Intensional infect proportion of newborn, with disease induced mortality rate

May 28, 2018

1 Motivation

Previous analysis showed no obvious advantage for intentional infection. But those are the cases where we ignored disease induced mortality. In reality, if we are taking smallpox for example, past researches have determined the mortality rate to be 30 percent for normally infected cases, but only 1 percent for variolated cases. Thus, it is possible that intentional infection has a positive effect on disease control.

2 Introduction

Again, we consider two intentional infect strategies. One is to intentional infect newborns and the other is to intentional infect susceptible. In this document, we discuss the first strategy only.

3 System of differential equations

Since we have to consider disease induced mortality rate, we need to adjust our model by adding extra terms representing mortality rate.

The following assumptions are used:

- Birth and natural death rate are the same.
- The latent period is short enough to be ignored.
- All susceptible individuals are equally likely to be infected, and all infected individuals are equally infectious.

$$
\begin{aligned}
\frac{\mathrm{d}S}{\mathrm{d}t} &= \mu(1-p) - \beta S(V+I) - \mu S\,, \\
\frac{\mathrm{d}V}{\mathrm{d}t} &= \beta SV + \mu p - \gamma V - \mu V\,, \\
\frac{\mathrm{d}I}{\mathrm{d}t} &= \beta SI - \gamma I - \mu I\,, \\
\frac{\mathrm{d}M}{\mathrm{d}t} &= p_V \gamma V + p_I \gamma I\,, \\
\frac{\mathrm{d}R}{\mathrm{d}t} &= (1-p_V)\gamma V + (1-p_I)\gamma I - \mu R\,,
\end{aligned}
\tag{1}
$$

Here, β is the transmission rate, γ is the recovery rate, μ is the *per capita* rate of birth and death, p is the proportion of newborns that are intentionally infected.

We non-dimensionalize Equation 1 by scaling time, by

$$\tau = (\gamma + \mu)t\,, \tag{2}$$

As the result, we obtain, ⟨***David:*** *Do not use hard-coded specific values of case fatality proportions. Use symbols.* p_V, p_I *for "probability of mortality in* V *or* I *classes respectively* ⟩

$$\frac{\mathrm{d}S}{\mathrm{d}\tau} = \epsilon(1-p) - \mathcal{R}_0 S(V+I) - \epsilon S\,, \tag{3a}$$

$$\frac{\mathrm{d}V}{\mathrm{d}\tau} = \mathcal{R}_0 SV + \epsilon p - V\,, \tag{3b}$$

$$\frac{\mathrm{d}I}{\mathrm{d}\tau} = \mathcal{R}_0 SI - I\,, \tag{3c}$$

$$\frac{\mathrm{d}M}{\mathrm{d}\tau} = p_V(1-\epsilon)V + p_I(1-\epsilon)I\,, \tag{3d}$$

$$\frac{\mathrm{d}R}{\mathrm{d}\tau} = (1-p_V)(1-\epsilon)V + (1-p_I)(1-\epsilon)I - \epsilon R\,, \tag{3e}$$

where $\epsilon = \frac{\mu}{\gamma+\mu}$, $\mathcal{R}_0 = \frac{\beta}{\gamma+\mu}$.

4 Equilibria

To solve for all equilibria, we let equations Equation 3a, Equation 3b and Equation 3c equal to 0, we solve solutions.

First by letting Equation 3c equal to 0, we have either $S = \frac{1}{\mathcal{R}_0}$, $I = 0$ or both. For the case where $S = \frac{1}{\mathcal{R}_0}$, Equation 3b returns,

$$\frac{\mathrm{d}V}{\mathrm{d}\tau} = \epsilon p = 0\,, \tag{4}$$

Which has no solution if $p \neq 0$, and since we consider various possibilities where $p \neq 0$, we conclude that $S \neq \frac{1}{\mathcal{R}_0}$. Therefore, $I = 0$.

Equipped with the above condition, we now solve the other two equations and the only solution we acquire is

$$\hat{S} = \frac{1}{\mathcal{R}_0} - \frac{2p}{(\mathcal{R}_0 - 1) + \sqrt{(\mathcal{R}_0 - 1)^2 + 4\mathcal{R}_0 p}}\,, \tag{5a}$$

$$\hat{V} = \frac{\epsilon(\mathcal{R}_0 - 1) + \epsilon\sqrt{(\mathcal{R}_0 - 1)^2 + 4\mathcal{R}_0 p}}{2\mathcal{R}_0}\,, \tag{5b}$$

$$\hat{I} = 0\,, \tag{5c}$$

At this equilibrium, since the infected population is non-zero, this is not a disease free equilibrium. It follows that the only equilibrium we found is an endemic equilibrium, and disease free equilibrium does not exist for this model.

