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A Mathematical Modeling of Tuberculosis Dynamics with Hygiene Consciousness as a Control Strategy

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Authors' contributions

This work was carried out in collaboration between bothauthors

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Abstract

Tuberculosis, an airborne infectious disease, remains a major threat to public health in Kenya. In this study, we derived a system of non-linear ordinary differential equations from the SLICR mathematical model of TB to study the effects of hygiene consciousness as a control strategy against TB in Kenya. The effective basic reproduction number (R_0) of the model was determined by the next generation matrix approach. We established and analyzed the equilibrium points. Using the Routh-Hurwitz criterion for local stability analysis and comparison theorem for global stability analysis, the disease-free equilibrium (DFE) was found to be locally asymptotically stable given that $R_0 < 1$.

Also by using the Routh-Hurwitz criterion for local stability analysis and Lyapunov function and LaSalle's invariance principle for global stability analysis, the endemic equilibrium (EE) point was found to be locally asymptotically stable given that $R_0 > 1$. Using MATLAB **ode45** solver, we simulated the model numerically and the results suggest that hygiene consciousness can help in controlling TB disease if incorporated effectively.

Keywords: Tuberculosis; Reproduction number; Stability; Numerical simulation.

2019 Mathematics Subject Classification: 53C25; 83C05; 57N16

1 Introduction

Tuberculosis is a bacterial infection disease that is common and potentially fatal if not timely treated. It is caused by the *mycobacterium tuberculosis*. Tuberculosis usually affects the lungs; it can also affect the brain, the central nervous system, the lymphatic system, the kidneys, and spine.

TB is transmitted from one person to another through the air. When a person with lung TB coughs, sneezes, speaks or spits, TB germs (bacterium) are propelled into the air. These germs enter into the respiratory system through the air inhaled. Any uninfected individual inhaling the bacteria is possibly exposed to infectious *mycobacterium tuberculosis*, An individual needs to inhale only a few of these bacteria to become infected.

Mathematical modeling is an important tool for a better understanding of the dynamics of infectious [1]. Modeling of tuberculosis was first carried out by Frost [2]. Since then, several researchers have continuously researched on how tuberculosis can be reduced using mathematical models by incorporating interventions such as treatment of tuberculosis-infected persons by using of drug chemotherapy [3], treatment of actively and latently infected persons [4], health education (see [5], [6],[7]), therapy of tuberculosis transmission [8], early treatment for latent patients and treatment of infective [9] and so on. Based on the result of these studies, the researchers found that tuberculosis could be controlled.

To this end, we seek to investigate analytically the dynamics of tuberculosis by incorporating hygiene as a control strategy in order to eradicate the disease.

2 Model Formulation

We formulate a model with total population N(t), which is divided into five classes: S(t)-Susceptible individuals, L(t)-Latently Infected individuals, I(t)- Infectious individuals (Actively Infected individuals), C(t)-Hygiene conscious individuals (Infectious individuals who have become hygiene conscious) and R(t)- Recovered individuals with natural death rate μ in all classes and TB related death rate δ in Infectious and hygiene consciousness compartments. People are recruited to Susceptible class at the rate Λ and become Latently infected at the rate of βI where β is the effective contact rate between susceptible and infectious individuals. $\nu\beta(0 < \nu < 1)$ is the reduced effective contact rate between susceptible and infectious individuals as a result of hygiene consciousness where ν is the rate at which infectious individuals become hygiene conscious. Latently infected individuals progress to infectious class at σ rate and these infectious individuals recover at the rate π and others progress to hygiene consciousness class at the rate ν . The hygiene conscious individuals recover at ω rate. From the above description we have the following assumptions and flow diagram;

Assumptions:

i The population birth and death rate occur at different rates. ii Infectious individuals observe hygiene willingly.

iii Infectious individuals recover as result of effective treatment. iv

There is permanent immunity after recovery.

- v All the newly born individuals join only susceptible class.
- vi Individuals in infectious and hygiene Conscious classes die naturally and as result of TB infection while those in other classes die naturally.

