomatic infection during intervention 1
 $\tau_{abi1} = \tau_{mag_{abi1}} * \text{propRx}_{ab} * \text{SQUAREPULSE}(\text{start}_{ab}, \text{dur}_{ab})$; rate of starting ACT in infected people during intervention 1
 $\tau_{mag_{abi1}} = 365/16$

;Artesunate monotherapy for symptomatic infection during intervention 1
 $\tau_{ai1} = \tau_{mag_{ai1}} * \text{propRx}_a * \text{SQUAREPULSE}(2009, \text{dur}_a)$; rate of starting artesunate in infected people during intervention 1
 $\tau_{mag_{ai1}} = 365/16$

;ACT for symptomatic infection during intervention 2

$\tau_{abi2} = \tau_{mag_{abi2}} * \text{propRx}_{ab} * \text{SQUAREPULSE}(2009, \text{dur}_{ab})$; rate of starting ACT during intervention 2

$\tau_{mag_{abi2}} = 365/16$

;Artesunate monotherapy for symptomatic infection during intervention 2

$\tau_{ai2} = \tau_{mag_{ai2}} * \text{propRx}_a * \text{SQUAREPULSE}(2009, \text{dur}_a)$; rate of starting artesunate in infected people during intervention 2

$\tau_{mag_{ai2}} = 365/16$

;Bed nets

$bn = bn_{mag} * \text{SQUAREPULSE}(2009, \text{dur}_{bn})$

$bn_{mag} = 0.3$; transmission reduction due to bed nets

$\text{dur}_{bn} = 0$; 0 or 4 years

;Initial conditions

$p_a = 0$; proportion resistant to artesunate in 1960

$p_b = 0.05$; proportion resistant to piperaquine in 1960 (and 2009)

$p_o = 1 - p_a - p_b$; proportion sensitive to artesunate and piperaquine in 1960

$p_{BI} = 0.0743$; proportion infected with detectable blood stage infection in 2009 (data is from 2006)

$p_{inf} = 0.16$; proportion any malaria infection in 1960

$p_p = 0$; proportion given primaquine = 0-1

$st = 1980$; time that artesunate resistance first arises - this is set relatively early to maximise p_a in 2009 so near 10%

$mst = 350$;

$\text{propRx}_a = \text{propRx}_{am} * \text{prop}_a * \text{adh}_a$; proportion that get effective artesunate treatment for symptomatic infection = 0.052

$\text{propRx}_{am} = 0.63$; proportion receiving antimalarials

$\text{prop}_a = 0.4$; proportion of antimalarials constituting artesunate monotherapy

$\text{adh}_a = 0.2$; proportion that take full 7 day course artesunate monotherapy

$\text{propRx}_{ab} = \text{IF TIME}<2009 \text{ THEN } 0 \text{ ELSE } p_{ab} * (\text{cov}_{ab} - \text{cov}_{ab} * \exp(-k * (\text{TIME} - 2009)))$; proportion that get effective ACT treatment for symptomatic infection

$\text{cov}_{ab} = 0.6$; maximum coverage with ACT for treatment

$p_{sab} = 0.85$; proportion of shops that sell modern drugs

$\text{adh}_{ab} = 0.77$; adherence to 3 day regime of ACT

$\text{adh}_{vg} = 0.77$; adherence to 3 day regime of atovaquone/proguanil

$p_{ab} = \text{cov}_{ab} * p_{sab} * \text{adh}_{ab}$; proportion that actually take 3 day course of ACT

$k = 30$; speed of introduction of intervention

$\text{propRx}_{i1} = \text{cov}_{i1} * \text{adh}_{ab}$; proportion that complete a 3 day course of MDA with ACT
 $\text{propRx}_{i2} = \text{cov}_{i2} * \text{adh}_{vg}$; proportion that complete a 3 day course of atovaquone/proguanil during MSAT
 $\text{propRx}_{i3} = \text{cov}_{i3} * \text{adh}_{ab}$; proportion that complete a 3 day course of ACT during MSAT
 $\text{cov}_{i1} = 0.8$; coverage with intervention 1
 $\text{cov}_{i2} = 0.8$; coverage with intervention 2
 $\text{cov}_{i3} = 0.8$; coverage with intervention 3

;Calculate intial conditions

$\text{ROOTI } \beta_n = \mu * N_0 - \beta_n * (p_{inf} * N_0) * S_{roodoio1} / N_0 + \delta * (p_{inf} * N_0) - \mu * S_{roodoio1}$
 $\text{ROOTI } S_{roodoio1} = \beta_n * (p_{inf} * N_0) * S_{roodoio1} / N_0 - (\gamma + \mu) * L_{roodoio1}$
 $\text{ROOTI } L_{roodoio1} = \gamma * L_{roodoio1} - (\sigma + \mu) * B_{roodoio1}$
 $\text{ROOTI } B_{roodoio1} = \sigma * B_{roodoio1} - (\delta + \mu) * (p_{inf} * N_0)$

GUESS $\beta_n = 10$

LIMIT $\beta_n \geq 0$

LIMIT $\beta_n \leq 1000$

GUESS $S_{roodoio1} = 2.1e+6$

LIMIT $S_{roodoio1} \geq 0$

LIMIT $S_{roodoio1} \leq N_0$

GUESS $L_{roodoio1} = 7e+4$

LIMIT $L_{roodoio1} \geq 0$

LIMIT $L_{roodoio1} \leq N_0$

GUESS $B_{roodoio1} = 2e+5$

LIMIT $B_{roodoio1} \geq 0$

LIMIT $B_{roodoio1} \leq N_0$

;MODEL

;Treatment with artesunate monotherapy

;N_{doio} and N_{daio}

;Box N_{doio}

init $S_{doio} = S_{roodoio1}$
 $I_{doio0} = N_0 * p_{inf}$

$$\begin{aligned}L_{\text{doio}} &= L_{\text{roodoio}} + L_{\text{radooio}} + L_{\text{rbodoio}} \\B_{\text{doio}} &= B_{\text{roodoio}} + B_{\text{radooio}} + B_{\text{rbodoio}} \\I_{\text{doio}} &= I_{\text{roodoio}} + I_{\text{radooio}} + I_{\text{rbodoio}}\end{aligned}$$

```
init Lroodoio = Lroodoio1
init Lradooio = 0
init Lrbodoio = 0
init Broodoio = Broodoio1
init Bradooio = 0
init Brbodoio = 0
init Iroodoio = po*Idoio0
init Iradooio = pa*Idoio0
init Irbodoio = pb*Idoio0
```

;S_{doio}

; Example of code for stochasticity S_{doio}(t+dt) = max(poission(S_{doio}+(μ*N₀ - μ*S_{doio} - β*S_{doio}*T_{Iro}/N₀ + δ*I_{roodoio} + c_{Brodo}*B_{roodoio} + c_{Irodo}*I_{roodoio} - β*S_{doio}*T_{Ira}/N₀ + δ*I_{radooio} + c_{Brado}*B_{radooio} + c_{Irado}*I_{radooio} - β*S_{doio}*T_{Ir_b}/N₀ + δ*I_{rbodoio} + c_{Brbdo}*B_{rbodoio} + c_{Irbd_o}*I_{rbodoio} + (1/X_{ao})*S_{daio} - τ₁*S_{doio} + (1/(X_b-X_{ai}))*S_{dbio})*dt),0)

; this form of code was substituted for every differential equation in the model to achieve stochasticity

d/dt(S_{doio}) = μ*N₀ - μ*S_{doio} - β*S_{doio}*T_{Iro}/N₀ + δ*I_{roodoio} + c_{Brodo}*B_{roodoio} + c_{Irodo}*I_{roodoio} - β*S_{doio}*T_{Ira}/N₀ + δ*I_{radooio} + c_{Brado}*B_{radooio} + c_{Irado}*I_{radooio} - β*S_{doio}*T_{Ir_b}/N₀ + δ*I_{rbodoio} + c_{Brbdo}*B_{rbodoio} + c_{Irbd_o}*I_{rbodoio} + (1/X_{ao})*S_{daio} - τ₁*S_{doio} + (1/(X_b-X_{ai}))*S_{dbio}

;roodoio

d/dt(L_{roodoio}) = β*S_{doio}*T_{Iro}/N₀ - (μ+γ)*L_{roodoio} + (1/X_{ao})*L_{roodaio} - τ₁*L_{roodoio} + (1/(X_b-X_{ai}))*L_{rodbio}
d/dt(B_{roodoio}) = γ*L_{roodoio} - (μ+σ)*B_{roodoio} - c_{Brodo}*B_{roodoio} - τ*B_{roodoio} + (1/X_{ao})*B_{roodaio} - τ₁*B_{roodoio} - τ₂*B_{roodoio} - τ_{ab}*B_{roodoio} + (1/(X_b-X_{ai}))*B_{rodbio}
d/dt(I_{roodoio}) = σ*B_{roodoio} - (μ+δ)*I_{roodoio} - c_{Irodo}*I_{roodoio} - τ*I_{roodoio} + (1/X_{ao})*I_{roodaio} - τ₁*I_{roodoio} - τ₂*I_{roodoio} - τ_{ab}*I_{roodoio} + (1/(X_b-X_{ai}))*I_{rodbio}

;radooio

d/dt(L_{radooio}) = β*S_{doio}*T_{Ira}/N₀ - (μ+γ)*L_{radooio} + (1/X_{ao})*L_{radaio} - τ₁*L_{radooio} + (1/(X_b-X_{ai}))*L_{radbio}
d/dt(B_{radooio}) = γ*L_{radooio} - (μ+σ)*B_{radooio} - c_{Brado}*B_{radooio} - τ*B_{radooio} + (1/X_{ao})*B_{roadaio} - τ₁*B_{radooio} - τ₂*B_{radooio} - τ_{ab}*B_{radooio} + (1/(X_b-X_{ai}))*B_{radbio}
d/dt(I_{radooio}) = σ*B_{radooio} - (μ+δ)*I_{radooio} - c_{Irado}*I_{radooio} - τ*I_{radooio} + (1/X_{ao})*I_{roadaio} - τ₁*I_{radooio} - τ₂*I_{radooio} - τ_{ab}*I_{radooio} + (1/(X_b-X_{ai}))*I_{radbio} + SQUAREPULSE(st,1/365)*mst

;rbodoio

d/dt(L_{rbodoio}) = β*S_{doio}*T_{Ir_b}/N₀ - (μ+γ)*L_{rbodoio} + (1/X_{ao})*L_{rbdaio} - τ₁*L_{rbodoio} + (1/(X_b-X_{ai}))*L_{rbdbio}

$$\begin{aligned} d/dt(B_{rbdoi}) &= \gamma * L_{rbdoi} - (\mu + \sigma) * B_{rbdoi} - c_{Brbdo} * B_{rbdoi} - \tau * B_{rbdoi} + (1/X_{ao}) * B_{rbdaio} - \tau_1 * B_{rbdoi} - \\ &\quad \tau_2 * B_{rbdoi} - \tau_{ab} * B_{rbdoi} + (1/(X_b - X_{ai})) * B_{rbdbio} \\ d/dt(I_{rbdoi}) &= \sigma * B_{rbdoi} - (\mu + \delta) * I_{rbdoi} - c_{Irbdo} * I_{rbdoi} - \tau * I_{rbdoi} + (1/X_{ao}) * I_{rbdaio} - \tau_1 * I_{rbdoi} - \tau_2 * I_{rbdoi} \\ &\quad - \tau_{ab} * I_{rbdoi} + (1/(X_b - X_{ai})) * I_{rbdbio} \end{aligned}$$

;Box N_daio

$$\begin{aligned} \text{init } S_{daio} &= 0 \\ \text{;init } B_{daio} &= B_{daio0} \\ \text{;init } L_{daio} &= L_{daio0} \\ \text{;init } I_{daio} &= I_{daio0} \end{aligned}$$

$$\begin{aligned} L_{daio} &= L_{rodaio} + L_{radaio} + L_{rbdaio} \\ B_{daio} &= B_{rodaio} + B_{radaio} + B_{rbdaio} \\ I_{daio} &= I_{rodaio} + I_{radaio} + I_{rbdaio} \end{aligned}$$

$$\begin{aligned} \text{init } L_{rodaio} &= 0 \\ \text{init } L_{radaio} &= 0 \\ \text{init } L_{rbdaio} &= 0 \\ \text{init } B_{rodaio} &= 0 \\ \text{init } B_{radaio} &= 0 \\ \text{init } B_{rbdaio} &= 0 \\ \text{init } I_{rodaio} &= 0 \\ \text{init } I_{radaio} &= 0 \\ \text{init } I_{rbdaio} &= 0 \end{aligned}$$

$$\begin{aligned} d/dt(S_{daio}) &= -\mu * S_{daio} - \beta * S_{daio} * T_{Iro}/N_0 + \delta * I_{rodaio} + c_{Broda} * B_{rodaio} + c_{Iroda} * I_{rodaio} - \beta * S_{daio} * T_{Ira}/N_0 + \\ &\quad \delta * I_{radaio} + c_{Brada} * B_{radaio} + c_{Irada} * I_{radaio} - \beta * S_{daio} * T_{Irb}/N_0 + \delta * I_{rbdaio} + c_{Brbda} * B_{rbdaio} + c_{Irbd} * I_{rbdaio} - \\ &\quad (1/X_{ao}) * S_{daio} - \tau_1 * S_{daio} \end{aligned}$$