5 Stability of Endemic Equilibrium

Stability analysis rely on Jacobian Matrix,

$$\mathcal{J} = \begin{bmatrix} -\mathcal{R}_0(V + I) - \epsilon & -\mathcal{R}_0 S & -\mathcal{R}_0 S \\ \mathcal{R}_0 V & \mathcal{R}_0 S - 1 & 0 \\ \mathcal{R}_0 I & 0 & \mathcal{R}_0 S - 1 \end{bmatrix}. \tag{6}$$

Eigenvalues of Jacobian are given as follow,

$$\lambda_1 = -1 + \mathcal{R}_0 S \tag{7a}$$

$$\lambda_2 = \frac{-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V - \sqrt{(-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V)^2 - 4(\mathcal{R}_0 + \epsilon - \mathcal{R}_0 S \epsilon)}}{2} \tag{7b}$$

$$\lambda_3 = \frac{-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V + \sqrt{(-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V)^2 - 4(\mathcal{R}_0 + \epsilon - \mathcal{R}_0 S \epsilon)}}{2} \tag{7c}$$

By using Equation 5a and Equation 7a, we obtain

$$-1 + \mathcal{R}_0 S = -\frac{2p\mathcal{R}_0}{(\mathcal{R}_0 - 1) + \sqrt{(\mathcal{R}_0 - 1)^2 + 4\mathcal{R}_0 p}} < 0 \tag{8}$$

Therefore, ⟨***David:*** *eigenvalues must be written in $a+ib$ form, where a, b are real.* ⟩ ⟨***David:*** *You have not calculated the real parts of λ_2 and λ_3.* ⟩

$$\Re(\lambda_1) = -1 + \mathcal{R}_0 S < 0\,, \tag{9}$$

To determine the real part of λ_2 and λ_3, we need to determine the sign of the quantity under the square root.

By using Equation 5a again, we have

$$\mathcal{R}_0 S \epsilon < \epsilon\,, \tag{10}$$

Therefore,

$$(\mathcal{R}_0 + \epsilon - \mathcal{R}_0 S \epsilon) > \mathcal{R}_0 > 0\,, \tag{11}$$

which means, if the sign of the quantity under the square root is positive, we necessarily have

$$\sqrt{(-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V)^2 - 4(\mathcal{R}_0 + \epsilon - \mathcal{R}_0 S \epsilon)} < |(-1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V)| \tag{12}$$

Therefore, $\Re(\lambda_2) < \Re(\lambda_3) < 0$.

Certainly, if the sign of the quantity under the square root is negative,

$$\Re(\lambda_2) = \Re(\lambda_3) = -1 + \mathcal{R}_0 S - \epsilon - \mathcal{R}_0 V < 0 \tag{13}$$

We are able to conclude that EE is stable.

6 Disease Free Equilibrium

As mentioned above in section 4, disease free equilibrium does not exist for this model.

7 Mortality rate at Endemic equilibrium

When performing epidemic analysis, it is important to observe the mortality rate of the population, since this parameter is crucial to the severity of this disease. Here, we emphasize the mortality rate at EE.

By substituting the corresponding values at EE into equation (3d), we obtain,

$$\frac{\mathrm{d}M}{\mathrm{d}\tau} = p_V(1-\epsilon)V = \frac{p_V(1-\epsilon)\epsilon(\mathcal{R}_0-1) + p_V(1-\epsilon)\epsilon\sqrt{(\mathcal{R}_0-1)^2+4\mathcal{R}_0 p}}{2\mathcal{R}_0}, \tag{14}$$

Equation 14 reveals 3 important points.

First, the mortality rate at the EE does increase as the proportion p of individuals who are intentionally infected is increased. Nevertheless, the total mortality over any period will be less with intentional infection if the mortality rate form intentional infection is much lower than the disease-induced mortality rate from natural infection. Consequently, it may be beneficial to have high p during an initial outbreak, but once an equilibrium is reached, it is likely to be better to avoid intentional infection.

