

## 1 Supplementary Methods: MBDETES

Let the behavioral state variable name from MBITES denote the expected number of mosquitoes of a given chronological age ( $a$ ), that are in each behavioral state; *e.g.*, the proportion in the post-prandial resting state (R) of age  $a$  at time  $t$  is  $R(a, t)$ . Similarly, let the waiting times to events be modeled as a rate that is dependent on the behavioral state ( $x$ ), age, and time ( $t$ ):  $\xi_x(a, t)$ . The proportion of mosquitoes that transition from state  $x$  to state  $y$  at the end of a bout is denoted  $P_{xy}$ . Death rates can be age dependent (*i.e.*, due to senescence), which affects the proportions transitioning to other states, so we write  $P_{xy}(a)$ . To deal with the event-driven nature of these bouts and the possibility some bouts may be repeated many times before transitioning to another state, we index mosquitoes by the  $i^{th}$  attempt to repeat the same event as a way of computing waiting times properly; for example, a mosquito repeating a blood feeding attempt bout transitions from  $B_n(a)$  to  $B_{n+1}(a)$ . Finally, we let  $\Lambda(t)$  represent the rate of emergence of adult female mosquitoes. The following system of coupled PDEs is homologous to MBITES:

$$F_1(0, t) = \Lambda(t)$$

$$\begin{aligned} \frac{\partial F_1(a, t)}{\partial t} + \frac{\partial F_1(a, t)}{\partial a} = & \xi_O(a, t)P_{OF}(a) \sum_i O_i(a, t) + \xi_B(a, t)P_{BF}(a) \sum_i B_i(a, t) \\ & + \xi_R(a, t)P_{RF}(a)R(a, t) - \xi_F(a, t)F_1(a, t) \end{aligned}$$

$$\frac{\partial F_i(a, t)}{\partial t} + \frac{\partial F_i(a, t)}{\partial a} = \xi_F(a, t)P_{FF}(a)F_{i-1}(a, t) - \xi_F(a, t)F_i(a, t)$$

$$\begin{aligned} \frac{\partial B_1(a, t)}{\partial t} + \frac{\partial B_1(a, t)}{\partial a} = & \xi_O(a, t)P_{OB}(a) \sum_i O_i(a, t) + \xi_F(a, t)P_{FB}(a) \sum_i F_i(a, t) \\ & + \xi_R(a, t)P_{RB}(a)R(a, t) \\ & - \xi_B(a, t)B_1(a, t) \end{aligned}$$

$$\frac{\partial B_i(a, t)}{\partial t} + \frac{\partial B_i(a, t)}{\partial a} = \xi_B(a, t)P_{BB}(a)B_{i-1}(a, t) - \xi_B(a, t)B_i(a, t)$$

$$\frac{\partial R(a, t)}{\partial t} + \frac{\partial R(a, t)}{\partial a} = \xi_B(a, t)P_{BR}(a) \sum_i B_i(a, t) - \xi_R(a, t)R(a, t)$$

$$\begin{aligned} \frac{\partial L_1(a, t)}{\partial t} + \frac{\partial L_1(a, t)}{\partial a} = & \xi_R(a, t)P_{RL}(a)R(a, t) + \xi_O(a, t)P_{OL}(a) \sum_i O_i(a, t) \\ & - \xi_L(a, t)L_1(a, t) \end{aligned}$$

$$\frac{\partial L_i(a, t)}{\partial t} + \frac{\partial L_i(a, t)}{\partial a} = \xi_L(a, t)P_{LL}(a)L_{i-1}(a, t) - \xi_L(a, t)L_i(a, t)$$

$$\begin{aligned} \frac{\partial O_1(a, t)}{\partial t} + \frac{\partial O_1(a, t)}{\partial a} = & \xi_L(a, t)P_{LO}(a) \sum_i L_i(a, t) + \xi_R(a, t)P_{RO}(a)R(a, t) \\ & - \xi_O(a, t)O_1(a, t) \end{aligned}$$

$$\frac{\partial O_i(a, t)}{\partial t} + \frac{\partial O_i(a, t)}{\partial a} = \xi_O(a, t)P_{OO}(a)O_{i-1}(a, t) - \xi_O(a, t)O_i(a, t) \quad (1)$$