;rodaio

$$\begin{aligned} d/dt(L_{rodaio}) &= \beta * S_{daio} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodaio} - (1/X_{ao}) * L_{rodaio} - \tau_1 * L_{rodaio} \\ d/dt(B_{rodaio}) &= \gamma * L_{rodaio} - (\mu + \sigma) * B_{rodaio} - c_{Broda} * B_{rodaio} + \tau * B_{rodoi} - (1/X_{ao}) * B_{rodaio} - \tau_1 * B_{rodaio} - \\ &\quad \tau_2 * B_{rodaio} \\ d/dt(I_{rodaio}) &= \sigma * B_{rodaio} - (\mu + \delta) * I_{rodaio} - c_{Iroda} * I_{rodaio} + \tau * I_{rodoi} - (1/X_{ao}) * I_{rodaio} - \tau_1 * I_{rodaio} - \tau_2 * I_{rodaio} \end{aligned}$$

;radaio

$$\begin{aligned} d/dt(L_{radaio}) &= \beta * S_{daio} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radaio} - (1/X_{ao}) * L_{radaio} - \tau_1 * L_{radaio} \\ d/dt(B_{radaio}) &= \gamma * L_{radaio} - (\mu + \sigma) * B_{radaio} - c_{Brada} * B_{radaio} + \tau * B_{radoio} - (1/X_{ao}) * B_{radaio} - \tau_1 * B_{radaio} - \\ &\quad \tau_2 * B_{radaio} \\ d/dt(I_{radaio}) &= \sigma * B_{radaio} - (\mu + \delta) * I_{radaio} - c_{Irada} * I_{radaio} + \tau * I_{radoio} - (1/X_{ao}) * I_{radaio} - \tau_1 * I_{radaio} - \tau_2 * I_{radaio} \end{aligned}$$

;rbdaio

$$\begin{aligned}
d/dt(L_{rbdaio}) &= \beta * S_{daio} * T_{Irb} / N_0 - (\mu + \gamma) * L_{rbdaio} - (1/X_{ao}) * L_{rbdaio} - \tau_1 * L_{rbdaio} \\
d/dt(B_{rbdaio}) &= \gamma * L_{rbdaio} - (\mu + \sigma) * B_{rbdaio} - c_{Brbda} * B_{rbdaio} + \tau * B_{rbdoio} - (1/X_{ao}) * B_{rbdaio} - \tau_1 * B_{rbdaio} - \tau_2 * B_{rbdaio} \\
d/dt(I_{rbdaio}) &= \sigma * B_{rbdaio} - (\mu + \delta) * I_{rbdaio} - c_{Irbda} * I_{rbdaio} + \tau * I_{rbdoio} - (1/X_{ao}) * I_{rbdaio} - \tau_1 * I_{rbdaio} - \tau_2 * I_{rbdaio}
\end{aligned}$$

$$\begin{aligned}
N_{doio} &= S_{doio} + L_{rodoio} + B_{rodoio} + I_{rodoio} + L_{radoio} + B_{radoio} + I_{radoio} + L_{rbdoio} + B_{rbdoio} + I_{rbdoio} \\
N_{daio} &= S_{daio} + L_{rodaio} + B_{rodaio} + I_{rodaio} + L_{radaio} + B_{radaio} + I_{radaio} + L_{rbdaio} + B_{rbdaio} + I_{rbdaio}
\end{aligned}$$

;Intervention 1
;MDA using artesunate/piperaquine +/- primaquine
;N_dabi1, N_dbil, N_doi1

;Box N_dabi1

```

init S_dabi1 = 0
;init B_dabi1 = B_dabi10
;init L_dabi1 = L_dabi10
;init I_dabi1 = I_dabi10

```

$$\begin{aligned}
L_{dabi1} &= L_{rodabi1} + L_{radabi1} + L_{rbdabi1} \\
B_{dabi1} &= B_{rodabi1} + B_{radabi1} + B_{rbdabi1} \\
I_{dabi1} &= I_{rodabi1} + I_{radabi1} + I_{rbdabi1}
\end{aligned}$$

$$\begin{aligned}
init L_{rodabi1} &= 0 \\
init L_{radabi1} &= 0 \\
init L_{rbdabi1} &= 0 \\
init B_{rodabi1} &= 0 \\
init B_{radabi1} &= 0 \\
init B_{rbdabi1} &= 0 \\
init I_{rodabi1} &= 0 \\
init I_{radabi1} &= 0 \\
init I_{rbdabi1} &= 0
\end{aligned}$$

$$\begin{aligned}
d/dt(S_{dabi1}) &= -\mu * S_{dabi1} - \beta * S_{dabi1} * T_{Iro} / N_0 + \delta * I_{rodabi1} + c_{Brodab} * B_{rodabi1} + c_{Irodab} * I_{rodabi1} - \\
&\beta * S_{dabi1} * T_{Ira} / N_0 + \delta * I_{radabi1} + c_{Bradab} * B_{radabi1} + c_{Iradab} * I_{radabi1} - \beta * S_{dabi1} * T_{Irb} / N_0 + \delta * I_{rbdabi1} + \\
&c_{Brbdab} * B_{rbdabi1} + c_{Irbdab} * I_{rbdabi1} + \tau_1 * S_{doio} + \tau_1 * S_{daio} - (1/X_{ai}) * S_{dabi1} + c_{Lp} * L_{rodabi1} + c_{Ip} * I_{rodabi1} + \\
&c_{Lp} * L_{radabi1} + c_{Ip} * I_{radabi1} + c_{Lp} * L_{rbdabi1} + c_{Ip} * I_{rbdabi1}
\end{aligned}$$

;rodabil

$d/dt(L_{rodabi1}) = \beta * S_{dabi1} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodabi1} + \tau_1 * L_{rodoio} + \tau_1 * L_{rodaio} - (1/X_{ai}) * L_{rodabi1} - c_{Lp} * L_{rodabi1}$
 $d/dt(B_{rodabi1}) = \gamma * L_{rodabi1} - (\mu + \sigma) * B_{rodabi1} - c_{Brodab} * B_{rodabi1} + \tau_1 * B_{rodoio} + \tau_1 * B_{rodaio} - (1/X_{ai}) * B_{rodabi1} + \tau_{abi1} * B_{rodoi1} + \tau_1 * B_{rodoi1}$
 $d/dt(I_{rodabi1}) = \sigma * B_{rodabi1} - (\mu + \delta) * I_{rodabi1} - c_{Irodab} * I_{rodabi1} + \tau_1 * I_{rodoio} + \tau_1 * I_{rodaio} - (1/X_{ai}) * I_{rodabi1} + \tau_{abi1} * I_{rodoi1} - c_{Ip} * I_{rodabi1} + \tau_1 * I_{rodoi1}$

;radabi1

$d/dt(L_{radabi1}) = \beta * S_{dabi1} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi1} + \tau_1 * L_{radoio} + \tau_1 * L_{radaio} - (1/X_{ai}) * L_{radabi1} - c_{Lp} * L_{radabi1}$
 $d/dt(B_{radabi1}) = \gamma * L_{radabi1} - (\mu + \sigma) * B_{radabi1} - c_{Bradab} * B_{radabi1} + \tau_1 * B_{radoio} + \tau_1 * B_{radaio} - (1/X_{ai}) * B_{radabi1} + \tau_{abi1} * B_{radoi1} + \tau_1 * B_{radoi1}$
 $d/dt(I_{radabi1}) = \sigma * B_{radabi1} - (\mu + \delta) * I_{radabi1} - c_{Iradab} * I_{radabi1} + \tau_1 * I_{radoio} + \tau_1 * I_{radaio} - (1/X_{ai}) * I_{radabi1} + \tau_{abi1} * I_{radoi1} - c_{Ip} * I_{radabi1} + \tau_1 * I_{radoi1}$

;rbdbabi1

$d/dt(L_{rbdbabi1}) = \beta * S_{dabi1} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdbabi1} + \tau_1 * L_{rbdoi} + \tau_1 * L_{rbdaio} - (1/X_{ai}) * L_{rbdbabi1} - c_{Lp} * L_{rbdbabi1}$
 $d/dt(B_{rbdbabi1}) = \gamma * L_{rbdbabi1} - (\mu + \sigma) * B_{rbdbabi1} - c_{Brbdab} * B_{rbdbabi1} + \tau_1 * B_{rbdoi} + \tau_1 * B_{rbdaio} - (1/X_{ai}) * B_{rbdbabi1} + \tau_{abi1} * B_{rbdoi1} + \tau_{abi1} * B_{rbdbi1} + \tau_1 * B_{rbdoi1}$
 $d/dt(I_{rbdbabi1}) = \sigma * B_{rbdbabi1} - (\mu + \delta) * I_{rbdbabi1} - c_{Irbdbab} * I_{rbdbabi1} + \tau_1 * I_{rbdoi} + \tau_1 * I_{rbdaio} - (1/X_{ai}) * I_{rbdbabi1} + \tau_{abi1} * I_{rbdoi1} + \tau_{abi1} * I_{rbdbi1} - c_{Ip} * I_{rbdbabi1} + \tau_1 * I_{rbdoi1}$

$$N_{dabi1} = S_{dabi1} + L_{rodabi1} + B_{rodabi1} + I_{rodabi1} + L_{radabi1} + B_{radabi1} + I_{radabi1} + L_{rbdbabi1} + B_{rbdbabi1} + I_{rbdbabi1}$$

;Box N_db1

init S_db1 = 0
 ;init B_db1 = B_db10
 ;init L_db1 = L_db10
 ;init I_db1 = I_db10

$L_{db1} = L_{rodbi1} + L_{radbi1} + L_{rbdbi1}$
 $B_{db1} = B_{rodbi1} + B_{radbi1} + B_{rbdbi1}$
 $I_{db1} = I_{rodbi1} + I_{radbi1} + I_{rbdbi1}$

init L_rodbi1 = 0
 init L_radbi1 = 0
 init L_rbdbi1 = 0
 init B_rodbi1 = 0
 init B_radbi1 = 0

```

init B_rdbdbi1 = 0
init I_rodbi1 = 0
init I_radbi1 = 0
init I_rdbdbi1 = 0

```

$$\frac{d}{dt}(S_{dbi1}) = -\mu * S_{dbi1} - \beta * S_{dbi1} * T_{Iro}/N_0 + \delta * I_{rodbi1} + c_{Brodb} * B_{rodbi1} + c_{Irod} * I_{rodbi1} - \beta * S_{dbi1} * T_{Ira}/N_0 + \delta * I_{radbi1} + c_{Brad} * B_{radbi1} + c_{Irad} * I_{radbi1} - \beta * S_{dbi1} * T_{Irb}/N_0 + \delta * I_{rdbdbi1} + c_{Brdb} * B_{rdbdbi1} + c_{Irdb} * I_{rdbdbi1} + (1/X_{ai}) * S_{dab1} - (1/(X_b - X_{ai})) * S_{dbi1}$$

;rodbi1

$$\begin{aligned}\frac{d}{dt}(L_{rodbi1}) &= \beta * S_{dbi1} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodabi1} + (1/X_{ai}) * L_{rodabi1} - (1/(X_b - X_{ai})) * L_{rodabi1} \\ \frac{d}{dt}(B_{rodbi1}) &= \gamma * L_{rodbi1} - (\mu + \sigma) * B_{rodbi1} - c_{Brodb} * B_{rodbi1} + (1/X_{ai}) * B_{rodabi1} - (1/(X_b - X_{ai})) * B_{rodabi1} \\ \frac{d}{dt}(I_{rodbi1}) &= \sigma * B_{rodbi1} - (\mu + \delta) * I_{rodbi1} - c_{Irod} * I_{rodbi1} + (1/X_{ai}) * I_{rodabi1} - (1/(X_b - X_{ai})) * I_{rodabi1}\end{aligned}$$

;radbi1

$$\begin{aligned}\frac{d}{dt}(L_{radbi1}) &= \beta * S_{dbi1} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi1} + (1/X_{ai}) * L_{radabi1} - (1/(X_b - X_{ai})) * L_{radabi1} \\ \frac{d}{dt}(B_{radbi1}) &= \gamma * L_{radbi1} - (\mu + \sigma) * B_{radbi1} - c_{Brad} * B_{radbi1} + (1/X_{ai}) * B_{radabi1} - (1/(X_b - X_{ai})) * B_{radabi1} \\ \frac{d}{dt}(I_{radbi1}) &= \sigma * B_{radbi1} - (\mu + \delta) * I_{radbi1} - c_{Irad} * I_{radbi1} + (1/X_{ai}) * I_{radabi1} - (1/(X_b - X_{ai})) * I_{radabi1}\end{aligned}$$

