

On the efficiency of HIV transmission: Insights through discrete time HIV models

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Appendix

Basic properties of the cell free viral spread model

The cell-free viral spread model is given by the following system of equations;

$$D_{t+1} = \beta_1 V_t, \quad (1)$$

$$P_{t+1} = \theta_1 D_t + \theta_3 P_t, \quad (2)$$

$$Q_{t+1} = \theta_2 P_t + \theta_3 Q_t, \quad (3)$$

$$V_{t+1} = \theta_2 \phi Q_t T_t^*, \quad (4)$$

$$T_{t+1} = s_T + \gamma T_t \exp\left(-\frac{T_t + T_t^*}{K} - \beta_1 V_t\right), \quad (5)$$

$$T_{t+1}^* = T_t [1 - \exp(-\beta_1 V_t)] + (1 - \mu) T_t^*, \quad (6)$$

The system of equations (1)-(6) will be studied in a biologically feasible region. We begin by showing that the solutions of this system of equations are bounded.

Proposition 1. *The system of equations (1)-(6) is bounded.*

Proof. For any $t_0 \in \mathbb{Z}^+$, eq (5), gives

$$\begin{aligned} T_{t_0+1} &\leq s_T + \gamma T_{t_0} \exp\left(-\frac{T_{t_0}}{K}\right), \\ &\leq s_T + \gamma K \exp(-1), \end{aligned} \quad (7)$$

where we have used $\max_{x \in \mathbb{R}} x \exp(-x/K) = K \exp(-1)$. We claim that

$$T_t \leq s_T + \gamma K \exp(-1) \quad \text{for } t \geq t_0.$$

We prove this by contradiction. Assume that there exist a number $\bar{l}_0 > t_0$ such that $T_{\bar{l}_0} > s_T + \gamma K \exp(-1)$. Then $\bar{l}_0 \geq t_0 + 2$. From the previous argument we have

$$\begin{aligned} T_{\bar{l}_0} &\leq s_T + \gamma T_{\bar{l}_0-1} \exp\left(-\frac{T_{\bar{l}_0-1}}{K}\right), \\ &\leq s_T + \gamma K \exp(-1), \end{aligned} \quad (8)$$

which is a contradiction.

From eq (6), it can be deduced that,

$$\begin{aligned} T_{t+1}^* &\leq T_t + (1 - \mu)T_t^* \\ &\leq s_T + \gamma K \exp(-1) + (1 - \mu)T_t^*. \end{aligned}$$

The solution of $T_{t+1}^* = s_T + \gamma K \exp(-1) + (1 - \mu)T_t^*$ is given by

$$T_t^* = (1 - \mu)^t T_0^* + (s_T + \gamma K \exp(-1)) \sum_{i=0}^{t-1} (1 - \mu)^i.$$

However, $\sum_{i=0}^{t-1} (1 - \mu)^i$ is a geometric series such that

$$T_t^* = \begin{cases} (1 - \mu)^t T_0^* + (s_T + \gamma K \exp(-1)) \left(\frac{1 - (1 - \mu)^t}{\mu} \right) & \text{if } (1 - \mu) \neq 1, \\ T_0^* + (s_T + \gamma K \exp(-1))t & \text{if } (1 - \mu) = 1. \end{cases}$$

The parameter $\mu \neq 0$ and thus we have

$$T_t^* \leq (1 - \mu)^t T_0^* + (s_T + \gamma K \exp(-1)) \left(\frac{1 - (1 - \mu)^t}{\mu} \right).$$

In this case

$$\limsup_{t \rightarrow \infty} T_t^* \leq \frac{s_T + \gamma K \exp(-1)}{\mu}.$$