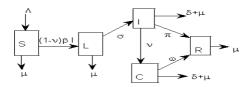


Figure 1: Flow chart

From Figure 1 we have the following equations of the model are:

$$\frac{dS}{dt} = \Lambda - (1 - v)\beta IS - \mu S$$

$$\frac{dL}{dt} = (1 - v)\beta IS - (\sigma + \mu)L$$

$$\frac{dI}{dt} = \sigma L - (v + \pi + \delta + \mu)I$$

$$\frac{dC}{dt} = vI - (\omega + \delta + \mu)C$$

$$\frac{dR}{dt} = \omega C + \pi I - \mu R$$
(2.1)

3 **Model Analysis**

Positivity and boundedness of solutions

The state variables of the model represent classes of human population and therefore should be non-negative for the model to be well posed. The model will make epidemiological sense where the feasible region is positively invariant. All the solutions are non-negative and bounded in the feasible region $\Omega = \{(S, L, I, \bar{C}, R)\}$

 $\in \mathbb{R}^5: S > 0; L, I, C, R \ge 0; N \le \frac{\Delta}{I}$ Because the first four equations of system (2.1) are independent of variable R(t), we reduce the system by to obtain;

$$\frac{dS}{dt} = \Lambda - (1 - v)\beta IS - \mu S$$

$$\frac{dL}{dt} = (1 - v)\beta IS - (\sigma + \mu)L$$

$$\frac{dI}{dt} = \sigma L - (v + \pi + \delta + \mu)I$$

$$\frac{dC}{dt} = vI - (\omega + \delta + \mu)C$$

$$\frac{dS}{dt} = (1 - v)\beta IS - \mu S$$

$$\frac{dL}{dt} = (1 - v)\beta IS - \mu S$$

$$\frac{dL}{dt} = (1 - v)\beta IS - \mu S$$

$$\frac{dL}{dt} = vI - (\omega + \delta + \mu)C$$

$$\frac{dS}{dt} = vI - (\omega + \delta + \mu)C$$

$$\frac{dS}{dt} = (1 - v)\beta IS - \mu S$$

(3.1)

Existence of Equilibrium points

At the equilibrium point, the right hand side of the system (3.1) is equal to zero. In that;

$$\Lambda - (1 - v)\beta IS - \mu S = 0 (1$$

$$- v)\beta IS - (\sigma + \mu)L = 0$$

$$\sigma L - (v + \pi + \delta + \mu)I = 0$$

$$vI - (\omega + \delta + \mu)C = 0$$
(3.2)

For Disease Free Equilibrium point E_0 , the disease is not present. Hence $S = S_0$, L = 0, I = 0 and C = 0. Upon substitution of these ($S = S_0$, L = 0, I = 0 and C = 0) in (3.2), we obtain $S_0 = {}^{\Lambda}$ Such that $E_0 = ({}^{\Lambda}, 0, 0, 0)$

Next, we investigate the existence of Endemic equilibrium point $E_e(S_e, L_e, I_e, C_e)$ by substituting $S = S_e$, $L = L_e$, $I = I_e$, $C = C_e$ in (3.2) to get

$$\Lambda - (1 - v)\beta I_e S_e - \mu S_e = 0 (1 - v)\beta I_e S_e - (\sigma + \mu) L_e = 0 \sigma L_e - (v + \pi + \delta + \mu) I_e = 0 v I_e - (\omega + \delta + \mu) C_e = 0$$
(3.3)

then by solving for S_e , L_e , I_e and C_e we obtain

$$E_{e} \begin{array}{c} & 2 \frac{(1-v)\beta\sigma\Lambda - \mu(\sigma+\mu)(v+\pi+\delta+\mu)}{(1-v)\mu\sigma\beta} \\ & S_{e} \\ & L_{e} \\ & I_{e} \\ & C_{e} \end{array} \begin{array}{c} \mu(\sigma+\mu)(v+\pi+\delta+\mu) - (1-v)\beta\sigma\Lambda \\ & \sigma(1-v)\beta(\sigma+\mu) \\ & \mu(\sigma+\mu)(v+\pi+\delta+\mu) - (1-v)\beta\sigma\Lambda \\ & \mu(\sigma+\mu)(v+\pi+\delta+\mu) - (1-v)\beta\sigma\Lambda \\ & \mu(\sigma+\mu)(v+\pi+\delta+\mu) - (1-v)\nu\beta\sigma\Lambda \\ & (1-v)\beta(\sigma+\mu)(v+\pi+\delta+\mu)(\omega+\delta+\mu) \end{array}$$

