Optimal control on a vaccine metapopulation SIR model

July 29, 2022

Abstract

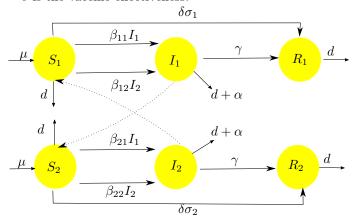
1 Introduction

We consider the following SIR model of metapopulations with migrations:

$$\begin{cases}
\frac{dS_{i}}{dt} = \mu N_{i} - S_{i}(t) \sum_{j=1}^{n} \beta_{ij} I_{j}(t) - \delta \sigma_{i} S_{i}(t) - dS_{i}(t) \\
\frac{dI_{i}}{dt} = S_{i}(t) \sum_{j=1}^{n} \beta_{ij} I_{j}(t) - (d + \alpha + \gamma) I_{i}(t) \\
\frac{dR_{i}}{dt} = \gamma I_{i}(t) + \delta \sigma_{i} S_{i}(t) - dR_{i}(t) \\
S(0) = S_{i0}; \quad I_{i}(0) = I_{i0}; \quad R_{i}(0) = R_{i0}
\end{cases} \tag{1}$$

where

- β_{ij} is the transmission rate per capita per time unit of individuals of the class i with the class j.
- μ is the birth rate.
- σ_i is the vaccination rate at the node i (control variable).
- γ is the recovery rate.
- \bullet d is the natural death rate.
- α is the death rate from the disease (case fatality).
- δ is the vaccine effectiveness.



Experimento método directo, minimiz

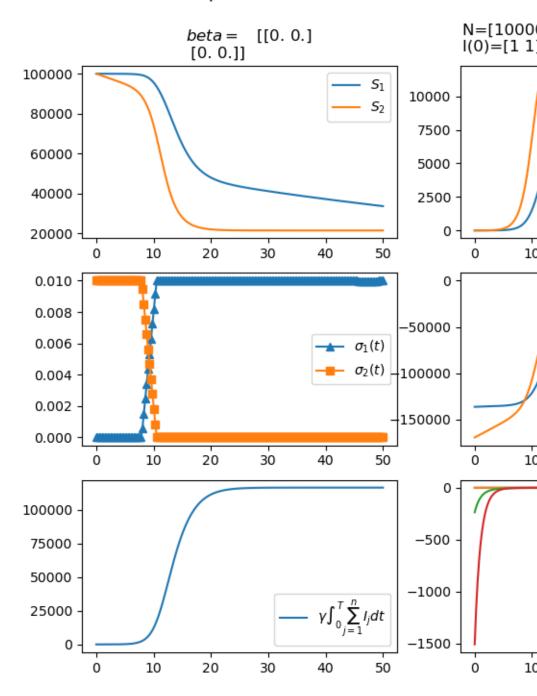


Figure 1: Caption

We suppose that $\sigma_T:[0,T]\to\mathbb{R}_+$ is a non-negative function. We consider the admissible control set

$$U_{ad} = \{ \sigma = (\sigma_1, \dots, \sigma_n) : \sigma_i \text{ measurable }, \sigma_i \geqslant 0, \sigma_1 + \dots + \sigma_n = \sigma_T(t) \}$$

Nota 26/07/2022:

 $\sigma_i = \#$ Vacunados nodo $i \times$ unidad de tiempo \times suceptible del nodo i.

La cantidad total de vacunas suministradas en el nodo i es $\sigma_i S_i$. Estamos considerando igual de admisibles distribuciones de vacunas que en total pueden resultar en cantidades de vacunados muy diferentes. Parece más razonable que el conjunto admisible sea algo del tipo:

$$\sigma_1 S_1 + \dots + \sigma_n S_n \leq \sigma_T(t)$$

Ahora $\sigma_T(t)$ representan cantidad de vacunas por unidad de tiempo en la población total. Quizás que la cantidad de vacunados no se exprese como $\sigma_i S_i$. Discutir un poco esto.

The objective function given by

$$J(\sigma) = \int_0^T \sum_{i=1}^n I_i(t)dt$$

we formulate the optimal control problem

find
$$\sigma^* \in U_{ad}$$
 such that $J(\sigma^*) = \min_{\sigma \in U_{ad}} J(\sigma)$ (2)

2 Some observations on dynamics of vaccine metapopulation model

We assume $\mu = d = \alpha = 0$ and $\delta = 1$. In this situation $N_i' := (S_i + I_i + R_i)' = 0$, i.e. the total population of node i remain constant. Cosequently $R_i = N - S_i - I_i$ and we can drop the equation for R_i of the system. Therefore, we can study the SIR metapopulation model with vaccination (SIRmv) system.

$$\begin{cases} S_i' = -S_i(t) \sum_{j=1}^n \beta_{ij} I_j(t) - \sigma_i(t) S_i(t) & (SIRmv1) \\ I_i' = S_i(t) \sum_{j=1}^n \beta_{ij} I_j(t) - \gamma I_i(t) & (SIRmv2) \end{cases}$$