If we let

$$n(t) = \begin{pmatrix} T_t & T_t^* \end{pmatrix}' \text{ and } \hat{n} = \begin{pmatrix} s_T + \gamma K \exp(-1) & \frac{s_T + \gamma K \exp(-1)}{\mu} \end{pmatrix}',$$

then

$$n(t) \leq \hat{n} \text{ for all } t.$$

Observe that

$$N(t+1) \leq \hat{A}N(t), \tag{9}$$

where

$$\hat{A} = \begin{pmatrix} 0 & 0 & 0 & \beta_1 \\ \theta_1 & \theta_3 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 \\ 0 & 0 & \theta_2 \phi \frac{s_T + \gamma K \exp(-1)}{\mu} & 0 \end{pmatrix}.$$

Define

$$N_1(t+1) = \hat{A}N_1(t), \quad t > 0, \quad N_1(0) = \hat{A}N(0),$$

then

$$N_1(t) = \hat{A}^t N_1(0).$$

The spectral radius of \hat{A} , is less than 1, and thus $N_1(t)$, is decreasing sequence. This means that $\hat{A}^t N_1(0) \rightarrow \mathbf{0}$, as $t \rightarrow \infty$, where $\mathbf{0} = \begin{pmatrix} 0 & 0 & 0 & 0 \end{pmatrix}$. We therefore get that

$$\liminf_{t \rightarrow \infty} N_1(t) \geq \mathbf{0} \quad (10)$$

From equality (9) and (10), it can be seen that $\mathbf{0} \leq N(t) \leq N_1(0)$. \square

We thus have the following proposition.

Proposition 2. *The set defined by*

$$\Omega = \{N(t), n(t) \in \mathbb{R}_+^4 \times \mathbb{R}_+^2 : \mathbf{0} \leq N(t) \leq N_1(0), \bar{\mathbf{0}} \leq n(t) \leq \hat{n}\},$$

where $\bar{\mathbf{0}} = \begin{pmatrix} 0 & 0 \end{pmatrix}$, is positively invariant under the flows of the system of equations (1)-(6).

Fixed points

The equilibrium points (fixed points) are obtained by equating the right hand side to the left hand side of the system of equations (1)-(6). If we denote the equilibrium point by

$\left(D^*, P^*, Q^*, V^*, T^*, T^{**} \right)'$, we have the following system of equations

$$D^* = \beta_1 V^*, \quad (11)$$

$$P^* = \theta_1 D^* + \theta_3 P^*, \quad (12)$$

$$Q^* = \theta_2 P^* + \theta_3 Q^*, \quad (13)$$

$$V^* = \theta_2 \phi Q^* T^{**}, \quad (14)$$

$$T^* = s_T + T^* \gamma \exp\left(-\beta_1 V^* - \frac{T^* + T^{**}}{K}\right), \quad (15)$$

$$T^{**} = T^* (1 - \exp(-\beta_1 V^*)) + (1 - \mu) T^{**}. \quad (16)$$

Substituting eq (11) into eq (12) we have

$$P^* = \frac{\theta_1}{1 - \theta_3} (\beta_1 V^*). \quad (17)$$

Eq (17) into eq (13) result in

$$Q^* = \frac{\theta_1 \theta_2}{(1 - \theta_3)^2} (\beta_1 V^*). \quad (18)$$

Substituting eq (18) into eq (14) we have

$$V^* = \frac{\theta_1 \theta_2^2 \phi}{(1 - \theta_3)^2} (\beta_1 V^*) T^{**}. \quad (19)$$