It is a nuisance to deal with an infinite set of equations, but if the state transitions are Markovian, then a change of variables to lump the the  $n^{th}$  states together:  $x = \sum_i x_i$ , with a rescaled rate variable,  $\gamma_x(a, t) = \xi(a, t)(1 - P_{xx})$

### Proof 1

To justify this summation, consider the infinite set of equations

$$\frac{dx_1}{dt} = -\lambda x_1$$

$$\frac{dx_i}{dt} = p\lambda x_{i-1} - \lambda x_i$$

with initial conditions  $x_1(0) = 1$ ,  $x_i(0) = 0$ . That is, initially all of the mosquitoes are in their first attempt for an exponentially distributed length of time. A proportion  $p$  are successful or leave frustrated, and  $1 - p$  attempt

again. This system can be solved iteratively;  $x_1$  has the solution

$$x_1(t) = e^{-\lambda t}$$

which, when plugged into the equation for  $x_2$ , gives

$$\frac{dx_2}{dt} = p\lambda e^{-\lambda t} - \lambda x_2$$

which can be solved using an integrating factor. This yields

$$x_2(t) = p\lambda t e^{-\lambda t}$$

This appears to be a weighted gamma distribution, which motivates the assumption for an inductive-step solution of

$$x_i(t) = \frac{(p\lambda t)^{i-1}}{(i-1)!} e^{-\lambda t}$$

Assuming this, we look at the  $x_{i+1}$  equation:

$$\frac{dx_{i+1}}{dt} = p\lambda x_i - \lambda x_{i+1}$$

Plugging in the assumed solution, we get

$$\begin{aligned} \frac{dx_{i+1}}{dt} &= p\lambda \frac{(p\lambda t)^{i-1}}{(i-1)!} e^{-\lambda t} - \lambda x_{i+1} \\ &= \frac{(p\lambda)^i}{i!} t^{i-1} e^{-\lambda t} - \lambda x_{i+1} \end{aligned}$$

Which is again amenable to an integrating factor combined with i-1 integration by parts, yielding

$$x_{i+1} = \frac{(p\lambda t)^i}{i!} e^{-\lambda t}$$

which completes the induction. Because we are interested in the total amount of time spent and not the time spent in any one compartment, we add the solutions together:

$$\begin{aligned} \sum_{i=1}^{\infty} x_i(t) &= \sum_{i=1}^{\infty} \frac{(p\lambda t)^{i-1}}{(i-1)!} e^{-\lambda t} \\ &= e^{-\lambda t} \sum_{i=0}^{\infty} \frac{(p\lambda t)^i}{i!} \\ &= e^{-\lambda t} e^{p\lambda t} \\ &= e^{-(1-p)\lambda t} \end{aligned}$$

Normalizing this gives us the total expected waiting time in this state for the mosquito is exponential with intensity  $(1 - p)\lambda$ .

As a note, this convergence is uniform as it is the Taylor series representation of the exponential function - this justifies summing the infinite equations and the differential operator which gives us

$$\begin{aligned}\frac{d}{dt} \sum_{i=1}^{\infty} x_i &= \sum_{i=1}^{\infty} \frac{dx_i}{dt} \\ &= \sum_{i=1}^{\infty} -\lambda(1 - p)x_i\end{aligned}$$

setting  $X = \sum_{i=1}^{\infty} x_i$ , we get a very simple single differential equation

$$\frac{dX}{dt} = -(1 - p)\lambda X$$

with the initial condition  $X(0) = 1$ . This has the same solution we found through induction.

### **Proof 2**

Say we want to find the total waiting time  $T$  a mosquito spends in a given state. Using the law of total probability, we can condition this on the number of attempts  $N$  a mosquito will make:

$$P(T = t) = \sum_{n=1}^{\infty} P(T = t | N = n) P(N = n)$$

The number of attempts is geometrically distributed, as it will succeed or give up with probability  $(1-p)$  and therefore try again with probability  $p$  - it will try until it succeeds or gives up. Therefore

$$P(N = n) = (1 - p)p^{n-1}$$

The waiting time between each attempt is iid with an exponentially distributed waiting time with intensity  $\lambda$ , so given it takes  $n$  trials the distribution follows a gamma distribution:

$$\begin{aligned}T | N &= \sum_{i=1}^n \text{Exp}_i(\lambda) \\ &\sim \Gamma(n, \lambda)\end{aligned}$$