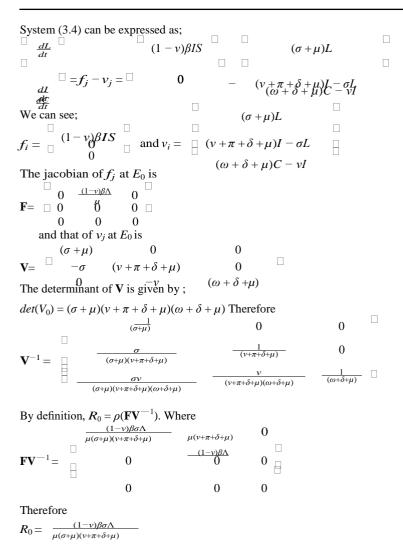
The Basic Reproduction Number (R_0)

The basic reproduction number R_0 is the average number of secondary infection cases arising from a typical primary infection case in an entirely susceptible population. Using the next generation matrix approach by Van den Driessche and Watmough [10], R_0 is given by the spectral radius $\rho(\mathbf{FV}^{-1})$ where \mathbf{F} is the Jacobian matrix of f_j at E_0 , where f_j contains terms that directly lead to new infections entering each compartment f_j and f_j is the Jacobian matrix of f_j at f_j at f_j where f_j is the rate of transfer of individuals into and out of compartment f_j . The disease compartments are f_j and f_j . The disease compartments are f_j and f_j and f_j are f_j and f_j are f_j and f_j are f_j and f_j are f_j and f_j are f_j and f_j are f_j

$$\frac{dL}{dt} = (1 - v)\beta IS - (\sigma + \mu)L$$

$$\frac{dI}{dt} = \sigma L - (v + \pi + \delta + \mu)I$$

$$\frac{dC}{dt} = vI - (\omega + \delta + \mu)C$$
(3.4)



4 Stability Analysis

In this section we will analyse local and global stability of the disease-free equilibrium and endemic equilibrium of the system (3.1).

Local Stability of the Disease Free Equilibrium Point

In this section we study local stability of the disease-free equilibrium of the system (3.1).

Theorem 1. The disease-free equilibrium of the system (3.1) is locally asymptotically stable when the reproduction number $R_0 < 1$.

Proof. Evaluating the jacobian matrix of system (3.1) at E_0 , we have;

$$J(E) = \begin{bmatrix} -\mu & 0 & -\frac{(1-\nu)\beta\Lambda}{\mu} & 0 \\ \theta & -(\sigma_{\sigma}^{+}\mu) & -\frac{(1-\psi)\beta\Lambda}{(1-\psi)\beta\Lambda} & \theta \\ 0 & 0 & \nu & -(\omega+\delta+\mu) \end{bmatrix}$$

From $J(E_0)$ the determinant is given by

$$Det J(E_0) = -\mu [-(\sigma + \mu)(\nu + \pi + \delta + \mu)(\omega + \delta + \mu) + \sigma(\omega + \delta + \mu)^{\frac{(1-\nu)\beta\Lambda}{2}}]$$
 and the trace is given by

$$tr(J(E_0) = -\mu - (\sigma + \mu) - (\nu + \pi + \delta + \mu) - (\omega + \delta + \mu)$$

Since all the parameters are positive, it can be seen that the $tr(J(E_0) < 0$ and $Det(J(E_0) > 0$ when

$$R_0 < 1$$
 i.e $(\sigma + \mu)(\nu + \pi + \delta + \mu) > \frac{(1-\nu)\beta\sigma\Lambda}{2}$

Applying Routh-Hurwitz criterion for necessary and sufficient conditions as in Enagi et al. [11], the characteristic polynomial has all roots with negative real part since $\operatorname{tr}(J(E_0)) < 0$ and $\operatorname{Det}(J(E_0)) > 0$ when $R_0 < 1$.

Thus, by Routh-Hurwitz criteria, the disease-free equilibrium is locally asymptotically stable when $R_0 < 1$

Global Stability of the Disease Free Equilibrium Point

Now we use comparison theorem as in [12], to prove the global stability of DFE

Theorem 2. The disease-free equilibrium of the system (3.1) is glocally asymptotically stable when the reproduction number $R_0 < 1$.