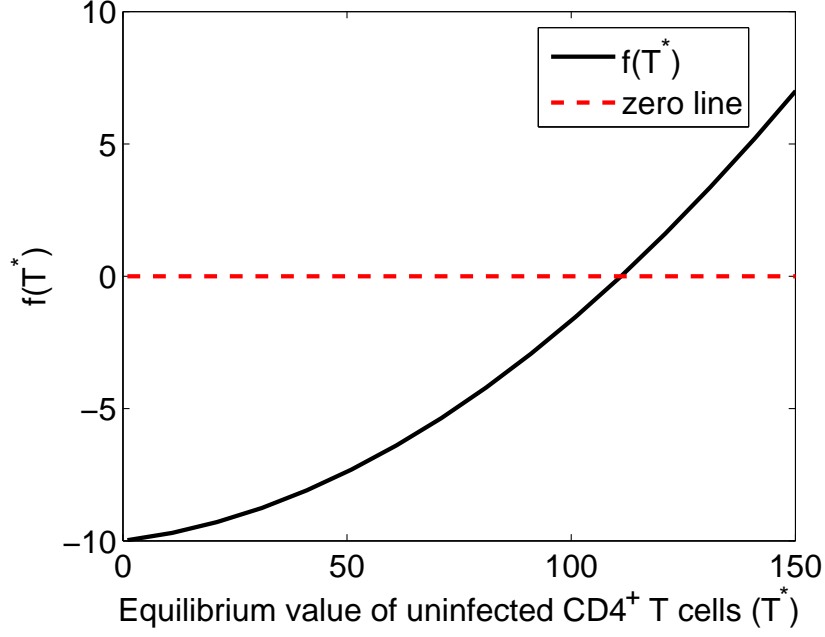


Figure 1: A graph of $f(T^*)$ against T^* . The parameters used were $s_T = 10$, $K = 1500$, $\gamma = 2.7$. It can be seen that the graph of $f(T^*)$ crosses the x-axis and thus there is solution for this parameter set.

From eq (19), we get $V^* = 0$, which gives the disease free equilibrium point or $T^{**} = \frac{(1-\theta_3)^2}{\theta_1\theta_2^2\phi\beta_1}$. Substituting $V^* = 0$ into equations (11)-(16) we get the disease free equilibrium

$$E_0 = \left(0, 0, 0, 0, T^*, 0 \right)',$$

where T^* is a solution of

$$T^* - s_T - T^*\gamma \exp\left(-\frac{T^*}{K}\right) = 0. \quad (20)$$

The analytical solution to the eq (20) is complex therefore, the graphical method is used to show the existence of the solution. Let $f(T^*) = T^* - s_T - T^*\gamma \exp\left(-\frac{T^*}{K}\right)$, then the graph of $f(T^*)$ against T^* is given in Figure 1. It can be seen from Figure 1 that the graph of $f(T^*)$ crosses the x-axis and thus there is a solution to the eq (20) for this parameter set.

From eq (16) we have

$$1 - \frac{\mu T^{**}}{T^*} = \exp(-\beta_1 V^*). \quad (21)$$

Substituting eq (21) into (15) result in

$$s_T - T^* + T^* \gamma \left(1 - \frac{\mu T^{**}}{T^*} \right) \exp\left(-\frac{T^*}{K}\right) = 0, \quad (22)$$

which can be simplified to give

$$s_T - T^* + \gamma (T^* - \mu T^{**}) \exp\left(-\frac{T^* + T^{**}}{K}\right) = 0, \quad (23)$$

where $T^{**} = \frac{(1-\theta_3)^2}{\theta_1 \theta_2^2 \phi \beta_1}$.

Stability of fixed points

The following theorem in [1] is used to study stability of the equilibrium points.

Theorem 1. *A fixed point x^* of a function $f(x)$ is asymptotically stable if all the eigenvalues μ of the first derivative, $Df(x)$ of $f(x)$ at x^* satisfy $|\mu| < 1$. The fixed point x^* is unstable if there exists an eigenvalue μ such that $|\mu| > 1$.*

Proposition 3. *The disease free equilibrium point*

$$E_0 = \left(0, 0, 0, 0, T^*, 0 \right)',$$

exist and is locally asymptotically stable when

$$\left| \gamma \left(1 - \frac{T^*}{K} \right) \exp\left(-\frac{T^*}{K}\right) \right| < 1,$$

where T^ is the solution of eq (20).*

Proof. Existence has been proved in the previous section. The Jacobian matrix at this equilibrium is given by

$$J_0 = \begin{pmatrix} 0 & 0 & 0 & \beta_1 & 0 & 0 \\ \theta_1 & \theta_3 & 0 & 0 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & 0 & M & G1 & 0 \\ 0 & 0 & 0 & -\beta_1 T^* & 0 & 1 - \mu \end{pmatrix},$$

where

$$M = -\beta_1 \gamma T^* \exp\left(-\frac{T^*}{K}\right),$$

and

$$G1 = \gamma \left(1 - \frac{T^*}{K}\right) \exp\left(-\frac{T^*}{K}\right).$$

All the eigenvalues J_0 (0 twice, θ_3 twice, G1 and $1 - \mu$) have magnitudes less than one if $|G1| < 1$. This completes the proof. \square