;rdbdbi1

$$\begin{aligned}\frac{d}{dt}(L_{rdbdbi1}) &= \beta * S_{dbi1} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rdbdbi1} + (1/X_{ai}) * L_{rdbabi1} - (1/(X_b - X_{ai})) * L_{rdbabi1} \\ \frac{d}{dt}(B_{rdbdbi1}) &= \gamma * L_{rdbdbi1} - (\mu + \sigma) * B_{rdbdbi1} - c_{Brdb} * B_{rdbdbi1} + (1/X_{ai}) * B_{rdbabi1} - (1/(X_b - X_{ai})) * B_{rdbabi1} - \tau_{abi1} * B_{rdbdbi1} - \tau_{ai1} * B_{rdbdbi1} \\ \frac{d}{dt}(I_{rdbdbi1}) &= \sigma * B_{rdbdbi1} - (\mu + \delta) * I_{rdbdbi1} - c_{Irdb} * I_{rdbdbi1} + (1/X_{ai}) * I_{rdbabi1} - (1/(X_b - X_{ai})) * I_{rdbabi1} - \tau_{abi1} * I_{rdbdbi1} - \tau_{ai1} * I_{rdbdbi1}\end{aligned}$$

$$N_{dbi1} = S_{dbi1} + L_{rodbi1} + B_{rodbi1} + I_{rodbi1} + L_{radbi1} + B_{radbi1} + I_{radbi1} + L_{rdbdbi1} + B_{rdbdbi1} + I_{rdbdbi1}$$

;Box N_doi1

```

init S_doi1 = 0;N_0-I_doi10
;init B_doi1 = B_doi10
;init L_doi1 = L_doi10
I_doi10 = 0;N_0*pinf

```

$$\begin{aligned}L_{doi1} &= L_{rodoi1} + L_{radoi1} + L_{rbdoi1} \\ B_{doi1} &= B_{rodoi1} + B_{radoi1} + B_{rbdoi1} \\ I_{doi1} &= I_{rodoi1} + I_{radoi1} + I_{rbdoi1}\end{aligned}$$

init L_rodoi1 = 0

```

init L_radoi1 = 0
init L_rbdoi1 = 0
init B_rodoi1 = 0
init B_radoi1 = 0
init B_rbdoi1 = 0
init I_rodoi1 = 0; p_o * I_doi1 0
init I_radoi1 = 0; p_a * I_doi1 0
init I_rbdoi1 = 0; p_b * I_doi1 0

```

$$d/dt(S_{doi1}) = -\mu * S_{doi1} - \beta * S_{doi1} * T_{Iro}/N_0 + \delta * I_{rodoi1} + c_{Brodo} * B_{rodoi1} + c_{Irodo} * I_{rodoi1} - \beta * S_{doi1} * T_{Ira}/N_0 + \delta * I_{radoi1} + c_{Brado} * B_{radoi1} + c_{Irado} * I_{radoi1} - \beta * S_{doi1} * T_{Irb}/N_0 + \delta * I_{rbdoi1} + c_{Brbdo} * B_{rbdoi1} + c_{Irbdo} * I_{rbdoi1} + (1/(X_b - X_{ai})) * S_{dbi1} + 1/(X_b - X_{ai} - X_{ao}) * S_{dbi1a} + (1/X_{ao}) * S_{dai1a}$$

;rodoi1

$$\begin{aligned} d/dt(L_{rodoi1}) &= \beta * S_{doi1} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodoi1} + (1/(X_b - X_{ai})) * L_{rodbi1} + 1/(X_b - X_{ai} - X_{ao}) * L_{rodbi1a} + (1/X_{ao}) * L_{radai1a} \\ d/dt(B_{rodoi1}) &= \gamma * L_{rodoi1} - (\mu + \sigma) * B_{rodoi1} - c_{Brodo} * B_{rodoi1} + (1/(X_b - X_{ai})) * B_{rodbi1} - \tau_{ab1} * B_{rodoi1} + 1/(X_b - X_{ai} - X_{ao}) * B_{rodbi1a} - \tau_{ai1} * B_{rodoi1} + (1/X_{ao}) * B_{radai1a} - \tau_1 * B_{rodoi1} \\ d/dt(I_{rodoi1}) &= \sigma * B_{rodoi1} - (\mu + \delta) * I_{rodoi1} - c_{Irodo} * I_{rodoi1} + (1/(X_b - X_{ai})) * I_{rodbi1} - \tau_{ab1} * I_{rodoi1} + 1/(X_b - X_{ai} - X_{ao}) * I_{rodbi1a} - \tau_{ai1} * I_{rodoi1} + (1/X_{ao}) * I_{radai1a} - \tau_1 * I_{rodoi1} \end{aligned}$$

;radoi1

$$\begin{aligned} d/dt(L_{radoi1}) &= \beta * S_{doi1} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radoi1} + (1/(X_b - X_{ai})) * L_{radbi1} + 1/(X_b - X_{ai} - X_{ao}) * L_{radbi1a} + (1/X_{ao}) * L_{radai1a} \\ d/dt(B_{radoi1}) &= \gamma * L_{radoi1} - (\mu + \sigma) * B_{radoi1} - c_{Brado} * B_{radoi1} + (1/(X_b - X_{ai})) * B_{radbi1} - \tau_{ab1} * B_{radoi1} + 1/(X_b - X_{ai} - X_{ao}) * B_{radbi1a} - \tau_{ai1} * B_{radoi1} + (1/X_{ao}) * B_{radai1a} - \tau_1 * B_{radoi1} \\ d/dt(I_{radoi1}) &= \sigma * B_{radoi1} - (\mu + \delta) * I_{radoi1} - c_{Irado} * I_{radoi1} + (1/(X_b - X_{ai})) * I_{radbi1} - \tau_{ab1} * I_{radoi1} + 1/(X_b - X_{ai} - X_{ao}) * I_{radbi1a} - \tau_{ai1} * I_{radoi1} + (1/X_{ao}) * I_{radai1a} - \tau_1 * I_{radoi1} \end{aligned}$$

;rbdoi1

$$\begin{aligned} d/dt(L_{rbdoi1}) &= \beta * S_{doi1} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdoi1} + (1/(X_b - X_{ai})) * L_{rbdbi1} + 1/(X_b - X_{ai} - X_{ao}) * L_{rbdbi1a} + (1/X_{ao}) * L_{rbdai1a} \\ d/dt(B_{rbdoi1}) &= \gamma * L_{rbdoi1} - (\mu + \sigma) * B_{rbdoi1} - c_{Brbdo} * B_{rbdoi1} + (1/(X_b - X_{ai})) * B_{rbdbi1} - \tau_{ab1} * B_{rbdoi1} + 1/(X_b - X_{ai} - X_{ao}) * B_{rbdbi1a} - \tau_{ai1} * B_{rbdoi1} + (1/X_{ao}) * B_{rbdai1a} - \tau_1 * B_{rbdoi1} \\ d/dt(I_{rbdoi1}) &= \sigma * B_{rbdoi1} - (\mu + \delta) * I_{rbdoi1} - c_{Irbdo} * I_{rbdoi1} + (1/(X_b - X_{ai})) * I_{rbdbi1} - \tau_{ab1} * I_{rbdoi1} + 1/(X_b - X_{ai} - X_{ao}) * I_{rbdbi1a} - \tau_{ai1} * I_{rbdoi1} + (1/X_{ao}) * I_{rbdai1a} - \tau_1 * I_{rbdoi1} \end{aligned}$$

$$N_{doi1} = S_{doi1} + L_{rodoi1} + B_{rodoi1} + I_{rodoi1} + L_{radoi1} + B_{radoi1} + I_{radoi1} + L_{rbdoi1} + B_{rbdoi1} + I_{rbdoi1}$$

;Treatment with artesunate in Intervention 1
; N_{dai1a}

;Box N_{dai1a}

```
init Sdai1a = 0
;init Bdai1a = Bdai1a0
;init Ldai1a = Ldai1a0
;init Idai1a = Idai1a0
```

$$\begin{aligned}L_{dai1a} &= L_{rodai1a} + L_{radai1a} + L_{rbdai1a} \\B_{dai1a} &= B_{rodai1a} + B_{radai1a} + B_{rbdai1a} \\I_{dai1a} &= I_{rodai1a} + I_{radai1a} + I_{rbdai1a}\end{aligned}$$

```
init Lrodai1a = 0
init Lradai1a = 0
init Lrbdai1a = 0
init Brodai1a = 0
init Bradai1a = 0
init Brbdai1a = 0
init Irodai1a = 0
init Iradai1a = 0
init Irbdai1a = 0
```

$$\begin{aligned}d/dt(S_{dai1a}) &= -\mu * S_{dai1a} - \beta * S_{dai1a} * T_{Iro}/N_0 + \delta * I_{rodai1a} + c_{Broda} * B_{rodai1a} + c_{Iroda} * I_{rodai1a} - \\&\quad \beta * S_{dai1a} * T_{Ira}/N_0 + \delta * I_{radai1a} + c_{Brada} * B_{radai1a} + c_{Irada} * I_{radai1a} - \beta * S_{dai1a} * T_{Irb}/N_0 + \delta * I_{rbdai1a} + \\&\quad c_{Brbda} * B_{rbdai1a} + c_{Irbd} * I_{rbdai1a} - (1/X_{ao}) * S_{dai1a}\end{aligned}$$

;rodai1a

$$\begin{aligned}d/dt(L_{rodai1a}) &= \beta * S_{dai1a} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodai1a} - (1/X_{ao}) * L_{rodai1a} \\d/dt(B_{rodai1a}) &= \gamma * L_{rodai1a} - (\mu + \sigma) * B_{rodai1a} - c_{Broda} * B_{rodai1a} + \tau_{ai1} * B_{rodoi1} - (1/X_{ao}) * B_{rodai1a} \\d/dt(I_{rodai1a}) &= \sigma * B_{rodai1a} - (\mu + \delta) * I_{rodai1a} - c_{Iroda} * I_{rodai1a} + \tau_{ai1} * I_{rodoi1} - (1/X_{ao}) * I_{rodai1a}\end{aligned}$$

;radai1a

$$\begin{aligned}d/dt(L_{radai1a}) &= \beta * S_{dai1a} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radai1a} - (1/X_{ao}) * L_{radai1a} \\d/dt(B_{radai1a}) &= \gamma * L_{radai1a} - (\mu + \sigma) * B_{radai1a} - c_{Brada} * B_{radai1a} + \tau_{ai1} * B_{rodoi1} - (1/X_{ao}) * B_{radai1a} \\d/dt(I_{radai1a}) &= \sigma * B_{radai1a} - (\mu + \delta) * I_{radai1a} - c_{Irada} * I_{radai1a} + \tau_{ai1} * I_{rodoi1} - (1/X_{ao}) * I_{radai1a}\end{aligned}$$

;rbdai1a

$$\begin{aligned}d/dt(L_{rbdai1a}) &= \beta * S_{dai1a} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdai1a} - (1/X_{ao}) * L_{rbdai1a} \\d/dt(B_{rbdai1a}) &= \gamma * L_{rbdai1a} - (\mu + \sigma) * B_{rbdai1a} - c_{Brbda} * B_{rbdai1a} + \tau_{ai1} * B_{rodoi1} - (1/X_{ao}) * B_{rbdai1a} \\d/dt(I_{rbdai1a}) &= \sigma * B_{rbdai1a} - (\mu + \delta) * I_{rbdai1a} - c_{Irbd} * I_{rbdai1a} + \tau_{ai1} * I_{rodoi1} - (1/X_{ao}) * I_{rbdai1a}\end{aligned}$$

$$N_{dai1a} = S_{dai1a} + L_{rodai1a} + B_{rodai1a} + I_{rodai1a} + L_{radai1a} + B_{radai1a} + I_{radai1a} + L_{rbdai1a} + B_{rbdai1a} + I_{rbdai1a}$$

;Treatment of infected patients with artesunate/piperaquine in Intervention 1
; N_{dabi1a}, N_{dbi1a}

;Box N_{dabi1a}

```
init Sdabi1a = 0
;init Bdabi1a = Bdabi1a0
;init Ldabi1a = Ldabi1a0
;init Idabi1a = Idabi1a0
```

$$\begin{aligned}L_{dabi1a} &= L_{rodabi1a} + L_{radabi1a} + L_{rbdabi1a} \\B_{dabi1a} &= B_{rodabi1a} + B_{radabi1a} + B_{rbdabi1a} \\I_{dabi1a} &= I_{rodabi1a} + I_{radabi1a} + I_{rbdabi1a}\end{aligned}$$

```
init Lrodabi1a = 0
init Lradabi1a = 0
init Lrbdabi1a = 0
init Brodabi1a = 0
init Bradabi1a = 0
init Brbdabi1a = 0
init Irodabi1a = 0
init Iradabi1a = 0
init Irbdabi1a = 0
```

$$\begin{aligned}d/dt(S_{dabi1a}) &= -\mu * S_{dabi1a} - \beta * S_{dabi1a} * T_{Iro}/N_0 + \delta * I_{rodabi1a} + c_{Brodab} * B_{rodabi1a} + c_{Irodab} * I_{rodabi1a} - \\&\beta * S_{dabi1a} * T_{Ira}/N_0 + \delta * I_{radabi1a} + c_{Bradb} * B_{radabi1a} + c_{Iradab} * I_{radabi1a} - \beta * S_{dabi1a} * T_{Irb}/N_0 + \delta * I_{rbdabi1a} + \\&c_{Brbdab} * B_{rbdabi1a} + c_{Irbdbab} * I_{rbdabi1a} - (1/X_{ao}) * S_{dabi1a}\end{aligned}$$