Therefore we have

$$P(T = t) = \sum_{i=1}^{\infty} P(T = t | N = n) P(N = n)$$

$$\begin{aligned}
&= \sum_{i=1}^{\infty} (1-p)p^{n-1} \frac{\lambda^n}{\Gamma(n)} t^{n-1} e^{-\lambda t} \\
&= (1-p)\lambda e^{-\lambda t} \sum_{i=0}^{\infty} \frac{(p\lambda t)^n}{n!} \\
&= \lambda(1-p)e^{-\lambda t} e^{p\lambda t} \\
&= (1-p)\lambda e^{-(1-p)\lambda t}
\end{aligned}$$

which is exactly the normalized solution to the previous system of ordinary differential equations - the waiting time for the mosquito to leave the state is exponentially distributed with intensity  $(1-p)\lambda$ . Importantly, this proof does not depend on  $(1-p)$  or  $\lambda$  being constant - they can be age- or time-dependent.

This means we can rewrite the infinite system of equations as a set of 5 differential equations:

$$F(0, t) = \Lambda(t)$$

$$\begin{aligned}
\frac{\partial F(a, t)}{\partial t} + \frac{\partial F(a, t)}{\partial a} &= \gamma_O(a, t)P_{OF}(a)O(a, t) + \gamma_B(a, t)P_{BF}(a)B(a, t) \\
&\quad + \gamma_R(a, t)P_{RF}(a)R(a, t) - \gamma_F(a, t)(1 - P_{FF}(a))F(a, t) \\
\frac{\partial B(a, t)}{\partial t} + \frac{\partial B(a, t)}{\partial a} &= \gamma_O(a, t)P_{OB}(a)O(a, t) + \gamma_F(a, t)P_{FB}(a)F(a, t) \\
&\quad + \gamma_R(a, t)P_{RB}(a)R(a, t) - \gamma_B(a, t)(1 - P_{BB}(a))B(a, t) \\
\frac{\partial R(a, t)}{\partial t} + \frac{\partial R(a, t)}{\partial a} &= \gamma_B(a, t)P_{BR}(a)B(a, t) - \gamma_R(a, t)R(a, t) \\
\frac{\partial L(a, t)}{\partial t} + \frac{\partial L(a, t)}{\partial a} &= \gamma_R(a, t)P_{RL}(a)R(a, t) + \gamma_O(a, t)P_{OL}(a)O(a, t) \\
&\quad - \gamma_L(a, t)(1 - P_{LL}(a))L(a, t) \\
\frac{\partial O(a, t)}{\partial t} + \frac{\partial O(a, t)}{\partial a} &= \gamma_L(a, t)P_{LO}(a)L(a, t) + \gamma_R(a, t)P_{RO}(a)R(a, t) \\
&\quad - \gamma_O(a, t)(1 - P_{OO}(a))O(a, t)
\end{aligned} \tag{2}$$

### 1.1 The MBITES-de for Cohorts

Finally, we want a version of these equations to model changes in a cohort of individuals with respect to age (assuming all the mosquitoes emerge from

aquatic habitats at the same time of day):

$$\begin{aligned}
F(0) &= 1 \\
\dot{F} &= \gamma_O(a)P_{OF}(a)O(a) + \gamma_B(a)P_{BF}(a)B(a) + \gamma_R(a,t)P_{RF}(a)R(a) \\
&\quad - \gamma_F(a)(1 - P_{FF}(a))F(a) \\
\dot{B} &= \gamma_O(a)P_{OB}(a)O(a) + \gamma_F(a)P_{FB}(a)F(a) + \gamma_R(a)P_{RB}(a)R(a) \\
&\quad - \gamma_B(a)(1 - P_{BB}(a))B(a) \\
\dot{R} &= \gamma_B(a)P_{BR}(a)B(a) - \gamma_R(a)R(a) \\
\dot{L} &= \gamma_R(a)P_{RL}(a)R(a) + \gamma_O(a)P_{OL}(a)O(a) \\
&\quad - \gamma_L(a)(1 - P_{LL}(a))L_e(a) \\
\dot{O} &= \gamma_L(a)P_{LO}(a)L(a) + \gamma_R(a)P_{RO}(a)R(a) \\
&\quad - \gamma_O(a)(1 - P_{OO}(a))O(a)
\end{aligned} \tag{3}$$