Proposition 4. *The system of equations (1)-(6), does not have an endermic equilibrium point for biologically feasible parameters.*

Basic properties of the cell-associated viral spread model

The cell-associated viral spread model is given by the following system of equations;

$$D_{t+1} = \beta_2 T_t^*, \quad (24)$$

$$P_{t+1} = \theta_1 D_t + \theta_3 P_t, \quad (25)$$

$$Q_{t+1} = \theta_2 P_t + \theta_3 Q_t, \quad (26)$$

$$V_{t+1} = \theta_2 \phi Q_t T_t^*, \quad (27)$$

$$T_{t+1} = s_T + \gamma T_t \exp\left(-\beta_2 T_t^* - \frac{T_t + T_t^*}{K}\right), \quad (28)$$

$$T_{t+1}^* = T_t (1 - \exp(-\beta_2 T_t^*)) + (1 - \mu) T_t^*. \quad (29)$$

Proposition 5. *The solutions of the system of equations (24)-(29) remain non negative and are bounded. The set Θ defined by $\Theta = \{N(t), n(t) \in \mathbb{R}^4 \times \mathbb{R}^2 : \mathbf{0} \leq N(t) \leq (I - \bar{G})^{-1} B \bar{n}, \bar{\mathbf{0}} \leq n(t) \leq \bar{n}\}$, where $\bar{n} = \left(s_T + \gamma K \exp(-1) \quad \frac{s_T + \gamma K \exp(-1)}{\mu} \right)'$, $\mathbf{0} = \left(0, 0, 0, 0 \right)'$, $\bar{G} = \begin{pmatrix} 0 & 0 & 0 & 0 \\ \theta_1 & \theta_3 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 \\ 0 & 0 & \theta_2 \phi \frac{s_T + \gamma K \exp(-1)}{\mu} & 0 \end{pmatrix}$ and $\bar{\mathbf{0}} = \left(0, 0 \right)'$ is positively invariant with respect to the flows of the system of equations (24)-(29).*

Proof. Using the procedure used to prove Proposition 1, it can easily be shown that

$$n(t) \leq \bar{n}$$

where

$$\bar{n} = \left(s_T + \gamma K \exp(-1) \quad \frac{s_T + \gamma K \exp(-1)}{\mu} \right)'.$$

Substituting inequality (9) and \bar{n} into equation of the virus at its different stages of the life cycle, yields

$$N(t+1) \leq \bar{G}N(t) + B\bar{n}. \quad (30)$$

We can prove by induction that $N(t) \leq \bar{G}^t N(0) + \sum_{i=0}^{t-1} (\bar{G}^i B\bar{n})$. It can then be shown that

$$\limsup_{t \rightarrow \infty} N(t) \leq (I - \bar{G})^{-1} B\bar{n}.$$

□

Equilibrium points

The equilibrium points (fixed points) are obtained by equating the right hand side to the left hand side of the system of equations (24)-(29). If we denote the equilibrium point by $(D, P, Q, V, T, T^*)'$, we have the following system of equations

$$D = \beta_2 T^*, \quad (31)$$

$$P = \theta_1 D + \theta_3 P, \quad (32)$$

$$Q = \theta_2 P + \theta_3 Q, \quad (33)$$

$$V = \theta_2 \phi Q T^*, \quad (34)$$

$$T = s_T + T\gamma \exp\left(-\beta_2 T^* - \frac{T + T^*}{K}\right), \quad (35)$$