;rodabi1a

$$\begin{aligned}d/dt(L_{rodabi1a}) &= \beta * S_{dabi1a} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodabi1a} - (1/X_{ao}) * L_{rodabi1a} \\d/dt(B_{rodabi1a}) &= \gamma * L_{rodabi1a} - (\mu + \sigma) * B_{rodabi1a} - c_{Brodab} * B_{rodabi1a} - (1/X_{ao}) * B_{rodabi1a} \\d/dt(I_{rodabi1a}) &= \sigma * B_{rodabi1a} - (\mu + \delta) * I_{rodabi1a} - c_{Irodab} * I_{rodabi1a} - (1/X_{ao}) * I_{rodabi1a}\end{aligned}$$

;radabi1a

$$\begin{aligned}d/dt(L_{radabi1a}) &= \beta * S_{dabi1a} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi1a} - (1/X_{ao}) * L_{radabi1a} \\d/dt(B_{radabi1a}) &= \gamma * L_{radabi1a} - (\mu + \sigma) * B_{radabi1a} - c_{Bradab} * B_{radabi1a} - (1/X_{ao}) * B_{radabi1a} \\d/dt(I_{radabi1a}) &= \sigma * B_{radabi1a} - (\mu + \delta) * I_{radabi1a} - c_{Iradab} * I_{radabi1a} - (1/X_{ao}) * I_{radabi1a}\end{aligned}$$

;rbdabi1a

$$d/dt(L_{rbdabi1a}) = \beta * S_{dabi1a} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdabi1a} - (1/X_{ao}) * L_{rbdabi1a}$$

$$\begin{aligned} d/dt(B_{rdbabi1a}) &= \gamma * L_{rdbabi1a} - (\mu + \sigma) * B_{rdbabi1a} - c_{Brdbab} * B_{rdbabi1a} + \tau_{ai1} * B_{rdbibi1a} - (1/X_{ao}) * B_{rdbabi1a} + \\ &\quad \tau_{ai1} * B_{rbdbil} \\ d/dt(I_{rdbabi1a}) &= \sigma * B_{rdbabi1a} - (\mu + \delta) * I_{rdbabi1a} - c_{Irdbab} * I_{rdbabi1a} + \tau_{ai1} * I_{rbdbil} - (1/X_{ao}) * I_{rdbabi1a} + \\ &\quad \tau_{ai1} * I_{rbdbil} \end{aligned}$$

$$N_{dabi1a} = S_{dabi1a} + L_{rodabi1a} + B_{rodabi1a} + I_{rodabi1a} + L_{radabi1a} + B_{radabi1a} + I_{radabi1a} + L_{rdbabi1a} + B_{rdbabi1a} + I_{rdbabi1a}$$

;Box N_{dbi1a}

```
init Sdbi1a = 0
;init Bdbi1a = Bdbi1a0
;init Ldbi1a = Ldbi1a0
;init Idbi1a = Idbi1a0
```

$$\begin{aligned} L_{dbi1a} &= L_{rodibi1a} + L_{radibi1a} + L_{rbdbil} \\ B_{dbi1a} &= B_{rodibi1a} + B_{radibi1a} + B_{rbdbil} \\ I_{dbi1a} &= I_{rodibi1a} + I_{radibi1a} + I_{rbdbil} \end{aligned}$$

```
init Lrodibi1a = 0
init Lradibi1a = 0
init Lrbdbil = 0
init Brodibi1a = 0
init Bradibi1a = 0
init Brbdbil = 0
init Irodibi1a = 0
init Iradibi1a = 0
init Irbdbil = 0
```

$$\begin{aligned} d/dt(S_{dbi1a}) &= -\mu * S_{dbi1a} - \beta * S_{dbi1a} * T_{Iro}/N_0 + \delta * I_{rodibi1a} + c_{Brodb} * B_{rodibi1a} + c_{Irodb} * I_{rodibi1a} - \\ &\quad \beta * S_{dbi1a} * T_{Ira}/N_0 + \delta * I_{radibi1a} + c_{Bradb} * B_{radibi1a} + c_{Iradb} * I_{radibi1a} - \beta * S_{dbi1a} * T_{Irb}/N_0 + \delta * I_{rbdbil} + \\ &\quad c_{Brdb} * B_{rbdbil} + c_{Irdb} * I_{rbdbil} + (1/X_{ao}) * S_{dabi1a} - 1/(X_b - X_{ai} - X_{ao}) * S_{dabi1a} \end{aligned}$$

;rodbi1a

$$\begin{aligned} d/dt(L_{rodbibi1a}) &= \beta * S_{dbi1a} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodbibi1a} + (1/X_{ao}) * L_{rodabi1a} - 1/(X_b - X_{ai} - X_{ao}) * L_{rodbibi1a} \\ d/dt(B_{rodbibi1a}) &= \gamma * L_{rodbibi1a} - (\mu + \sigma) * B_{rodbibi1a} - c_{Brodb} * B_{rodbibi1a} + (1/X_{ao}) * B_{rodabi1a} - 1/(X_b - X_{ai} - X_{ao}) * B_{rodbibi1a} \\ d/dt(I_{rodbibi1a}) &= \sigma * B_{rodbibi1a} - (\mu + \delta) * I_{rodbibi1a} - c_{Irodb} * I_{rodbibi1a} + (1/X_{ao}) * I_{rodabi1a} - 1/(X_b - X_{ai} - X_{ao}) * I_{rodbibi1a} \end{aligned}$$

;radbi1a

$$\begin{aligned} d/dt(L_{radabi1a}) &= \beta * S_{dbi1a} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi1a} + (1/X_{ao}) * L_{radabi1a} - 1/(X_b - X_{ai} - X_{ao}) * L_{radabi1a} \\ d/dt(B_{radabi1a}) &= \gamma * L_{radabi1a} - (\mu + \sigma) * B_{radabi1a} - c_{Bradb} * B_{radabi1a} + (1/X_{ao}) * B_{radabi1a} - 1/(X_b - X_{ai} - X_{ao}) * B_{radabi1a} \end{aligned}$$

$$d/dt(I_{radbi1a}) = \sigma * B_{radbi1a} - (\mu + \delta) * I_{radbi1a} - c_{Iradb} * I_{radbi1a} + (1/X_{ao}) * I_{radabi1a} - 1/(X_b - X_{ai} - X_{ao}) * I_{radbi1a}$$

;rbdbi1a

$$d/dt(L_{rbdbi1a}) = \beta * S_{dbi1a} * T_{lrb} / N_0 - (\mu + \gamma) * L_{rbdbi1a} + (1/X_{ao}) * L_{rbdabi1a} - 1/(X_b - X_{ai} - X_{ao}) * L_{rbdbi1a}$$

$$d/dt(B_{rbdbi1a}) = \gamma * L_{rbdbi1a} - (\mu + \sigma) * B_{rbdbi1a} - c_{Brbdb} * B_{rbdbi1a} - \tau_{ai1} * B_{rbdbi1a} + (1/X_{ao}) * B_{rbdabi1a} - 1/(X_b - X_{ai} - X_{ao}) * B_{rbdbi1a}$$

$$d/dt(I_{rbdbi1a}) = \sigma * B_{rbdbi1a} - (\mu + \delta) * I_{rbdbi1a} - c_{Irbdb} * I_{rbdbi1a} - \tau_{ai1} * I_{rbdbi1a} + (1/X_{ao}) * I_{rbdabi1a} - 1/(X_b - X_{ai} - X_{ao}) * I_{rbdbi1a}$$

$$N_{dbi1a} = S_{dbi1a} + L_{rodbi1a} + B_{rodbi1a} + I_{rodbi1a} + L_{radbi1a} + B_{radbi1a} + I_{radbi1a} + L_{rbdbi1a} + B_{rbdbi1a} + I_{rbdbi1a}$$

;Switch to ACT (artesunate + piperaquine) for treatment in place of artesunate monotherapy

;also Intervention 3: MSAT with ACT

;N_dabio, N_dbio

;Box N_dabio

init S_dabio = 0

;init B_dabio = B_dabio0

;init L_dabio = L_dabio0

;init I_dabio = I_dabio0

$$L_{dabio} = L_{rodabio} + L_{radabio} + L_{rbdabio}$$

$$B_{dabio} = B_{rodabio} + B_{radabio} + B_{rbdabio}$$

$$I_{dabio} = I_{rodabio} + I_{radabio} + I_{rbdabio}$$

init L_rodabio = 0

init L_radabio = 0

init L_rbdabio = 0

init B_rodabio = 0

init B_radabio = 0

init B_rbdabio = 0

init I_rodabio = 0

init I_radabio = 0

init I_rbdabio = 0

$$d/dt(S_{dabio}) = -\mu * S_{dabio} - \beta * S_{dabio} * T_{lro} / N_0 + \delta * I_{rodabio} + c_{Brodab} * B_{rodabio} + c_{Irodab} * I_{rodabio} - \beta * S_{dabio} * T_{lra} / N_0 + \delta * I_{radabio} + c_{Bradab} * B_{radabio} + c_{Iradab} * I_{radabio} - \beta * S_{dabio} * T_{lrb} / N_0 + \delta * I_{rbdabio} + c_{Brbdab} * B_{rbdabio} + c_{Irbdab} * I_{rbdabio} - (1/X_{ai}) * S_{dabio} + c_{Lp} * (L_{rodabio} + L_{radabio} + L_{rbdabio}) + c_{Ip} * (I_{rodabio} + I_{radabio} + I_{rbdabio})$$

;rodabio

$$\begin{aligned} \frac{d}{dt}(L_{\text{rodabio}}) &= \beta * S_{\text{dabio}} * T_{\text{Iro}} / N_0 - (\mu + \gamma) * L_{\text{rodabio}} - (1/X_{\text{ai}}) * L_{\text{rodabio}} - c_{Lp} * L_{\text{rodabio}} \\ \frac{d}{dt}(B_{\text{rodabio}}) &= \gamma * L_{\text{rodabio}} - (\mu + \sigma) * B_{\text{rodabio}} - c_{\text{Brodab}} * B_{\text{rodabio}} + \tau_{ab} * B_{\text{rodoio}} - (1/X_{\text{ai}}) * B_{\text{rodabio}} \\ \frac{d}{dt}(I_{\text{rodabio}}) &= \sigma * B_{\text{rodabio}} - (\mu + \delta) * I_{\text{rodabio}} - c_{\text{Irodab}} * I_{\text{rodabio}} + \tau_{ab} * I_{\text{rodoio}} - (1/X_{\text{ai}}) * I_{\text{rodabio}} - c_{Ip} * I_{\text{rodabio}} \end{aligned}$$

;radabio

$$\begin{aligned} \frac{d}{dt}(L_{\text{radabio}}) &= \beta * S_{\text{dabio}} * T_{\text{Ira}} / N_0 - (\mu + \gamma) * L_{\text{radabio}} - (1/X_{\text{ai}}) * L_{\text{radabio}} - c_{Lp} * L_{\text{radabio}} \\ \frac{d}{dt}(B_{\text{radabio}}) &= \gamma * L_{\text{radabio}} - (\mu + \sigma) * B_{\text{radabio}} - c_{\text{Bradab}} * B_{\text{radabio}} + \tau_{ab} * B_{\text{radoio}} - (1/X_{\text{ai}}) * B_{\text{radabio}} \\ \frac{d}{dt}(I_{\text{radabio}}) &= \sigma * B_{\text{radabio}} - (\mu + \delta) * I_{\text{radabio}} - c_{\text{Iradab}} * I_{\text{radabio}} + \tau_{ab} * I_{\text{radoio}} - (1/X_{\text{ai}}) * I_{\text{radabio}} - c_{Ip} * I_{\text{radabio}} \end{aligned}$$

