

A model of Partial Differential Equations for HIV propagation in lymph nodes

by: E.B.S. Marinho, F.S. Bacelar, R.F.S Andrade (2012)
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Mario González Galindo

ESILV

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Introduction

- Modelling of the interaction between HIV and $CD4^+$ T cells, which are the main target of HIV.
- Infection seen inside the lymph nodes, which is the previous location of IS cells before going to the bloodstream.
- Addition of space dependence to previous models considering only time based solutions.

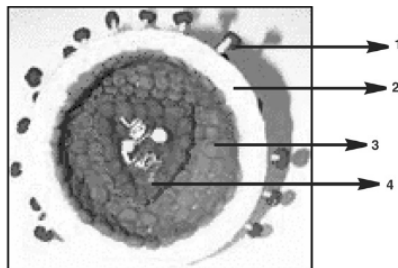


Figure: HIV Structure. 1. Envelope composed of proteins gp120 and gp41; 2. Internal capsule composed of protein p17; 3. Membrane composed of protein p24 and 4. Genetic material composed of RNA and proteins. [2]

Methods: Partial Differential Equation System

Model dependant on time t and space

$\vec{r} = \langle x\hat{x}, y\hat{y} \rangle$; $\hat{x}, \hat{y} \in [0, L], [0, L]$

$$\frac{\partial H_1}{\partial t} = -k_5 H_1 A^p - k_6 H_1 B^n + k_9 H_1 \nabla^2 A,$$

$$\frac{\partial H_2}{\partial t} = k_3 D - k_5 H_2 A^q - k_6 H_2 B^n + k_9 H_2 \nabla^2 A,$$

$$\frac{\partial A}{\partial t} = -k_1 A(t - \tau) + k_4 D + k_5 (H_1 A^p + H_2 A^q) + k_6 (H_1 + H_2) B^n - k_9 (H_1 + H_2) \nabla^2 A,$$

$$\frac{\partial B}{\partial t} = k_1 A(t - \tau) - k_2 B,$$

$$\frac{\partial D}{\partial t} = -k_3 D - k_4 D + k_2 B$$

Figure: Partial Differential Equation System describing the HIV infection in the lymph nodes. [3]

Model considering 4 variables:

- $H_1(t, \vec{r})$: Healthy cells already in the region
- $H_2(t, \vec{r})$: Healthy cells newly differentiated
- $A(t, \vec{r})$: Newly infected cells
- $B(t, \vec{r})$: Old infected cells
- $D(t, \vec{r})$: Dead cells

parameter	meaning	value
k_1	Transition rate from A to B	0.163
k_2	Transition rate from B to D	0.228
k_3	Replenishment rate from D to H_2	0.650
k_4	Replenishment rate from D to A	3.25×10^{-5}
k_5	Infection rate from H_1 and H_2 to A	0.650
K_6	Infection rate from H_1 and H_2 to B	0.169
K_9	Diffusion rate of A cells	$\in (0.2, 8) \times 10^{-3}$
p	Infection order from H_1 to A	1
q	Infection order from H_2 to A	1.15
n	Infection order from H_1 and H_2 to B	4
τ	Time delay from A to B	4 weeks

Table: The parameters taken after the *Dynamics of HIV infection* literature, presented with their functions and meanings along the model. [1]

Results: Temporal evolution

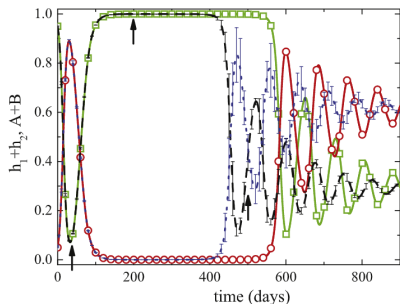


Figure: Time evolution of $(H_1 + H_2)$ and $(A + B)$ predicted by the system of **Figure 2** in a square lattice of tissue of area L^2 . The squares $(H_1 + H_2)$ and circles $(A + B)$ are the solution for uniform initial conditions $(H_1, H_2, A, B, D) = (0.95, 0, 0.05, 0, 0)$; $(\hat{H}_1, \hat{H}_2, \hat{A}, \hat{B}, \hat{D}) = (0, 1, 0, 0)$. The dashed $(H_1 + H_2)$ and dotted $(A + B)$ lines are the solution for non uniform initial conditions; $A_0 = 0.06$ in a small square of the lattice [3].

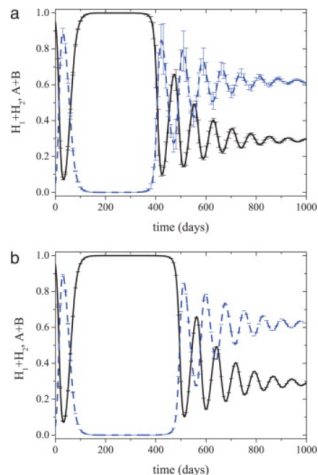


Figure: Time evolution for the fluctuation of the parameter K_9 with values of $K_9 = 2 \times 10^{-4}$ (a) and $k_9 = 2 \times 10^{-3}$ (b), with solid and dashed lines representing $(H_1 + H_2)$ and $A + B$ respectively [3].

Results: Spatial Evolution

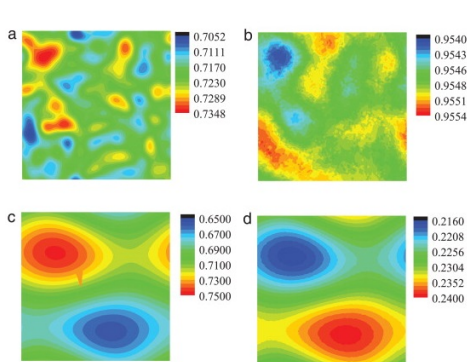


Figure: Four spatial patterns for the fraction of $H_1 + H_2$ at $t_1 = 10$ (a), $t_2 = 100$ (b), $t_3 = 400$ (c) and $t_4 = 600$ days (d) resulting from random non homogeneous initial conditions. The same parameters as the **Figure 3** are considered [3].

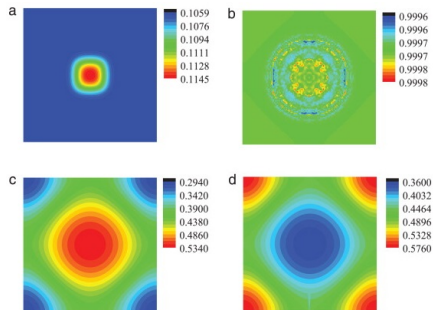


Figure: Representation of the dynamic concentration of healthy cells ($H_1 + H_2$) in a square lattice presented with non-homogeneous initial concentrations at $t_1 = 40$ days (a), $t_2 = 200$ days (b), $t_3 = 500$ days (c), using a diffusion coefficient $k_9 = 2 \times 10^{-4}$. Panel (d) shows concentration of infected cells ($A + B$) at $t = t_3$. [3]

Conclusions

- The time and space evolution of the infection is widely influenced by the diffusion factors and initial distributions over the evaluation area.
- Identification of the influence of non linear diffusion terms to describe the movement of the infected cells across a given area of the lymph nodes.
- Identification of fluctuations and changes in behavior with given changes in diffusion parameters and the introduction of non uniform initial conditions over the evaluation area.



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