$$T^* = T(1 - \exp(-\beta_2 T^*)) + (1 - \mu)T^*. \quad (36)$$

Substituting eq (31) into eq (32) we have

$$P = \frac{\theta_1}{1 - \theta_3} (\beta_2 T^*). \quad (37)$$

Eq (37) into eq (33) result in

$$Q = \frac{\theta_1 \theta_2}{(1 - \theta_3)^2} (\beta_2 T^*). \quad (38)$$

Substituting eq (38) into eq (34) we have

$$V = \frac{\theta_1 \theta_2^2 \phi}{(1 - \theta_3)^2} (\beta_2 T^*) T^*. \quad (39)$$

$V = 0$ in (39) implies $T^* = Q = P = 0$. Substituting $T^* = 0$ into equations (35) and (36) and solving the two equation for T , we get T which is a solution of equation

$$T - s_T - T\gamma \exp\left(-\frac{T}{K}\right) = 0. \quad (40)$$

Proposition 6. *The disease free equilibrium point exist and is locally asymptotically stable for*

$$\left| \gamma \left(1 - \frac{T}{K} \right) \exp \left(-\frac{T}{K} \right) \right| < 1$$

and

$$|-\beta_2 T + 1 - \mu_T| < 1$$

where T is a solution of eq (40).

Proof. The disease free equilibrium is given by

$$E_1 = \left(0, 0, 0, 0, T, 0 \right)',$$

where T is a solution of eq (40).

The Jacobian at E_1 is given by

$$J_{E_1} = \begin{pmatrix} 0 & 0 & 0 & 0 & 0 & \beta_2 \\ \theta_1 & \theta_3 & 0 & 0 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & G1 & -\gamma\beta_2 T \exp(-\frac{T}{K}) \\ 0 & 0 & 0 & 0 & 0 & -\beta_2 T + 1 - \mu_T \end{pmatrix}.$$

All the eigenvalues of J_{E_1} , (0 (twice), $G1$, θ_3 (twice) and $-\beta_2 T^* + 1 - \mu$) have magnitudes less than one if and only if $|G1| < 1$ and $|-\beta_2 T + 1 - \mu_T| < 1$. \square

The disease equilibrium point is obtained by equating the right hand side to the left hand side of the system of equations (24)-(29). If we denote the equilibrium point by $\left(D^*, P^*, Q^*, V^*, T^*, T^{**} \right)$ we have the following system of equations

$$D^* = \beta_2 T^{**}, \tag{41}$$

$$P^* = \theta_1 D^* + \theta_3 P^*, \tag{42}$$

$$Q^* = \theta_2 P^* + \theta_3 Q^*, \tag{43}$$

$$V^* = \theta_2 \phi Q^* T^{**}, \tag{44}$$

$$T^* = s_T + T^* \gamma \exp \left(-\beta_2 T^{**} - \frac{T^* + T^{**}}{K} \right), \tag{45}$$

$$T^{**} = T^* (1 - \exp(-\beta_2 T^{**})) + (1 - \mu) T^{**}. \tag{46}$$

Substituting eq (41) into eq (42) we have

$$P^* = \frac{\theta_1}{1 - \theta_3} (\beta_2 T^{**}). \tag{47}$$

Substituting eq (47) into eq (43) result in

$$Q^* = \frac{\theta_1 \theta_2}{(1 - \theta_3)^2} (\beta_2 T^{**}). \quad (48)$$

Substituting eq (48) into eq (44) we have

$$V^* = \frac{\theta_1 \theta_2^2 \phi}{(1 - \theta_3)^2} (\beta_2 T^{**}) T^{**}. \quad (49)$$

Making T^* subject of formula in eq (46) we get

$$T^* = \frac{\mu T^{**}}{1 - \exp(-\beta_2 T^{**})}. \quad (50)$$

Substituting this expression of T^* into eq (45) and simplifying the resulting equation we get

$$\mu T^{**} - s_T (1 - \exp(-\beta_2 T^{**})) - \gamma \mu T^{**} \exp\left(-\beta_2 T^{**} - \frac{\mu T^{**}}{K(1 - \exp(-\beta_2 T^{**}))} - \frac{T^{**}}{K}\right) = 0. \quad (51)$$