;rbdbdio

$$\begin{aligned} \frac{d}{dt}(L_{\text{rbdbdio}}) &= \beta * S_{\text{dabio}} * T_{\text{Irb}} / N_0 - (\mu + \gamma) * L_{\text{rbdbdio}} - (1/X_{\text{ai}}) * L_{\text{rbdbdio}} - c_{Lp} * L_{\text{rbdbdio}} \\ \frac{d}{dt}(B_{\text{rbdbdio}}) &= \gamma * L_{\text{rbdbdio}} - (\mu + \sigma) * B_{\text{rbdbdio}} - c_{\text{Brbdb}} * B_{\text{rbdbdio}} + \tau_{ab} * B_{\text{rbdoio}} - (1/X_{\text{ai}}) * B_{\text{rbdbdio}} + \\ &\quad \tau_{ab} * B_{\text{rbdbio}} \\ \frac{d}{dt}(I_{\text{rbdbdio}}) &= \sigma * B_{\text{rbdbdio}} - (\mu + \delta) * I_{\text{rbdbdio}} - c_{\text{Irbdb}} * I_{\text{rbdbdio}} + \tau_{ab} * I_{\text{rbdoio}} - (1/X_{\text{ai}}) * I_{\text{rbdbdio}} + \tau_{ab} * I_{\text{rbdbio}} - \\ &\quad c_{Ip} * I_{\text{rbdbio}} \end{aligned}$$

$$N_{\text{dabio}} = S_{\text{dabio}} + L_{\text{rodabio}} + B_{\text{rodabio}} + I_{\text{rodabio}} + L_{\text{radabio}} + B_{\text{radabio}} + I_{\text{radabio}} + L_{\text{rbdbdio}} + B_{\text{rbdbdio}} + I_{\text{rbdbdio}}$$

;Box N_dbio

$$\begin{aligned} \text{init } S_{\text{dabio}} &= 0 \\ \text{init } B_{\text{dbio}} &= B_{\text{dbio}}0 \\ \text{init } L_{\text{dbio}} &= L_{\text{dbio}}0 \\ \text{init } I_{\text{dbio}} &= I_{\text{dbio}}0 \end{aligned}$$

$$\begin{aligned} L_{\text{dbio}} &= L_{\text{rodbio}} + L_{\text{radbio}} + L_{\text{rbdbio}} \\ B_{\text{dbio}} &= B_{\text{rodgio}} + B_{\text{radgio}} + B_{\text{rbdbio}} \\ I_{\text{dbio}} &= I_{\text{rodgio}} + I_{\text{radgio}} + I_{\text{rbdbio}} \end{aligned}$$

$$\begin{aligned} \text{init } L_{\text{rodbio}} &= 0 \\ \text{init } L_{\text{radbio}} &= 0 \\ \text{init } L_{\text{rbdbio}} &= 0 \\ \text{init } B_{\text{rodbio}} &= 0 \\ \text{init } B_{\text{radbio}} &= 0 \\ \text{init } B_{\text{rbdbio}} &= 0 \\ \text{init } I_{\text{rodbio}} &= 0 \\ \text{init } I_{\text{radbio}} &= 0 \\ \text{init } I_{\text{rbdbio}} &= 0 \end{aligned}$$

$$\begin{aligned} d/dt(S_{dbio}) = & -\mu * S_{dbio} - \beta * S_{dbio} * T_{Iro}/N_0 + \delta * I_{rodbio} + c_{Brodb} * B_{rodbio} + c_{Irodb} * I_{rodbio} - \beta * S_{dbio} * T_{Ira}/N_0 + \\ & \delta * I_{radbio} + c_{Bradb} * B_{radbio} + c_{Iradb} * I_{radbio} - \beta * S_{dbio} * T_{rbdb}/N_0 + \delta * I_{rbdbio} + c_{Brbdb} * B_{rbdbio} + c_{Irbdb} * I_{rbdbio} + \\ & (1/X_{ai}) * S_{dabio} - (1/(X_b - X_{ai})) * S_{dbio} \end{aligned}$$

;rodbio

$$d/dt(L_{rodbio}) = \beta * S_{dbio} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodbio} + (1/X_{ai}) * L_{rodbio} - (1/(X_b - X_{ai})) * L_{rodbio}$$

$$d/dt(B_{rodbio}) = \gamma * L_{rodbio} - (\mu + \sigma) * B_{rodbio} - c_{Brodb} * B_{rodbio} + (1/X_{ai}) * B_{rodbio} - (1/(X_b - X_{ai})) * B_{rodbio}$$

$$d/dt(I_{rodbio}) = \sigma * B_{rodbio} - (\mu + \delta) * I_{rodbio} - c_{Irodb} * I_{rodbio} + (1/X_{ai}) * I_{rodbio} - (1/(X_b - X_{ai})) * I_{rodbio}$$

;radbio

$$d/dt(L_{radbio}) = \beta * S_{dbio} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radbio} + (1/X_{ai}) * L_{radbio} - (1/(X_b - X_{ai})) * L_{radbio}$$

$$d/dt(B_{radbio}) = \gamma * L_{radbio} - (\mu + \sigma) * B_{radbio} - c_{Bradb} * B_{radbio} + (1/X_{ai}) * B_{radbio} - (1/(X_b - X_{ai})) * B_{radbio}$$

$$d/dt(I_{radbio}) = \sigma * B_{radbio} - (\mu + \delta) * I_{radbio} - c_{Iradb} * I_{radbio} + (1/X_{ai}) * I_{radbio} - (1/(X_b - X_{ai})) * I_{radbio}$$

;rbdbio

$$d/dt(L_{rbdbio}) = \beta * S_{dbio} * T_{rbdb}/N_0 - (\mu + \gamma) * L_{rbdbio} + (1/X_{ai}) * L_{rbdbio} - (1/(X_b - X_{ai})) * L_{rbdbio}$$

$$d/dt(B_{rbdbio}) = \gamma * L_{rbdbio} - (\mu + \sigma) * B_{rbdbio} - c_{Brbdb} * B_{rbdbio} + (1/X_{ai}) * B_{rbdbio} - (1/(X_b - X_{ai})) * B_{rbdbio} - \tau_{ab} * B_{rbdbio}$$

$$d/dt(I_{rbdbio}) = \sigma * B_{rbdbio} - (\mu + \delta) * I_{rbdbio} - c_{Irbdb} * I_{rbdbio} + (1/X_{ai}) * I_{rbdbio} - (1/(X_b - X_{ai})) * I_{rbdbio} - \tau_{ab} * I_{rbdbio}$$

$$N_{dbio} = S_{dbio} + L_{rodbio} + B_{rodbio} + I_{rodbio} + L_{radbio} + B_{radbio} + I_{radbio} + L_{rbdbio} + B_{rbdbio} + I_{rbdbio}$$

;Intervention 2

;MSAT using atovaquone/proguanil

; N_dvgpi2, N_dvgpi2, N_dvi2, N_doi2

;Box N_dvgpi2

init S_dvgpi2 = 0

;init B_dvgpi2 = B_dvgpi20

;init L_dvgpi2 = L_dvgpi20

;init I_dvgpi2 = I_dvgpi20

L_dvgpi2 = L_rodvgpi2 + L_radvgpi2 + L_rbdvgpi2

B_dvgpi2 = B_rodvgpi2 + B_radvgpi2 + B_rbdvgpi2

I_dvgpi2 = I_rodvgpi2 + I_radvgpi2 + I_rbdvgpi2

init L_rodvgpi2 = 0

init L_radvgpi2 = 0

init L_rbdvgpi2 = 0

init B_rodvgpi2 = 0

```

init Bradvgpi2 = 0
init Brbdvgpi2 = 0
init Irodvgpi2 = 0
init Iradvgpi2 = 0
init Irbdvgpi2 = 0

```

$$\begin{aligned}
d/dt(S_{dvgpi2}) = & -\mu * S_{dvgpi2} - \beta * S_{dvgpi2} * T_{Iro}/N_0 + \delta * I_{rodvgpi2} + c_{Ldvg} * L_{rodvgpi2} + c_{Bdvg} * B_{rodvgpi2} + \\
& c_{Idvg} * I_{rodvgpi2} - \beta * S_{dvgpi2} * T_{Ira}/N_0 + \delta * I_{radvgpi2} + c_{Ldvg} * L_{radvgpi2} + c_{Bdvg} * B_{radvgpi2} + c_{Idvg} * I_{radvgpi2} - \\
& \beta * S_{dvgpi2} * T_{Irb}/N_0 + \delta * I_{rbdvgpi2} + c_{Ldvg} * L_{rbdvgpi2} + c_{Bdvg} * B_{rbdvgpi2} + c_{Idvg} * I_{rbdvgpi2} - (1/X_p) * S_{dvgpi2} + \\
& c_{Lp} * (L_{rodvgpi2} + L_{radvgpi2} + L_{rbdvgpi2}) + c_{Ip} * (I_{rodvgpi2} + I_{radvgpi2} + I_{rbdvgpi2})
\end{aligned}$$

;rodvgpi2

$$\begin{aligned}
d/dt(L_{rodvgpi2}) = & \beta * S_{dvgpi2} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodvgpi2} - (1/X_p) * L_{rodvgpi2} - c_{Lp} * L_{rodvgpi2} - c_{Ldvg} * L_{rodvgpi2} \\
d/dt(B_{rodvgpi2}) = & \gamma * L_{rodvgpi2} - (\mu + \sigma) * B_{rodvgpi2} + \tau_2 * B_{rodoio} + \tau_2 * B_{rodaio} - (1/X_p) * B_{rodvgpi2} - \\
& c_{Bdvg} * B_{rodvgpi2} + \tau_2 * B_{rodoi2} + \tau_2 * B_{radabi2} + \tau_2 * B_{radbi2} \\
d/dt(I_{rodvgpi2}) = & \sigma * B_{rodvgpi2} - (\mu + \delta) * I_{rodvgpi2} + \tau_2 * I_{rodoio} + \tau_2 * I_{rodaio} - (1/X_p) * I_{rodvgpi2} - c_{Ip} * I_{rodvgpi2} - \\
& c_{Idvg} * I_{rodvgpi2} + \tau_2 * I_{rodoi2} + \tau_2 * I_{radabi2} + \tau_2 * I_{radbi2}
\end{aligned}$$

;radvgpi2

$$\begin{aligned}
d/dt(L_{radvgpi2}) = & \beta * S_{dvgpi2} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radvgpi2} - (1/X_p) * L_{radvgpi2} - c_{Lp} * L_{radvgpi2} - c_{Ldvg} * L_{radvgpi2} \\
d/dt(B_{radvgpi2}) = & \gamma * L_{radvgpi2} - (\mu + \sigma) * B_{radvgpi2} + \tau_2 * B_{rodoio} + \tau_2 * B_{rodaio} - (1/X_p) * B_{radvgpi2} - \\
& c_{Bdvg} * B_{radvgpi2} + \tau_2 * B_{rodoi2} + \tau_2 * B_{radabi2} + \tau_2 * B_{radbi2} \\
d/dt(I_{radvgpi2}) = & \sigma * B_{radvgpi2} - (\mu + \delta) * I_{radvgpi2} + \tau_2 * I_{rodoio} + \tau_2 * I_{rodaio} - (1/X_p) * I_{radvgpi2} - c_{Ip} * I_{radvgpi2} - \\
& c_{Idvg} * I_{radvgpi2} + \tau_2 * I_{rodoi2} + \tau_2 * I_{radabi2} + \tau_2 * I_{radbi2}
\end{aligned}$$

;rbdvgpi2

$$\begin{aligned}
d/dt(L_{rbdvgpi2}) = & \beta * S_{dvgpi2} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdvgpi2} - (1/X_p) * L_{rbdvgpi2} - c_{Lp} * L_{rbdvgpi2} - c_{Ldvg} * L_{rbdvgpi2} \\
d/dt(B_{rbdvgpi2}) = & \gamma * L_{rbdvgpi2} - (\mu + \sigma) * B_{rbdvgpi2} + \tau_2 * B_{rbdoi} + \tau_2 * B_{rbdaio} - (1/X_p) * B_{rbdvgpi2} - \\
& c_{Bdvg} * B_{rbdvgpi2} + \tau_2 * B_{rbdoi2} + \tau_2 * B_{rbdabi2} + \tau_2 * B_{rbdbi2} \\
d/dt(I_{rbdvgpi2}) = & \sigma * B_{rbdvgpi2} - (\mu + \delta) * I_{rbdvgpi2} + \tau_2 * I_{rbdoi} + \tau_2 * I_{rbdaio} - (1/X_p) * I_{rbdvgpi2} - c_{Ip} * I_{rbdvgpi2} - \\
& c_{Idvg} * I_{rbdvgpi2} + \tau_2 * I_{rbdoi2} + \tau_2 * I_{rbdabi2} + \tau_2 * I_{rbdbi2}
\end{aligned}$$

$$\begin{aligned}
N_{dvgpi2} = & S_{dvgpi2} + L_{rodvgpi2} + B_{rodvgpi2} + I_{rodvgpi2} + L_{radvgpi2} + B_{radvgpi2} + I_{radvgpi2} + L_{rbdvgpi2} + B_{rbdvgpi2} \\
& + I_{rbdvgpi2}
\end{aligned}$$

;Box N_{dvgi2}

```

init Sdvgi2 = 0
;init Bdvgi2 = Bdvgi20
;init Ldvgi2 = Ldvgi20