## 1.2 Infection Dynamics in the MBITES-de Equations

To simulate infection dynamics in MBITES-de, we subdivide each variable  $X$  into new variables  $X_x$ ,  $x \in \{U, Y, Z\}$ , to represent the fraction of mosquitoes in behavioral state  $X$  that are uninfected,  $U$ , infected,  $Y$ , or infected and infectious  $Z$ . These lead to the following systems of coupled differential equations that remain unchanged, but for the equation describing resting mosquitoes. We let  $Q\kappa(t)$  the proportion of mosquitoes becoming infected after blood feeding at time  $t$ .

$$\begin{aligned}
\frac{\partial R_U(a,t)}{\partial t} + \frac{\partial R_U(a,t)}{\partial a} &= (1 - Q\kappa(t)) \gamma_B(a,t)P_{BR}(a)B_U(a,t) - \gamma_R(a,t)R_U(a,t) \\
\frac{\partial R_Y(a,t)}{\partial t} + \frac{\partial R_Y(a,t)}{\partial a} &= Q\kappa(t)\gamma_B(a,t)P_{BR}(a)B_U(a,t) \\
&\quad + \gamma_B(a,t)P_{BR}(a)B_Y(a,t) - \gamma_R(a,t)R_Y(a,t)
\end{aligned} \tag{4}$$

We let  $\tau(t)$  denote the (possibly time-dependent) extrinsic incubation period. Because  $\tau(t)$  is time dependent, we let  $\hat{t}$  denote that point in the past when the mosquito became infected in order to become infectious at time  $t$ : *i.e.*,  $t = \hat{t} + \tau(\hat{t})$ . Let  $\rho(t)$  the proportion of mosquitoes surviving through the extrinsic incubation period (*i.e.*, from  $\hat{t}$  to  $t = \hat{t} + \tau(\hat{t})$ ). An equation describing the proportion of infectious mosquitoes is:

$$\begin{aligned}
\frac{\partial R_Z(a,t)}{\partial t} + \frac{\partial R_Z(a,t)}{\partial a} &= \rho(t)Q\kappa(\hat{t}) \gamma_B(a,t)P_{BR}(a)B_U(a,t) \\
&\quad + \gamma_B(a,t)P_{BR}(a)B_Z(a,t) - \gamma_R(a,t)R_Z(a,t)
\end{aligned} \tag{5}$$

The remaining equations remain as follows:

$$\begin{aligned}
F_x(0, t) &= \Lambda(t) \\
\frac{\partial F_x(a, t)}{\partial t} + \frac{\partial F_x(a, t)}{\partial a} &= \gamma_O(a, t)P_{OF}(a)O_x(a, t) + \gamma_B(a, t)P_{BF}(a)B_x(a, t) \\
&\quad + \gamma_R(a, t)P_{RF}(a)R_x(a, t) - \gamma_F(a, t)(1 - P_{FF}(a))F_x(a, t) \\
\frac{\partial B_{O,x}(a, t)}{\partial t} + \frac{\partial B_x(a, t)}{\partial a} &= \gamma_O(a, t)P_{OB}(a)O_x(a, t) + \gamma_F(a, t)P_{FB}(a)F(a, t) \\
&\quad + \gamma_R(a, t)P_{RB}(a)R_x(a, t) - \gamma_B(a, t)(1 - P_{BB}(a))B_x(a, t) \\
\frac{\partial L_x(a, t)}{\partial t} + \frac{\partial L_x(a, t)}{\partial a} &= \gamma_R(a, t)P_{RL}(a)R_x(a, t) + \gamma_O(a, t)P_{OL}(a)O_x(a, t) \\
&\quad - \gamma_L(a, t)(1 - P_{LL}(a))L_x(a, t) \\
\frac{\partial O_{O,x}(a, t)}{\partial t} + \frac{\partial O_x(a, t)}{\partial a} &= \gamma_L(a, t)P_{LO}(a)L(a, t) + \gamma_R(a, t)P_{RO}(a)R_x(a, t) \\
&\quad - \gamma_O(a, t)(1 - P_{OO}(a))O_x(a, t)
\end{aligned} \tag{6}$$