Second, as expected, the probability p_V of mortality due to intentional infection plays a major role in the mortality rate at the EE. Meaning intentional infection is unwise if p_V is too high.

Third, mortality rate $\mathrm{d}M/\mathrm{d}\tau$ also increases as $\mathcal{R}_0$ increases.

More specifically, we now consider the case of smallpox, i.e., the parameter values in Table 1.

Table 1: Model parameters and smallpox values.

Symbol	Meaning	Value
μ	Natural *per capita* death rate	$\frac{1}{50*365}$ per day
γ	Recovery rate	$\frac{1}{22}$ per day
$\mathcal{R}_0$	Basic reproductive number	4.5

Therefore, we can calculate $\epsilon = \frac{\mu}{\mu+\gamma} = 0.0012$

$$\frac{\mathrm{d}M}{\mathrm{d}\tau} = 0.00111111(0.00420902 + \frac{100375\sqrt{12.25+18p}}{83466496}), \tag{15}$$

⟨***David:*** *write that as* $a + b\sqrt{c+dp}$*, where* a, b, c, d *are number given to three significant figures.* ⟩ So we plot $\frac{\mathrm{d}M}{\mathrm{d}\tau}$ as a function of p,

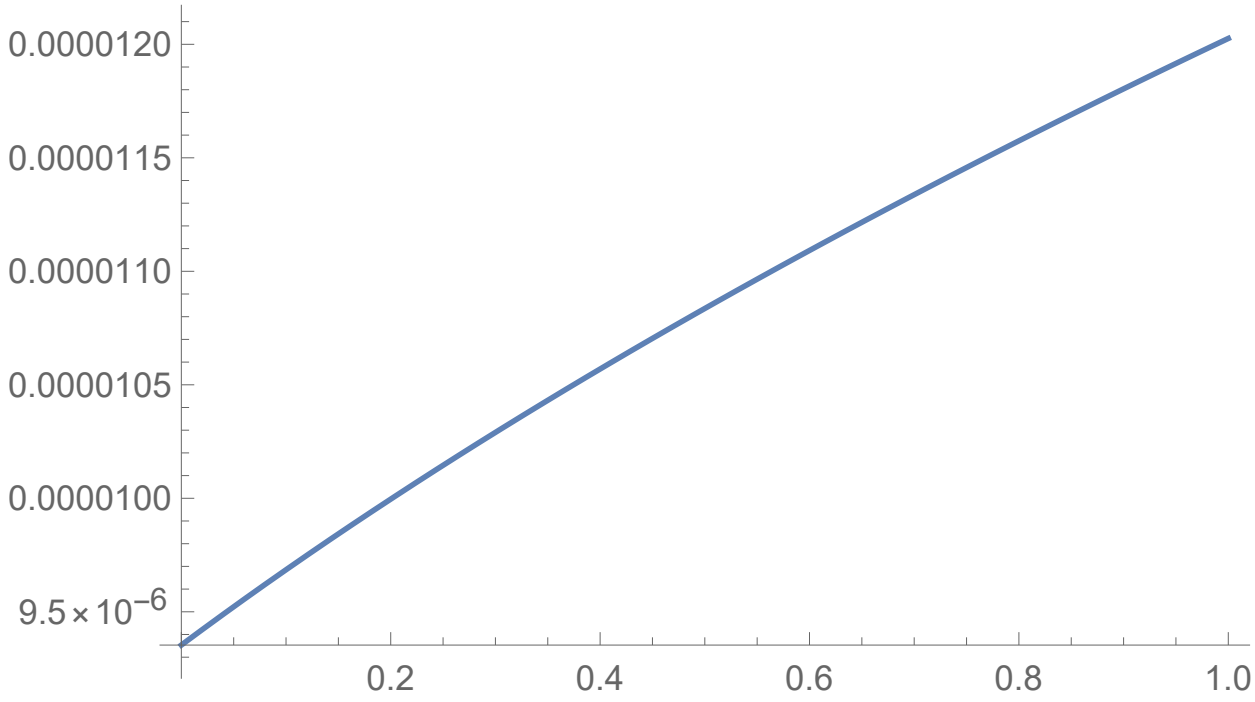

Figure 1: $\frac{\mathrm{d}M}{\mathrm{d}\tau}$ at EE as a function of p.