```

;init $I_{dvgi2} = I_{dvgi20}$

$$\begin{aligned} L_{dvgi2} &= L_{rodvgi2} + L_{radvgi2} + L_{rbdvgi2} \\ B_{dvgi2} &= B_{rodvgi2} + B_{radvgi2} + B_{rbdvgi2} \\ I_{dvgi2} &= I_{rodvgi2} + I_{radvgi2} + I_{rbdvgi2} \end{aligned}$$

init $L_{rodvgi2} = 0$
 init $L_{radvgi2} = 0$
 init $L_{rbdvgi2} = 0$
 init $B_{rodvgi2} = 0$
 init $B_{radvgi2} = 0$
 init $B_{rbdvgi2} = 0$
 init $I_{rodvgi2} = 0$
 init $I_{radvgi2} = 0$
 init $I_{rbdvgi2} = 0$

$$\begin{aligned} d/dt(S_{dvgi2}) &= -\mu * S_{dvgi2} - \beta * S_{dvgi2} * T_{Iro}/N_0 + \delta * I_{rodvgi2} - \beta * S_{dvgi2} * T_{Ira}/N_0 + \delta * I_{radvgi2} - \\ &\beta * S_{dvgi2} * T_{IrB}/N_0 + \delta * I_{rbdvgi2} + (1/X_p) * S_{dvgp2} - (1/(X_g-X_p)) * S_{dvgi2} + c_{Ldvg} * (L_{rodvgi2} + L_{radvgi2} + \\ &L_{rbdvgi2}) + c_{Bdvg} * (B_{rodvgi2} + B_{radvgi2} + B_{rbdvgi2}) + c_{Idvg} * (I_{rodvgi2} + I_{radvgi2} + I_{rbdvgi2}) \end{aligned}$$

;rodvgi2

$$\begin{aligned} d/dt(L_{rodvgi2}) &= \beta * S_{dvgi2} * T_{Iro}/N_0 - (\mu+\gamma) * L_{rodvgi2} + (1/X_p) * L_{rodvgp2} - (1/(X_g-X_p)) * L_{rodvgi2} - \\ &c_{Ldvg} * L_{rodvgi2} \\ d/dt(B_{rodvgi2}) &= \gamma * L_{rodvgi2} - (\mu+\sigma) * B_{rodvgi2} + (1/X_p) * B_{rodvgp2} - (1/(X_g-X_p)) * B_{rodvgi2} - c_{Bdvg} * B_{rodvgi2} \\ d/dt(I_{rodvgi2}) &= \sigma * B_{rodvgi2} - (\mu+\delta) * I_{rodvgi2} + (1/X_p) * I_{rodvgp2} - (1/(X_g-X_p)) * I_{rodvgi2} - c_{Idvg} * I_{rodvgi2} \end{aligned}$$

;radvgi2

$$\begin{aligned} d/dt(L_{radvgi2}) &= \beta * S_{dvgi2} * T_{Ira}/N_0 - (\mu+\gamma) * L_{radvgi2} + (1/X_p) * L_{radvgp2} - (1/(X_g-X_p)) * L_{radvgi2} - \\ &c_{Ldvg} * L_{radvgi2} \\ d/dt(B_{radvgi2}) &= \gamma * L_{radvgi2} - (\mu+\sigma) * B_{radvgi2} + (1/X_p) * B_{radvgp2} - (1/(X_g-X_p)) * B_{radvgi2} - c_{Bdvg} * B_{radvgi2} \\ d/dt(I_{radvgi2}) &= \sigma * B_{radvgi2} - (\mu+\delta) * I_{radvgi2} + (1/X_p) * I_{radvgp2} - (1/(X_g-X_p)) * I_{radvgi2} - c_{Idvg} * I_{radvgi2} \end{aligned}$$

;rbdvgi2

$$\begin{aligned} d/dt(L_{rbdvgi2}) &= \beta * S_{dvgi2} * T_{IrB}/N_0 - (\mu+\gamma) * L_{rbdvgi2} + (1/X_p) * L_{rbdvgp2} - (1/(X_g-X_p)) * L_{rbdvgi2} - \\ &c_{Ldvg} * L_{rbdvgi2} \\ d/dt(B_{rbdvgi2}) &= \gamma * L_{rbdvgi2} - (\mu+\sigma) * B_{rbdvgi2} + (1/X_p) * B_{rbdvgp2} - (1/(X_g-X_p)) * B_{rbdvgi2} - c_{Bdvg} * B_{rbdvgi2} \\ d/dt(I_{rbdvgi2}) &= \sigma * B_{rbdvgi2} - (\mu+\delta) * I_{rbdvgi2} + (1/X_p) * I_{rbdvgp2} - (1/(X_g-X_p)) * I_{rbdvgi2} - c_{Idvg} * I_{rbdvgi2} \end{aligned}$$

$$N_{dvgi2} = S_{dvgi2} + L_{rodvgi2} + B_{rodvgi2} + I_{rodvgi2} + L_{radvgi2} + B_{radvgi2} + I_{radvgi2} + L_{rbdvgi2} + B_{rbdvgi2} + I_{rbdvgi2}$$

```

;Box N_dvi2

init S_dvi2 = 0
;init B_dvi2 = B_dvi20
;init L_dvi2 = L_dvi20
;init I_dvi2 = I_dvi20

L_dvi2 = L_rodvi2+L_radvvi2+L_rbdvi2
B_dvi2 = B_rodvi2+B_radvvi2+B_rbdvi2
I_dvi2 = I_rodvi2+I_radvvi2+I_rbdvi2

init L_rodvi2 = 0
init L_radvvi2 = 0
init L_rbdvi2 = 0
init B_rodvi2 = 0
init B_radvvi2 = 0
init B_rbdvi2 = 0
init I_rodvi2 = 0
init I_radvvi2 = 0
init I_rbdvi2 = 0

d/dt(S_dvi2) = - μ*S_dvi2 - β*S_dvi2*T_Iro/N₀ + δ*I_rodvi2 - β*S_dvi2*T_Ira/N₀ + δ*I_radvvi2 - β*S_dvi2*T_Irb/N₀ +
δ*I_rbdvi2 + (1/(X_g-X_p))*S_dvgi2 - (1/(X_v-X_g-X_p))*S_dvi2) + c_Ldv*(L_rodvi2 + L_radvvi2 + L_rbdvi2) +
c_Bdv*(B_rodvi2 + B_radvvi2 + B_rbdvi2) + c_Idv*(I_rodvi2 + I_radvvi2 + I_rbdvi2)

;rodvi2
d/dt(L_rodvi2) = β*S_dvi2*T_Iro/N₀ - (μ+γ)*L_rodvi2 - c_Ldv*L_rodvi2 + (1/(X_g-X_p))*L_radvgi2 - (1/(X_v-X_g-
X_p))*L_rodvi2)
d/dt(B_rodvi2) = γ*L_rodvi2 - (μ+σ)*B_rodvi2 - c_Bdv*B_rodvi2 + (1/(X_g-X_p))*B_radvgi2 - (1/(X_v-X_g-X_p))*B_rodvi2)
d/dt(I_rodvi2) = σ*B_rodvi2 - (μ+δ)*I_rodvi2 - c_Idv*I_rodvi2 + (1/(X_g-X_p))*I_radvgi2 - (1/(X_v-X_g-X_p))*I_rodvi2)

;radvvi2
d/dt(L_radvvi2) = β*S_dvi2*T_Ira/N₀ - (μ+γ)*L_radvvi2 - c_Ldv*L_radvvi2 + (1/(X_g-X_p))*L_radvgi2 - (1/(X_v-X_g-
X_p))*L_radvvi2)
d/dt(B_radvvi2) = γ*L_radvvi2 - (μ+σ)*B_radvvi2 - c_Bdv*B_radvvi2 + (1/(X_g-X_p))*B_radvgi2 - (1/(X_v-X_g-X_p))*B_radvvi2)
d/dt(I_radvvi2) = σ*B_radvvi2 - (μ+δ)*I_radvvi2 - c_Idv*I_radvvi2 + (1/(X_g-X_p))*I_radvgi2 - (1/(X_v-X_g-X_p))*I_radvvi2)

;rbdvi2

```

$$\begin{aligned}
d/dt(L_{rbdoi2}) &= \beta * S_{dvi2} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdoi2} - c_{Ldv} * L_{rbdoi2} + (1/(X_g - X_p)) * L_{rbdvgi2} - (1/(X_v - X_g - X_p)) * L_{rbdoi2} \\
d/dt(B_{rbdoi2}) &= \gamma * L_{rbdoi2} - (\mu + \sigma) * B_{rbdoi2} - c_{Bdv} * B_{rbdoi2} + (1/(X_g - X_p)) * B_{rbdvgi2} - (1/(X_v - X_g - X_p)) * B_{rbdoi2} \\
d/dt(I_{rbdoi2}) &= \sigma * B_{rbdoi2} - (\mu + \delta) * I_{rbdoi2} - c_{Idv} * I_{rbdoi2} + (1/(X_g - X_p)) * I_{rbdvgi2} - (1/(X_v - X_g - X_p)) * I_{rbdoi2}
\end{aligned}$$

$$N_{dvi2} = S_{dvi2} + L_{rodoi2} + B_{radoi2} + I_{rodoi2} + L_{radvi2} + B_{radvi2} + I_{radvi2} + L_{rbdoi2} + B_{rbdoi2} + I_{rbdoi2}$$

;Box N_{doi2}

$$\begin{aligned}
\text{init } S_{doi2} &= 0 \\
I_{doi2}0 &= 0
\end{aligned}$$

$$\begin{aligned}
L_{doi2} &= L_{rodoi2} + L_{radoi2} + L_{rbdoi2} \\
B_{doi2} &= B_{rodoi2} + B_{radoi2} + B_{rbdoi2} \\
I_{doi2} &= I_{rodoi2} + I_{radoi2} + I_{rbdoi2}
\end{aligned}$$

$$\begin{aligned}
\text{init } L_{rodoi2} &= 0 \\
\text{init } L_{radoi2} &= 0 \\
\text{init } L_{rbdoi2} &= 0 \\
\text{init } B_{rodoi2} &= 0 \\
\text{init } B_{radoi2} &= 0 \\
\text{init } B_{rbdoi2} &= 0 \\
\text{init } I_{rodoi2} &= 0 \\
\text{init } I_{radoi2} &= 0 \\
\text{init } I_{rbdoi2} &= 0
\end{aligned}$$

$$\begin{aligned}
d/dt(S_{doi2}) &= -\mu * S_{doi2} - \beta * S_{doi2} * T_{Iro}/N_0 + \delta * I_{rodoi2} + c_{Brodo} * B_{rodoi2} + c_{Irodo} * I_{rodoi2} - \beta * S_{doi2} * T_{Ira}/N_0 + \\
&\delta * I_{radoi2} + c_{Brado} * B_{radoi2} + c_{Irado} * I_{radoi2} - \beta * S_{doi2} * T_{Irb}/N_0 + \delta * I_{rbdoi2} + c_{Brbdo} * B_{rbdoi2} + c_{Irbdo} * I_{rbdoi2} + \\
&(1/(X_v - X_g - X_p)) * S_{dvi2} + 1/(X_b - X_{ai}) * S_{dbi2} + (1/X_{ao}) * S_{dai2}
\end{aligned}$$

;rodoi2

$$\begin{aligned}
d/dt(L_{rodoi2}) &= \beta * S_{doi2} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodoi2} + (1/(X_v - X_g - X_p)) * L_{rodoi2} + 1/(X_b - X_{ai}) * L_{rodbi2} + \\
&(1/X_{ao}) * L_{rodai2} \\
d/dt(B_{rodoi2}) &= \gamma * L_{rodoi2} - (\mu + \sigma) * B_{rodoi2} - c_{Brodo} * B_{rodoi2} + (1/(X_v - X_g - X_p)) * B_{rodoi2} - \tau_{abi2} * B_{rodoi2} + \\
&1/(X_b - X_{ai}) * B_{rodbi2} - \tau_{ai2} * B_{rodoi2} + (1/X_{ao}) * B_{rodai2} - \tau_2 * B_{rodoi2} \\
d/dt(I_{rodoi2}) &= \sigma * B_{rodoi2} - (\mu + \delta) * I_{rodoi2} - c_{Irodo} * I_{rodoi2} + (1/(X_v - X_g - X_p)) * I_{rodoi2} - \tau_{abi2} * I_{rodoi2} + 1/(X_b - \\
&X_{ai}) * I_{rodbi2} - \tau_{ai2} * I_{rodoi2} + (1/X_{ao}) * I_{rodai2} - \tau_2 * I_{rodoi2}
\end{aligned}$$