Proof. Using the comparison theorem in [12] we rewrite the disease compartments as

$$\frac{dL}{dt} = L = (1 - v)\beta I(S_0 - S)$$

$$= (\mathbf{F} \cdot \mathbf{V}) = I C = 0$$

$$\frac{dI}{dt} = 0$$

$$0$$

Where **F** and **V**_A are defined in section (3.2.4) Since $S \subseteq S_0 = \stackrel{\Lambda}{\longrightarrow} \forall t > 0$, it follows that

The characteristic equation is given by

$$-((\sigma+\mu)+\lambda)((\nu+\pi+\delta+\mu)+\lambda)((\omega+\delta+\mu)+\lambda)+\sigma((\omega+\delta+\mu)+\lambda)^{(1-\nu)\beta\Lambda}=0$$
 Which upon expansion gives

$$((\omega + \delta + \mu) + \lambda)[\lambda^2 + a_1\lambda + a_2] = 0 \tag{4.1}$$

where

$$a_1 = (\sigma + \mu) + (\nu + \pi + \delta + \mu)$$

$$a_2 = (\sigma + \mu)(\nu + \pi + \delta + \mu) - \frac{(1-\nu)\beta\sigma\Lambda}{2}$$

From equation (4.1) it is clear that one of the eigenvalues is $-(\omega + \delta + \mu)$ and other eigenvalues are given by the roots of

$$\lambda^2 + a_1\lambda + a_2 = 0 \tag{4.2}$$

By using Routh Hurwitz criterion, equation (4.2) has roots with negative real part if $a_1a_2 > 0$. We can see that $a_1 > 0$, this implies that for $a_1a_2 > 0$ to be satisfied, $a_2 > 0$.

Clearly $a_2 > 0$ when $R_0 < 1$ i.e $(\sigma + \mu)(\nu + \pi + \delta + \mu) > \frac{(1-\nu)\beta\sigma\Lambda}{2}$ Since we have shown that $\mathbf{F} - \mathbf{V}$ has negative eigenvalues when $R_0 < 1^{\mu}$, it means that $(L(t), I(t), C(t)) - \rightarrow (0, 0, 0)$ as t = 0. $\rightarrow \infty$. Using comparison theorem, it follows that $(L(t), I(t), C(t)) \longrightarrow (0, 0, 0)$ and $S(t) \longrightarrow^{\Lambda}_{\mu} as \ t \longrightarrow^{\infty} (S(t), L(t), I(t), C(t)) \longrightarrow E_0$ Therefore the disease free equilibrium is globally asymptotically stable when $R_0 < 1$.

Local Stability of the Endemic Equilibrium Point

Theorem 3. The endemic equilibrium point (E_e) of system (3.1) is locally asymptotically stable when $R_0 > 1$

Proof. We prove the theorem by determining if there exist negative eigenvalues when $R_0 > 1$. We start by evaluating the Jacobian matrix of system (3.1) at the endemic equilibrium point. Thus

$$J(E_e) = \begin{bmatrix} -(1-v)\beta I_e - \mu & 0 & -(1-v)\beta S_e & 0 \\ (1-v)\beta I_e & -(\sigma+\mu) & (1-v)\beta S_e & 0 \\ 0 & \sigma & -(v+\pi+\delta+\mu) & 0 \\ 0 & 0 & v & -(\omega+\delta+\mu) \end{bmatrix}$$
We see that $-(\omega+\delta+\mu)$ is one of the eigenvalues, the rest can be determined by reducing $J(E_e)$ to
$$J_1(E_e) = \begin{bmatrix} -[(1-v)\beta I_e + \mu] & 0 & -(1-v)\beta S_e \\ 0 & \sigma & -(v+\pi+\delta+\mu) \end{bmatrix}$$

$$J_{1}(E_{e}) = \begin{array}{c} \square & -\left[(1-v)\beta I_{e} + \mu\right] & 0 & -(1-v)\beta S_{e} \\ \square & (1-v)\beta I_{e} & -(\sigma+\mu) & (1-v)\beta S_{e} \\ 0 & \sigma & -(v+\pi+\delta+\mu) \end{array}$$