A major observation from this plot is, the magnitude of mortality rate at endemic equilibrium is far less than natural death rate. That is,

$$\left.\frac{\mathrm{d}M}{\mathrm{d}t}\right|_{\mathrm{EE}} \ll \epsilon \tag{16}$$

Consequently, once an equilbirium is reached, disease induced mortality will be negligible. In Figure 3 we demonstrate the total mortality counts as a function of time, using parameters from Table 1.

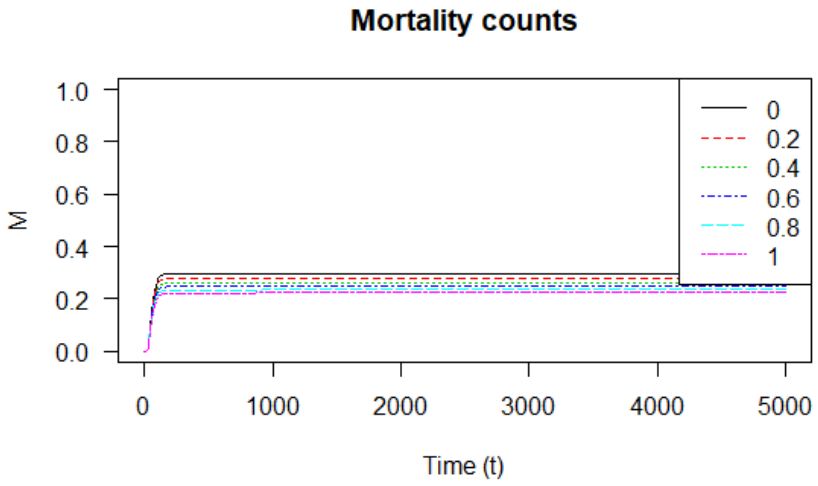

Figure 2: $\frac{\mathrm{d}M}{\mathrm{d}\tau}$ at EE as a function of p.

Another plot to show the disadvantages of having a high proportion of intentional infection. This time, the variolation mortality is 20 percent instead of 1 percent.

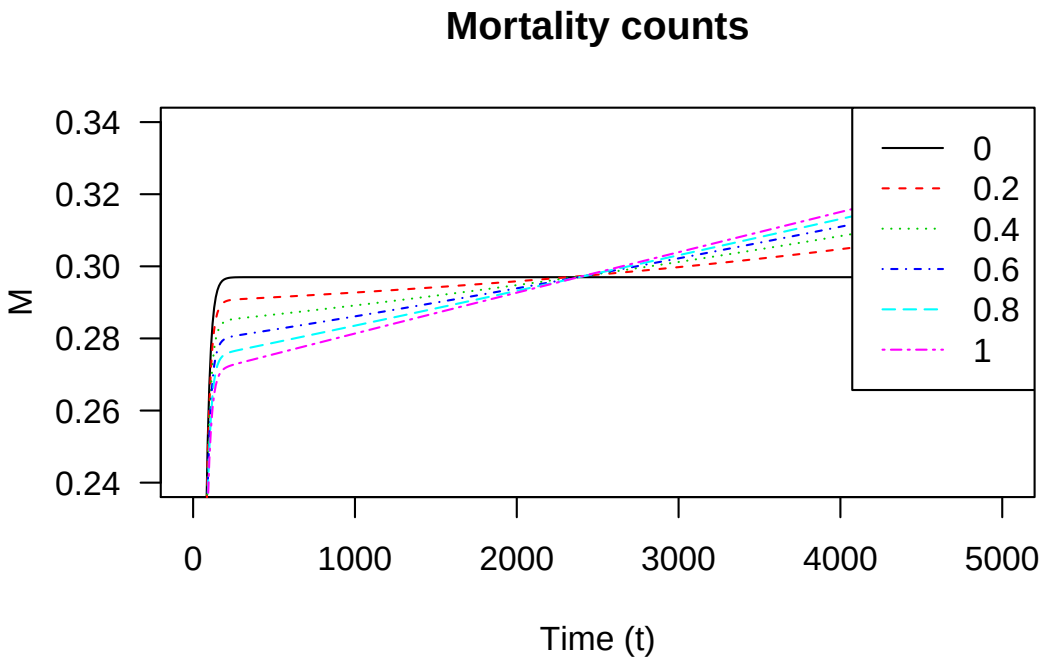

Figure 3: $\frac{\mathrm{d}M}{\mathrm{d}\tau}$ at EE as a function of p.