;radoi2

$$\begin{aligned}
d/dt(L_{radoi2}) &= \beta * S_{doi2} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radoi2} + (1/(X_v - X_g - X_p)) * L_{radvi2} + 1/(X_b - X_{ai}) * L_{radbi2} + \\
&(1/X_{ao}) * L_{radai2} \\
d/dt(B_{radoi2}) &= \gamma * L_{radoi2} - (\mu + \sigma) * B_{radoi2} - c_{Brado} * B_{radoi2} + (1/(X_v - X_g - X_p)) * B_{radvi2} - \tau_{abi2} * B_{radoi2} + \\
&1/(X_b - X_{ai}) * B_{radbi2} - \tau_{ai2} * B_{radoi2} + (1/X_{ao}) * B_{radai2} - \tau_2 * B_{radoi2} \\
d/dt(I_{radoi2}) &= \sigma * B_{radoi2} - (\mu + \delta) * I_{radoi2} - c_{Irado} * I_{radoi2} + (1/(X_v - X_g - X_p)) * I_{radvi2} - \tau_{abi2} * I_{radoi2} + 1/(X_b - \\
&X_{ai}) * I_{radbi2} - \tau_{ai2} * I_{radoi2} + (1/X_{ao}) * I_{radai2} - \tau_2 * I_{radoi2}
\end{aligned}$$

;rbdoi2

$$\begin{aligned}
d/dt(L_{rbdoi2}) &= \beta * S_{doi2} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdoi2} + (1/(X_v - X_g - X_p)) * L_{rbdvi2} + 1/(X_b - X_{ai}) * L_{rbdbi2} + \\
&(1/X_{ao}) * L_{rbdai2} \\
d/dt(B_{rbdoi2}) &= \gamma * L_{rbdoi2} - (\mu + \sigma) * B_{rbdoi2} - c_{Brbdo} * B_{rbdoi2} + (1/(X_v - X_g - X_p)) * B_{rbdvi2} - \tau_{abi2} * B_{rbdoi2} + \\
&1/(X_b - X_{ai}) * B_{rbdbi2} - \tau_{ai2} * B_{rbdoi2} + (1/X_{ao}) * B_{rbdai2} - \tau_2 * B_{rbdoi2} \\
d/dt(I_{rbdoi2}) &= \sigma * B_{rbdoi2} - (\mu + \delta) * I_{rbdoi2} - c_{Irbd} * I_{rbdoi2} + (1/(X_v - X_g - X_p)) * I_{rbdvi2} - \tau_{abi2} * I_{rbdoi2} + 1/(X_b - \\
&X_{ai}) * I_{rbdbi2} - \tau_{ai2} * I_{rbdoi2} + (1/X_{ao}) * I_{rbdai2} - \tau_2 * I_{rbdoi2}
\end{aligned}$$

$$N_{doi2} = S_{doi2} + L_{radoi2} + B_{radoi2} + I_{radoi2} + L_{radai2} + B_{radai2} + I_{radai2} + L_{rbdoi2} + B_{rbdoi2} + I_{rbdoi2}$$

;Treatment with artesunate in intervention 2

; N_dai2

;Box N_dai2

$$\begin{aligned}
\text{init } S_{dai2} &= 0 \\
\text{init } B_{dai2} &= B_{dai2}0 \\
\text{init } L_{dai2} &= L_{dai2}0 \\
\text{init } I_{dai2} &= I_{dai2}0
\end{aligned}$$

$$\begin{aligned}
L_{dai2} &= L_{rodai2} + L_{radai2} + L_{rbdai2} \\
B_{dai2} &= B_{rodai2} + B_{radai2} + B_{rbdai2} \\
I_{dai2} &= I_{rodai2} + I_{radai2} + I_{rbdai2}
\end{aligned}$$

$$\begin{aligned}
\text{init } L_{rodai2} &= 0 \\
\text{init } L_{radai2} &= 0 \\
\text{init } L_{rbdai2} &= 0 \\
\text{init } B_{rodai2} &= 0 \\
\text{init } B_{radai2} &= 0 \\
\text{init } B_{rbdai2} &= 0 \\
\text{init } I_{rodai2} &= 0 \\
\text{init } I_{radai2} &= 0 \\
\text{init } I_{rbdai2} &= 0
\end{aligned}$$

$$\begin{aligned}
d/dt(S_{dai2}) &= -\mu * S_{dai2} - \beta * S_{dai2} * T_{Iro}/N_0 + \delta * I_{rodai2} + c_{Broda} * B_{rodai2} + c_{Iroda} * I_{rodai2} - \beta * S_{dai2} * T_{Ira}/N_0 + \\
&\delta * I_{radai2} + c_{Brada} * B_{radai2} + c_{Irada} * I_{radai2} - \beta * S_{dai2} * T_{Irb}/N_0 + \delta * I_{rbdai2} + c_{Brbda} * B_{rbdai2} + c_{Irbd} * I_{rbdai2} - \\
&(1/X_{ao}) * S_{dai2}
\end{aligned}$$

```

;rodai2
d/dt(Lrodai2) = β*Sdai2*TIro/N0 - (μ+γ)*Lrodai2 - (1/Xao)*Lrodai2
d/dt(Brodai2) = γ*Lrodai2 - (μ+σ)*Brodai2 - cBrada*Brodai2 + τai2*Brodoi2 - (1/Xao)*Brodai2
d/dt(Irodai2) = σ*Brodai2 - (μ+δ)*Irodai2 - cIroda*Irodai2 + τai2*Irodoi2 - (1/Xao)*Irodai2

```

```

;radai2
d/dt(Lradai2) = β*Sdai2*TIra/N0 - (μ+γ)*Lradai2 - (1/Xao)*Lradai2
d/dt(Bradai2) = γ*Lradai2 - (μ+σ)*Bradai2 - cBrada*Bradai2 + τai2*Brodoi2 - (1/Xao)*Bradai2
d/dt(Iradai2) = σ*Bradai2 - (μ+δ)*Iradai2 - cIrada*Iradai2 + τai2*Irodoi2 - (1/Xao)*Iradai2

```

```

;rbdai2
d/dt(Lrbdai2) = β*Sdai2*TIrb/N0 - (μ+γ)*Lrbdai2 - (1/Xao)*Lrbdai2
d/dt(Brbdai2) = γ*Lrbdai2 - (μ+σ)*Brbdai2 - cBrbda*Brbdai2 + τai2*Brbdoi2 - (1/Xao)*Brbdai2
d/dt(Irbdai2) = σ*Brbdai2 - (μ+δ)*Irbdai2 - cIrbda*Irbdai2 + τai2*Irbdoi2 - (1/Xao)*Irbdai2

```

$$N_{dai2} = S_{dai2} + L_{rodai2} + B_{rodai2} + I_{rodai2} + L_{radai2} + B_{radai2} + I_{radai2} + L_{rbdai2} + B_{rbdai2} + I_{rbdai2}$$

```

;Treatment with ACT (artesunate + piperaquine) during intervention 2
; Ndabi2, Ndbi2

```

;Box N_{dabi2}

```

init Sdabi2 = 0
;init Bdabi2 = Bdabi20
;init Ldabi2 = Ldabi20
;init Idabi2 = Idabi20

```

```

Ldabi2 = Lrodabi2+Lradabi2+Lrbdabi2
Bdabi2 = Brodabi2+Bradabi2+Brbdabi2
Idabi2 = Irodabi2+Iradabi2+Irbdabi2

```

```

init Lrodabi2 = 0
init Lradabi2 = 0
init Lrbdabi2 = 0
init Brodabi2 = 0
init Bradabi2 = 0
init Brbdabi2 = 0
init Irodabi2 = 0
init Iradabi2 = 0
init Irbdabi2 = 0

```

$d/dt(S_{dabi2}) = -\mu * S_{dabi2} - \beta * S_{dabi2} * T_{Iro}/N_0 + \delta * I_{rodabi2} + c_{Brodab} * B_{rodabi2} + c_{Irodab} * I_{rodabi2} - \beta * S_{dabi2} * T_{Ira}/N_0 + \delta * I_{radabi2} + c_{Bradab} * B_{radabi2} + c_{Iradab} * I_{radabi2} - \beta * S_{dabi2} * T_{Irb}/N_0 + \delta * I_{rbdbabi2} + c_{Brbdab} * B_{rbdbabi2} + c_{Irbdbab} * I_{rbdbabi2} - (1/X_{ai}) * S_{dabi2}$

;rodabi2

$d/dt(L_{rodabi2}) = \beta * S_{dabi2} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodabi2} - (1/X_{ai}) * L_{rodabi2}$

$d/dt(B_{rodabi2}) = \gamma * L_{rodabi2} - (\mu + \sigma) * B_{rodabi2} - c_{Brodab} * B_{rodabi2} - (1/X_{ai}) * B_{rodabi2} + \tau_{abi2} * B_{rodoi2} - \tau_2 * B_{rodabi2}$

$d/dt(I_{rodabi2}) = \sigma * B_{rodabi2} - (\mu + \delta) * I_{rodabi2} - c_{Irodab} * I_{rodabi2} - (1/X_{ai}) * I_{rodabi2} + \tau_{abi2} * I_{rodoi2} - \tau_2 * I_{rodabi2}$

;radabi2

$d/dt(L_{radabi2}) = \beta * S_{dabi2} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi2} - (1/X_{ai}) * L_{radabi2}$

$d/dt(B_{radabi2}) = \gamma * L_{radabi2} - (\mu + \sigma) * B_{radabi2} - c_{Bradab} * B_{radabi2} - (1/X_{ai}) * B_{radabi2} + \tau_{abi2} * B_{rodoi2} - \tau_2 * B_{radabi2}$

$d/dt(I_{radabi2}) = \sigma * B_{radabi2} - (\mu + \delta) * I_{radabi2} - c_{Iradab} * I_{radabi2} - (1/X_{ai}) * I_{radabi2} + \tau_{abi2} * I_{rodoi2} - \tau_2 * I_{radabi2}$

;rbdbabi2

$d/dt(L_{rbdbabi2}) = \beta * S_{dabi2} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rbdbabi2} - (1/X_{ai}) * L_{rbdbabi2}$

$d/dt(B_{rbdbabi2}) = \gamma * L_{rbdbabi2} - (\mu + \sigma) * B_{rbdbabi2} - c_{Brbdab} * B_{rbdbabi2} + \tau_{ab} * B_{rbdbabi2} - (1/X_{ai}) * B_{rbdbabi2} + \tau_{abi2} * B_{rbdoi2} - \tau_2 * B_{rbdbabi2}$

$d/dt(I_{rbdbabi2}) = \sigma * B_{rbdbabi2} - (\mu + \delta) * I_{rbdbabi2} - c_{Irbdbab} * I_{rbdbabi2} + \tau_{ab} * I_{rbdbabi2} - (1/X_{ai}) * I_{rbdbabi2} + \tau_{abi2} * I_{rbdoi2} - \tau_2 * I_{rbdbabi2}$

$N_{dabi2} = S_{dabi2} + L_{rodabi2} + B_{rodabi2} + I_{rodabi2} + L_{radabi2} + B_{radabi2} + I_{radabi2} + L_{rbdbabi2} + B_{rbdbabi2} + I_{rbdbabi2}$

;Box N_db2

init S_db2 = 0

;init B_db2 = B_db20

;init L_db2 = L_db20

;init I_db2 = I_db20

$L_{db2} = L_{rodabi2} + L_{radabi2} + L_{rbdbabi2}$

$B_{db2} = B_{rodabi2} + B_{radabi2} + B_{rbdbabi2}$

$I_{db2} = I_{rodabi2} + I_{radabi2} + I_{rbdbabi2}$

init L_rodabi2 = 0

init L_radabi2 = 0

init L_rbdbabi2 = 0

init B_rodabi2 = 0

init B_radabi2 = 0

```

init B_rdbdbi2 = 0
init I_rodibi2 = 0
init I_radabi2 = 0
init I_rdbdbi2 = 0

```

$$\frac{d}{dt}(S_{dbi2}) = -\mu * S_{dbi2} - \beta * S_{dbi2} * T_{Iro}/N_0 + \delta * I_{rodibi2} + c_{Brodb} * B_{rodibi2} + c_{Irodib} * I_{rodibi2} - \beta * S_{dbi2} * T_{Ira}/N_0 + \delta * I_{radabi2} + c_{Bradb} * B_{radabi2} + c_{Iradab} * I_{radabi2} - \beta * S_{dbi2} * T_{Irb}/N_0 + \delta * I_{rdbdbi2} + c_{Brdbdb} * B_{rdbdbi2} + c_{Irbdb} * I_{rdbdbi2} + (1/X_{ai} * S_{dabi2}) - 1/(X_b - X_{ai}) * S_{dbi2}$$