Solving for T^{**} is complex and we resort to graphical solutions. We let

$$f(T^{**}) = \mu T^{**} - s_T (1 - \exp(-\beta_2 T^{**})) - \gamma \mu T^{**} \exp\left(-\beta_2 T^{**} - \frac{\mu T^{**}}{K(1 - \exp(-\beta_2 T^{**}))} - \frac{T^{**}}{K}\right). \quad (52)$$

The cell-free and cell-associated viral spread model

The model that considers both forms transmission takes the form;

$$D_{t+1} = \beta_1 V_t + \beta_2 T_t^*, \quad (53)$$

$$P_{t+1} = \theta_1 D_t + \theta_3 P_t, \quad (54)$$

$$Q_{t+1} = \theta_2 P_t + \theta_3 Q_t, \quad (55)$$

$$V_{t+1} = \theta_2 \phi Q_t T_t^*, \quad (56)$$

$$T_{t+1} = s_T + \gamma T_t \exp\left(-\beta_1 V_t - \beta_2 T_t^* - \frac{T_t + T_t^*}{K}\right), \quad (57)$$

$$T_{t+1}^* = T_t (1 - \exp(-\beta_1 V_t - \beta_2 T_t^*)) + (1 - \mu) T_t^*. \quad (58)$$

Proposition 7. *The system of equations (53)-(58) has a disease free equilibrium point given by*

$$E_3 = \left(0, 0, 0, 0, T^*, 0 \right)',$$

where T^* is a solution of eq (20).

The Jacobian matrix at E_3 is given by

$$J_{E_3} = \begin{pmatrix} 0 & 0 & 0 & \beta_1 & 0 & \beta_2 \\ \theta_1 & \theta_3 & 0 & 0 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & 0 & -\gamma\beta_1 T^* \exp\left(-\frac{T^*}{K}\right) & G1 & -\gamma\beta_2 T^* \exp\left(-\frac{T^*}{K}\right) \\ 0 & 0 & 0 & -\beta_1 T^* & 0 & -\beta_2 T^* + 1 - \mu \end{pmatrix}.$$

The eigenvalues are 0 (twice), θ_3 (twice), $G1$, and $-\beta_2 T^* + 1 - \mu$. All the eigenvalues have magnitudes less than one if and only if $|G1| < 1$ and $|-\beta_2 T^* + 1 - \mu| < 1$. Showing that the system of equations (53)-(58) has an endemic equilibrium is maybe difficult, we use permanence to show the existence of the disease equilibrium.

Definition 1. *The system (53)-(58) is permanent if there exist positive constants m and M which are independent of the solution of system (53)-(58), such that any positive solution $\{D_t, P_t, Q_t, V_t, T_t, T_t^*\}$ of system (53)-(58) satisfies*

$$m \leq \liminf_{t \rightarrow \infty} \{D_t, P_t, Q_t, V_t, T_t, T_t^*\} \leq \limsup_{t \rightarrow \infty} \{D_t, P_t, Q_t, V_t, T_t, T_t^*\} \leq M.$$

Lemma 1. *Every solution $\{D_t, P_t, Q_t, V_t, T_t, T_t^*\}$ of system (53)-(58) satisfies $\limsup_{t \rightarrow \infty} n(t) \leq \bar{n}$ and $\limsup_{t \rightarrow \infty} N(t) \leq (I - \bar{G})^{-1} B \bar{n}$, where \bar{n} , \bar{G} are as previously defined.*