From $J_1(E_e)$,

 $tr(J_1(E_e)) = -((1 - v)\beta I_e + 3\mu + \sigma + v + \pi + \delta)$ $Det(J_1(E_e)) = -[(1 - v)\beta I_e + \mu] \left\{ \frac{2\mu(\sigma + \mu)(v + \pi + \delta + \mu) - 2(1 - v)\beta\sigma\Delta}{\mu} \right\}$

Clearly $Det(J_1(E_e)) > 0$ for $R_0 > 1$ i.e $(1 - v)\beta\sigma\Lambda > \mu(\sigma + \mu)(v + \pi + \delta + \mu)$

Applying Routh-Hurwitz criterion for necessary and sufficient conditions as in Enagi et al. [11], $tr(J_1(E_e))$ <0 and $Det(J_1(E_e))>0$ when $R_0>1$. Implying that all eigenvalues of $J_1(E_e)$, have negative real part when $R_0 > 1$. Thus E_e is locally asymptotically stable when $R_0 > 1$.

Global Stability of the Endemic Equilibrium Point

We study the global asymptotic stability of the endemic equilibrium using LaSalles invariance principle [13].

Theorem 4. The Endemic Equilibrium Point E_e of the system (3.1) is globally asymptotically stable if $R_0 > 1$.

Proof. We apply [13] approach to prove global stability of E_e . Consider the following Lyapunov function $G(S, L, I, C) = (S - S_e l n) + M(L - L_e l n) + P(I - I_e l n) + Q(C - C_e l n)$ The derivative of G is: S_e S_e The derivative of G is;

$$d \frac{d G}{dt} = \begin{pmatrix} 1 & -\frac{S_e}{S} \end{pmatrix} (\Delta - (1 - v)\beta IS - \mu S) + M \begin{pmatrix} 1 & -\frac{L_e}{L} \end{pmatrix} ((1 - v)\beta IS - (\sigma + \mu)L) + (\sigma L + \frac{C_e}{L}) (vI - (\omega + \delta + \mu)C)$$

$$(4.3)$$

7

At endemic equilibrium, system (3.1) become;

$$\Lambda = (1 - v)\beta I_e S_e + \mu S_e$$

$$(\sigma + \mu) = \frac{(1 - v)\beta I_e S_e}{L_e}$$

$$(v + \pi + \delta + \mu) = \frac{\sigma L_e}{I_e}$$

$$(\omega + \delta + \mu) = \frac{v I_e}{C_e}$$
(4.4)

Substituting (4.4)in (4.3), we get;

$$\frac{dG}{dt} = 1 - \frac{S_e}{S} [(1 - v)\beta I_e S_e + \mu S_e - (1 - v)\beta I S - \mu S]
+ M 1 - \frac{I_e}{L} [(1 - v)\beta I S - \frac{(1 - v)\beta I_e S_e L}{L_e}]
+ P 1 - \frac{I_e}{I} [\sigma L - \frac{\sigma L_e I}{I_e}] + Q 1 - \frac{C_e}{C} [vI - \frac{v I_e C}{C_e}]$$
(4.5)

Equation (4.5) can be written as

where $w = \frac{S}{S_e}$, $x = \frac{L}{L_e}$, $y = \frac{I}{I_e}$ and $z = \frac{C}{L_e}$ Further, equation (4.6) can be written as $\frac{dG}{dt} = -\mu \frac{(S - Se)^2}{S} + f(w, x, y, z)$

Where

 $+M(1 v)\vec{\beta}IS (1+wy-x)\vec{\beta}IS (1+$ we obtain,

$$M (1 - v)\beta I_e S_e = 0$$
$$-P\sigma L_e = 0$$
$$-Qv I_e = 0$$

let M=1, solving for P and Q, we have;

$$P = \frac{(1-v)\beta I_e S_e}{\sigma L_e}, \ Q = \frac{(1-v)\beta S_e}{v}$$

Substituting for M, P and Q in equation (4.7), we obtain
$$f(w, x, y, z) = 1 - \frac{1}{w} (1 - v)\beta I_{e}S_{e} - 1 - \frac{1}{w} (1 - v)wy\beta I_{e}S_{e}S_{e} - 1 - \frac{1}{w} (1 - v)wy\beta I_{e}S_{e}S_{e} - 1 - \frac{1}{w} (1 - v)wy\beta I_{e}S_{e}S_{e} - 1 - \frac{1}{w} (1 - v)wy\beta I_{e}S_{e}S_{e}S_{e}$$