;rodibi2

$$\begin{aligned}\frac{d}{dt}(L_{rodibi2}) &= \beta * S_{dbi2} * T_{Iro}/N_0 - (\mu + \gamma) * L_{rodabi2} + (1/X_{ai}) * L_{rodabi2} - 1/(X_b - X_{ai}) * L_{rodibi2} \\ \frac{d}{dt}(B_{rodibi2}) &= \gamma * L_{rodibi2} - (\mu + \sigma) * B_{rodibi2} - c_{Brodb} * B_{rodibi2} + (1/X_{ai}) * B_{rodabi2} - 1/(X_b - X_{ai}) * B_{rodibi2} - \tau_2 * B_{rodibi2} \\ \frac{d}{dt}(I_{rodibi2}) &= \sigma * B_{rodibi2} - (\mu + \delta) * I_{rodibi2} - c_{Irodib} * I_{rodibi2} + (1/X_{ai}) * I_{rodabi2} - 1/(X_b - X_{ai}) * I_{rodibi2} - \tau_2 * I_{rodibi2}\end{aligned}$$

;radabi2

$$\begin{aligned}\frac{d}{dt}(L_{radabi2}) &= \beta * S_{dbi2} * T_{Ira}/N_0 - (\mu + \gamma) * L_{radabi2} + (1/X_{ai}) * L_{radabi2} - 1/(X_b - X_{ai}) * L_{radabi2} \\ \frac{d}{dt}(B_{radabi2}) &= \gamma * L_{radabi2} - (\mu + \sigma) * B_{radabi2} - c_{Bradb} * B_{radabi2} + (1/X_{ai}) * B_{radabi2} - 1/(X_b - X_{ai}) * B_{radabi2} - \tau_2 * B_{radabi2} \\ \frac{d}{dt}(I_{radabi2}) &= \sigma * B_{radabi2} - (\mu + \delta) * I_{radabi2} - c_{Iradab} * I_{radabi2} + (1/X_{ai}) * I_{radabi2} - 1/(X_b - X_{ai}) * I_{radabi2} - \tau_2 * I_{radabi2}\end{aligned}$$

;rdbdbi2

$$\begin{aligned}\frac{d}{dt}(L_{rdbdbi2}) &= \beta * S_{dbi2} * T_{Irb}/N_0 - (\mu + \gamma) * L_{rdbdbi2} + (1/X_{ai}) * L_{rdbdbi2} - 1/(X_b - X_{ai}) * L_{rdbdbi2} \\ \frac{d}{dt}(B_{rdbdbi2}) &= \gamma * L_{rdbdbi2} - (\mu + \sigma) * B_{rdbdbi2} - c_{Brdbdb} * B_{rdbdbi2} - \tau_{ab} * B_{rdbdbi2} + (1/X_{ai}) * B_{rdbdbi2} - 1/(X_b - X_{ai}) * B_{rdbdbi2} - \tau_2 * B_{rdbdbi2} \\ \frac{d}{dt}(I_{rdbdbi2}) &= \sigma * B_{rdbdbi2} - (\mu + \delta) * I_{rdbdbi2} - c_{Irbdb} * I_{rdbdbi2} - \tau_{ab} * I_{rdbdbi2} + (1/X_{ai}) * I_{rdbdbi2} - 1/(X_b - X_{ai}) * I_{rdbdbi2} - \tau_2 * I_{rdbdbi2}\end{aligned}$$

$$N_{dbi2} = S_{dbi2} + L_{rodibi2} + B_{rodibi2} + I_{rodibi2} + L_{radabi2} + B_{radabi2} + I_{radabi2} + L_{rdbdbi2} + B_{rdbdbi2} + I_{rdbdbi2}$$

;CHECKS AND TOTALS

$$\text{check} = N_0 - (N_{doio} + N_{daio} + N_{dabi1} + N_{dbi1} + N_{doi1} + N_{dabio} + N_{dbio} + N_{dai1a} + N_{dabi1a} + N_{dbi1a} + N_{dvgp1} + N_{dvgi2} + N_{dvi2} + N_{doi2} + N_{dai2} + N_{dabi2} + N_{dbi2})$$

$$\begin{aligned}T_N &= N_{doio} + N_{daio} + N_{dabi1} + N_{dbi1} + N_{doi1} + N_{dabio} + N_{dbio} + N_{dai1a} + N_{dabi1a} + N_{dbi1a} + N_{dvgp1} + N_{dvgi2} + N_{dvi2} + N_{doi2} + N_{dai2} + N_{dabi2} + N_{dbi2}\end{aligned}$$

$$N_{ra} = (I_{radoio} + I_{radaio} + I_{radabi1} + I_{radbi1} + I_{radoi1} + I_{radabio} + I_{radbio} + I_{radai1a} + I_{radabi1a} + I_{radbi1a}) + (B_{radoio} + B_{radaio} + B_{radabi1} + B_{radbi1} + B_{radoi1} + B_{radabio} + B_{radbio} + B_{radai1a} + B_{radabi1a} + B_{radbi1a}) + (I_{radvgpi2} + I_{radvgi2} + I_{radvi2} + I_{radoi2} + I_{radai2} + I_{radabi2} + I_{radbi2}) + (B_{radvgpi2} + B_{radvgi2} + B_{radvi2} + B_{radoi2} + B_{radai2} + B_{radabi2} + B_{radbi2})$$

$$N_{rb} = (I_{rbdoio} + I_{rbdaio} + I_{rbdabi1} + I_{rbdbi1} + I_{rbdoi1} + I_{rbdabio} + I_{rbdai1a} + I_{rbdabi1a} + I_{rbdbi1a}) + (B_{rbdai1a} + B_{rbdoio} + B_{rbdaio} + B_{rbdabi1} + B_{rbdbi1} + B_{rbdoi1} + B_{rbdabio} + B_{rbdbio} + B_{rbdai1a} + B_{rbdabi1a} + B_{rbdbi1a}) + (I_{rbdvgpi2} + I_{rbdvgi2} + I_{rbdvi2} + I_{rbdoi2} + I_{rbdai2} + I_{rbdabi2} + I_{rbdbi2}) + (B_{rbdai2} + B_{rbdvgpi2} + B_{rbdvgi2} + B_{rbdvi2} + B_{rbdoi2} + B_{rbdai2} + B_{rbdabi2} + B_{rbdbi2})$$

$$T_{IB} = (I_{doio} + I_{daio} + I_{dabi1} + I_{dbi1} + I_{doi1} + I_{dabio} + I_{dbio} + I_{dai1a} + I_{dabi1a} + I_{dbi1a}) + (B_{doio} + B_{daio} + B_{dabi1} + B_{dbi1} + B_{doi1} + B_{dabio} + B_{dbio} + B_{dai1a} + B_{dabi1a} + B_{dbi1a}) + (I_{dvvgpi2} + I_{dvvgi2} + I_{dvgi2} + I_{doi2} + I_{dai2} + I_{dabi2} + I_{dbi2}) + (B_{dvvgpi2} + B_{dvvgi2} + B_{dvgi2} + B_{doi2} + B_{dai2} + B_{dabi2} + B_{dbi2})$$

$$\beta = \text{amp} * ((1-\text{bn}) * \beta_n) * \cos(2 * 3.14159 * (\text{time} - \pi)) + ((1-\text{bn}) * \beta_n)$$

$\pi = 0.5$; peak time for malaria transmission with 0=January

$\text{amp} = 0.67$; amplitude of seasonal variation with value 0 to 1

$$q = (\mu + \gamma) * (\mu + \sigma) * (\mu + \delta) / (\sigma * \gamma)$$

$\text{cost}_a = 0$; cost of resistance to artesunate

$\text{cost}_b = 0$; cost of resistance to piperaquine

$$T_{Iro} = I_{rodoio} + I_{rodaio} + I_{rodabil} + I_{rodbi1} + I_{rodoi1} + I_{rodabio} + I_{rodbio} + I_{rodai1a} + I_{radabi1a} + I_{radbi1a} + I_{rodvgpi2} + I_{rodvgi2} + I_{rodvi2} + I_{rodoi2} + I_{rodai2} + I_{radabi2} + I_{radbi2}$$

$$T_{Ira} = (1 - \text{cost}_a) * (I_{radoio} + I_{radaio} + I_{radabi1} + I_{radbi1} + I_{radoi1} + I_{radabio} + I_{radbio} + I_{radai1a} + I_{radabi1a} + I_{radbi1a} + I_{radvgpi2} + I_{radvgi2} + I_{radvi2} + I_{radoi2} + I_{radai2} + I_{radabi2} + I_{radbi2})$$

$$T_{Irb} = (1 - \text{cost}_b) * (I_{rbdoio} + I_{rbdaio} + I_{rbdabi1} + I_{rbdbi1} + I_{rbdoi1} + I_{rbdabio} + I_{rbdbio} + I_{rbdai1a} + I_{rbdabi1a} + I_{rbdbi1a} + I_{rbdvgpi2} + I_{rbdvgi2} + I_{rbdvi2} + I_{rbdoi2} + I_{rbdai2} + I_{rbdabi2} + I_{rbdbi2})$$

$$T_I = T_{Iro} + T_{Ira} + T_{Irb}$$

$$T_{i1cum}' = \tau_1 * (N_{doio} + N_{daio})$$

$$T_{i1cumcov} = (100 * T_{i1cum}) / T_N$$

$$\text{init } T_{i1cum} = 0$$

$\text{perc}_{ra} = 100 * N_{ra} / T_{IB}$; percent of blood stage infections resistant to artesunate

$\text{perc}_{rb} = 100 * N_{rb} / T_{IB}$, percent of blood stage infections resistant to piperaquine

$\text{perc}_{IB} = 100 * T_{IB} / T_N$; percent of infections that are blood stage

$$\text{init}(IT_{roda}) = 0$$

$$IT_{roda} = B_{rodaio} + B_{rodai1a} + L_{rodaio} + L_{rodai1a} + I_{rodaio} + I_{rodai1a} + B_{rodabio} + B_{rodabi1} + B_{rodabi1a} + L_{rodabio} + L_{rodabi1} + L_{rodabi1a} + I_{rodabio} + I_{rodabi1} + I_{rodabi1a} + B_{rodabi2} + L_{rodabi2} + I_{rodabi2} + B_{rodai2} + L_{rodai2} + I_{rodai2}$$

$$\text{init}(IT_{rodb}) = 0$$

$$IT_{rodb}' = B_{rodbio} + B_{rodbi1} + B_{rodbi1a} + L_{rodbio} + L_{rodbi1} + L_{rodbi1a} + I_{rodbio} + I_{rodbi1} + I_{rodbi1a} + B_{rodabio} + B_{rodabi1} + B_{rodabi1a} + L_{rodabio} + L_{rodabi1} + L_{rodabi1a} + I_{rodabio} + I_{rodabi1} + I_{rodabi1a} + B_{rodabi2} + L_{rodabi2} + I_{rodabi2} + B_{rodbi2} + L_{rodbi2} + I_{rodbi2}$$

$init(IT_{rada})=0$

$$IT_{rada}' = B_{radaio} + B_{radai1a} + L_{radaio} + L_{radai1a} + I_{radaio} + I_{radai1a} + B_{radabio} + B_{radabi1} + B_{radabi1a} + L_{radabio} + L_{radabi1} + L_{radabi1a} + I_{radabio} + I_{radabi1} + I_{radabi1a} + B_{radabi2} + L_{radabi2} + I_{radabi2} + B_{radai2} + L_{radai2} + I_{radai2}$$

$init(IT_{radb})=0$

$$IT_{radb}' = B_{radbio} + B_{radbi1} + B_{radbi1a} + L_{radbio} + L_{radbi1} + L_{radbi1a} + I_{radbio} + I_{radbi1} + I_{radbi1a} + B_{radabio} + B_{radabi1} + B_{radabi1a} + L_{radabio} + L_{radabi1} + L_{radabi1a} + I_{radabio} + I_{radabi1} + I_{radabi1a} + B_{radabi2} + L_{radabi2} + I_{radabi2} + B_{radbi2} + L_{radbi2} + I_{radbi2}$$

$init(IT_{rbda})=0$

$$IT_{rbda}' = B_{rbdaio} + B_{rbdai1a} + L_{rbdaio} + L_{rbdai1a} + I_{rbdaio} + I_{rbdai1a} + B_{rbdabio} + B_{rbdabi1} + B_{rbdabi1a} + L_{rbdabio} + L_{rbdabi1} + L_{rbdabi1a} + I_{rbdabio} + I_{rbdabi1} + I_{rbdabi1a} + B_{rbdabi2} + L_{rbdabi2} + I_{rbdabi2} + B_{rbdai2} + L_{rbdai2} + I_{rbdai2}$$

$init(IT_{rbdb})=0$

$$IT_{rbdb}' = B_{rbdbio} + B_{rbdbi1} + B_{rbdbi1a} + L_{rbdbio} + L_{rbdbi1} + L_{rbdbi1a} + I_{rbdbio} + I_{rbdbi1} + I_{rbdbi1a} + B_{rbdabio} + B_{rbdabi1} + B_{rbdabi1a} + L_{rbdabio} + L_{rbdabi1} + L_{rbdabi1a} + I_{rbdabio} + I_{rbdabi1} + I_{rbdabi1a} + B_{rbdabi2} + L_{rbdabi2} + I_{rbdabi2} + B_{rbdbi2} + L_{rbdbi2} + I_{rbdbi2}$$