Proof. From the proof of Proposition 1 we get that

$$\limsup_{t \rightarrow \infty} T_t \leq s_T + \gamma K \exp(-1).$$

Using the same procedure as the one used in proving Proposition 1, we get

$$\limsup_{t \rightarrow \infty} T_t^* \leq \frac{s_T + \gamma K \exp(-1)}{\mu}$$

and

$$\limsup_{t \rightarrow \infty} N(t) \leq (I - \bar{G})^{-1} B \bar{n},$$

where \bar{n} , \bar{G} are as previously defined. □

Lemma 2. *Every solution $\{D_t, P_t, Q_t, V_t, T_t, T_t^*\}$ of system (53)-(58) satisfies $h \leq \liminf_{t \rightarrow \infty} n(t)$*

$$\text{and } H \leq \liminf_{t \rightarrow \infty} N(t) \text{ where } h = (H_1 \ H_2), \ H = (I - \tilde{G}) B h, \ \tilde{G} = \begin{pmatrix} 0 & 0 & 0 & 0 \\ \theta_1 & \theta_3 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 \\ 0 & 0 & \theta_2 \phi H_2 & 0 \end{pmatrix},$$

$H_1 = \gamma K \exp(-\beta_1 \bar{V} - \beta_2 \bar{T}^* - 1)$ and

$$H_2 = \gamma K \exp(-\beta_1 \bar{V} - \beta_2 \bar{T}^* - 1) \left(1 - \ln \left(\frac{\mu}{\beta_2 H_1} \right) \right) + \frac{\mu}{\beta_2} \ln \left(\frac{\mu}{\beta_2 H_1} \right).$$

Proof. According to Lemma 1, there exist a $t^* \in \mathbb{Z}^+$, such that

$$n(t) < \bar{n} + (\epsilon, \epsilon)', \quad N(t) < \bar{N} + (\epsilon, \epsilon, \epsilon, \epsilon)', \quad t \geq t^*$$

where $\bar{N} = (I - \bar{G})^{-1}B\bar{n}$. We first show that $\liminf_{t \rightarrow \infty} T_t \geq H_1$, where H_1 is to be determined. Assume that there exist a $t_0 \geq t^*$ such that

$$\begin{aligned} T_{t_0+1} &\geq s_T + \gamma T_{t_0} \exp\left(-\beta_1(\bar{V} + \epsilon) - \beta_2(\bar{T}^* + \epsilon) - \frac{T_{t_0}}{K}\right), \\ &\geq s_T + \gamma T_{t_0} \exp\left(-F - \frac{T_{t_0}}{K}\right), \\ &\geq \gamma T_{t_0} \exp\left(-F - \frac{T_{t_0}}{K}\right), \\ &\geq \gamma K \exp(-F - 1), \end{aligned}$$

where $-\beta_1(\bar{V} + \epsilon) - \beta_2(\bar{T}^* + \epsilon) = F$. We claim that

$$T_t \geq K \exp(-F - 1), \quad \text{for } t \geq t_0.$$

By way of contradiction, assume there exist a $\tau_0 > t_0$ such that $T_{\tau_0} < \gamma K \exp(-F - 1)$. Let $\bar{\tau}_0 = t_0 + 2$ be the smallest number such that $T_{\bar{\tau}_0} < \gamma K \exp(-F - 1)$. The argument presented above produces a contradiction and this proves the claim. Thus

$$\liminf_{t \rightarrow \infty} T_t \geq \gamma K \exp(-F - 1).$$

Setting $\epsilon \rightarrow 0$ leads to

$$\liminf_{t \rightarrow \infty} T_t \geq H_1 = \gamma K \exp(-\beta_1 \bar{V} - \beta_2 \bar{T}^* - 1).$$