Which upon simplification gives
$$f(w, x, y, z) = (1 - v)\beta I_e S_e (4 + y - \frac{y}{z} - z - \frac{x}{y} - \frac{1}{w} - \frac{wy}{x})$$
Using geometric mean inequality, we obtain

Using geometric mean inequality, we obtain

$$\frac{y}{z} + z + \frac{x}{y} + \frac{1}{y} + \frac{wy}{w} - xy > 4$$

Thus
$$f(w, x, y, z) \le 0$$
. Hence $\frac{dG}{\leq 0} = 0$ in Ω The equality $\frac{dG}{dz} = 0$

Thus $f(w, x, y, z) \le 0$. Hence $\frac{dG}{dt} \le 0$ in Ω The equality $\frac{dG}{dt} = 0$ iff w=x=y=z=1 and $S = S_e$, $L = L_e$, $L = C_e$. Therefore, using LaSalle's invariance principle, the endemic equilibrium

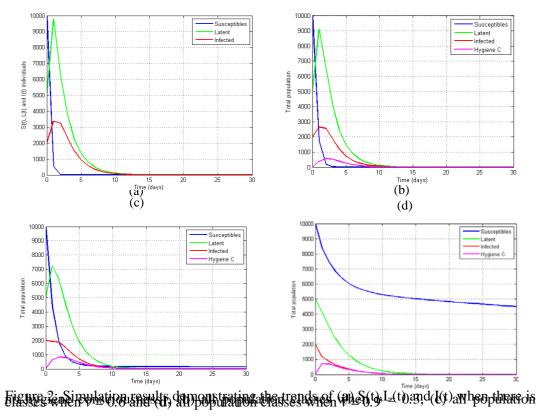
point of the system (3.1) is globally asymptotically stable.

5 **Numerical Simulation**

We carry out numerical simulations of the model (3.1), using MATLAB ode45 solver. The parameter values used are presented in Table 1. Simulation results are presented in Figures (a) to (d).

Parameter symbolished: 1: Parameterevalues of the model Source		
Λ	$8.7 \times 10^{-3} / \text{day}$	CIA (2014)
μ	$7.0 \times 10^{-3} / day$	CIA (2014)

Figure (a) is graphical representation indicating the trends of population classes in the absence of hygiene consciousness. It can be seen that susceptibles fall to zero because they are infected by infectious individuals whereas latently Infacted and Infection 1800 is the properties of latently Infected and Infectious individuals 81.7 eas \$0.70 / Heaviain point and then Estimatering, implying that v = 0.3 is not effective in controlling tuberculosis. From figure (c) it can be observed that x = 0.6, number of latently Infected and Infectious individuals increases to divertain point and then start declining but the peaks 0.6, number of of the two graphs in Figure (c) are a bit lower as compared to those in Figure (b) Figure (d) shows that for v = 0.9, the numbers of all population classes fall off and eventually converge to zero. This suggest that effective hygiene consciousness is sufficient in controlling TB epidemic. The infectious individuals can observe hygiene through cough etiquette, ensuring that windows are open while in congested area and not spitting everywhere. Also, hygiene can be observed through proper cleaning and sterilization of hospital equipments.



6 Conclusion

A mathematical model of TB which captures the effects of hygiene consciousness is formulated. We proved the positivity of the solutions and determined the equilibrium points. We carried out stability analysis of the model and it showed that the disease-free equilibrium point is both locally and globally asymptotically stable given that $R_0 < 1$, implying that TB can be eliminated from the population. Whereas the endemic equilibrium point is both locally and globally asymptotically stable given that $R_0 > 1$, implying that TB will eventually be able to invade the population but its transmission levels can be kept at manageable levels in the presence of hygiene consciousness. The obtained numerical simulations results suggest that in the absence of hygiene consciousness TB disease could invade the population while in the presence of effective hygiene consciousness, the development of TB disease is slowed down. Hence this implies that there is a need to embrace hygiene consciousness as a control measure of TB.

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Competing Interest

We declare that no competing interests exist.

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