We now need to show that $\liminf_{t \rightarrow \infty} T_t^* \geq H_2$, where H_2 is to be determined. From the above steps we have that for $t^* \in \mathbb{Z}^+$,

$$n(t) < \bar{n} + (\epsilon, \epsilon)', \quad N(t) < \bar{N} + (\epsilon, \epsilon, \epsilon, \epsilon)',$$

and $T_t \geq H_1 - \epsilon$, $t \geq t^*$. Assume that there exist $t_0 \geq t^*$ such that

$$\begin{aligned} T_{t_0+1}^* &\geq (H_1 - \epsilon) \left(1 - \exp(-\beta_1 V_{t_0} - \beta_2 T_{t_0}^*)\right) + (1 - \mu)T_{t_0}^*, \\ &\geq (H_1 - \epsilon) \left(1 - \exp(-\beta_2 T_{t_0}^*)\right) + (1 - \mu)T_{t_0}^*, \end{aligned} \tag{59}$$

$$\geq (H_1 - \epsilon) \left(1 - \exp(-\beta_2 T_{t_0}^*)\right) - \mu T_{t_0}^*, \tag{60}$$

$$\geq (H_1 - \epsilon) \left(1 - \ln\left(\frac{\mu}{\beta_2(H_1 - \epsilon)}\right)\right) + \frac{\mu}{\beta_2} \ln\left(\frac{\mu}{\beta_2(H_1 - \epsilon)}\right). \tag{61}$$

We claim that

$$T_t^* \geq H1_2, \text{ where } H1_2 = (H_1 - \epsilon) \left(1 - \ln \left(\frac{\mu}{\beta_2(H_1 - \epsilon)} \right) \right) + \frac{\mu}{\beta_2} \ln \left(\frac{\mu}{\beta_2(H_1 - \epsilon)} \right)$$

for $t \geq t_0$.

By way of contradiction assume there exists a $\tau_0 > t_0$ such that $T_{\tau_0}^* < H1_2$, then $\tau_0 \geq t_0 + 2$.

Let $\bar{\tau}_0 = t_0 + 2$ be the smallest number such that $T_{\bar{\tau}_0}^* < H1_2$. The above argument produces $T_{\bar{\tau}_0}^* \leq H1_2$, a contradiction and this proves the claim. Thus $\liminf_{t \rightarrow \infty} T_t^* \geq H1_2$. Setting $\epsilon \rightarrow 0$ leads to

$$\liminf_{t \rightarrow \infty} T_t \geq H_2 = H_1 \left(1 - \ln \left(\frac{\mu}{\beta_2 H_1} \right) \right) + \frac{\mu}{\beta_2} \ln \left(\frac{\mu}{\beta_2 H_1} \right)$$

We now show that $\liminf N(t) \geq H_3$ where H_3 is to be determined. We now have that $n(t) < \bar{n} + (\epsilon, \epsilon)'$, $N(t) < \bar{N} + (\epsilon, \epsilon, \epsilon, \epsilon)'$, and $T_t \geq H_1 - \epsilon$, $T_t^* \geq H_2 - \epsilon$, $t \geq t^*$.

Define $\hat{G} = \begin{pmatrix} 0 & 0 & 0 & 0 \\ \theta_1 & \theta_3 & 0 & 0 \\ 0 & \theta_2 & \theta_3 & 0 \\ 0 & 0 & \theta_2 \phi(H_2 - \epsilon) & 0 \end{pmatrix}$, and $\hat{n} = (H_1 - \epsilon, H_2 - \epsilon)'$. For any $t \geq 0$ we have that

$$N(t+1) = GN(t) + B\hat{n} \tag{62}$$

$$\geq \hat{G}N(t) + B\hat{n} \tag{63}$$

We can prove by induction that $N(t) \geq \hat{G}^t N(0) + \sum_{i=0}^{t-1} \hat{G}^i B\hat{n}$, from which it can be deduced that $\liminf_{t \rightarrow \infty} N(t) \geq (I - \hat{G})B\hat{n}$. Setting $\epsilon \rightarrow 0$ leads to $\liminf_{t \rightarrow \infty} N(t) \geq H_3 = (I - \tilde{G})B\hat{n}$. \square

Proposition 8. *The system of equations (53)-(58) is permanent.*

Proof. The result follows from Lemma 1 and Lemma 2. \square

The implication of this result is that an endemic equilibrium point exist.

References

- [1] Guckenheimer J. and Holmes P. 1983. Nonlinear oscillations, Dynamical systems, and bifurcations of vector fields. Springer-Verlag.