We note that the system of equations (SIRmv1), (SIRmv1) possibly has a distinuous right hand side. I this case, I this case, following to [Filippoy, 1988]

continuous right hand side. I this case, I this case, following to [Filippov, 1988]. Poner el significado de solución

We will use the following well known and elementary formula for the solution of a linear scalar eqution of first order x'(t) + p(t)x(t) = q(t):

$$x(t) = e^{-\int_0^t p(s)ds} \left\{ x(0) + \int_0^t e^{\int_0^s p(r)dr} q(s)ds \right\}$$
 (3)

From (SIRmv1) and (3) we obtain that

$$S_i(t) = S_i(0) \exp\left(-\int_0^t \sum_{j=1}^n \beta_{ij} I_j(s) + \sigma(s) ds\right)$$

The following conjecture establish that if is a irreducible matrix (see [Meyer, 2000]) the epidemic is transmitted from any node to the rest with infinite speed.

Proposition 2.1. Suppose that β is a irreducible matrix. If there exists i such that $I_i(0) > 0$ then for every j and t > 0 we have $I_j(t) > 0$.

Proof. Agregar
$$+$$

Lemma 2.2. Let $h \in L^{\infty}([0, +\infty))$ be a function. Suppose that there exists $\lim_{t\to\infty} h(t)$ and denote it by h_{∞} . Then

$$h_{\infty} = \lim_{t \to \infty} \gamma \int_{0}^{t} h(s)e^{\gamma(s-t)}ds. \tag{4}$$

Proof. Given $\varepsilon > 0$ there exists $t_0 \geq 0$ such that

$$t \ge t_0 \Rightarrow |h(t) - h_{\infty}| < \varepsilon$$
.

Using the identity

$$\gamma \int_0^t e^{\gamma(s-t)} ds = 1 - e^{-\gamma t},$$

we can deduce

$$\begin{split} \left|h_{\infty} - \gamma \int_0^t h(s)e^{\gamma(s-t)}ds\right| &= \left|e^{-\gamma t}h_{\infty} + \gamma \int_0^t \left(h_{\infty} - h(s)\right)e^{\gamma(s-t)}ds\right| \\ &\leq e^{-\gamma t}|h_{\infty}| + \gamma \int_0^{t_0} |h_{\infty} - h(s)|e^{\gamma(s-t)}ds + \gamma \int_{t_0}^t |h_{\infty} - h(s)|e^{\gamma(s-t)}ds \\ &\qquad \qquad e^{-\gamma t}|h_{\infty}| + 2\gamma ||h||_{L^{\infty}}e^{\gamma(t_0-t)}t_0 + \varepsilon. \end{split}$$

This inequality implies the result in lemma.

Remark 1. Of course, the existence of the limit on the right hand side in equation (4) does not guarantee the existence of limit of h(t) for $t \to \infty$. An example of this fact is obtained as follow. Let

$$h = \sum_{n=0}^{\infty} \mathbb{1}_{[n,n+\frac{1}{2}]},$$

where $\mathbbm{1}_A$ denotes the characteristic function of the set A. Then $h(s)+h(s-\frac{1}{2})=1$, for $s\in[0,+\infty]$. We define $\varphi(s)=h(e^s-1)$. Then

$$1 = \lim_{t \to \infty} \gamma \int_0^t \left(\varphi(s) + \varphi\left(s - \frac{1}{2}\right) \right) e^{\gamma(s-t)} ds$$

$$= \lim_{t \to \infty} \gamma \int_0^t \varphi(s) e^{\gamma(s-t)} ds + \gamma \int_0^t \varphi\left(s - \frac{1}{2}\right) e^{\gamma(s-t)} ds \quad (5)$$

On the other hand

$$\gamma \int_{0}^{t} \varphi\left(s - \frac{1}{2}\right) e^{\gamma(s-t)} ds = \gamma e^{\frac{\gamma}{2}} \int_{0}^{t - \frac{1}{2}} \varphi\left(r\right) e^{\gamma(r-t)} dr$$
$$= \gamma e^{\frac{\gamma}{2}} \int_{0}^{t} \varphi\left(r\right) e^{\gamma(r-t)} dr - \gamma e^{\frac{\gamma}{2}} \int_{t - \frac{1}{2}}^{t} \varphi\left(r\right) e^{\gamma(r-t)} dr \quad (6)$$

In my understanding

$$\lim_{t\to\infty}\int_{t-\frac{1}{2}}^{t}\varphi\left(r\right)e^{\gamma\left(r-t\right)}dr=\frac{1}{2\gamma}(1-e^{-\gamma/2}).$$

Taking account of (5), (6) we infer that

$$\lim_{t \to \infty} \gamma \int_0^t \varphi(s) e^{\gamma(s-t)} ds = \frac{e^{\gamma/2}}{2(1 + e^{\gamma/2})}$$

The following proposition to expresses the fact that epidemic is extinguished when $t \to \infty$ and, in certain sense, the total quantity of applied vaccines $\sigma_i S_i$ goes to zero when $t \to \infty$.

Proposition 2.3.

$$\lim_{t \to \infty} \gamma \int_0^t \sigma_i(s) S_i(s) e^{\gamma(s-t)} ds = I_i(\infty) = 0.$$

Proof. Adding equations (SIRmv1) and (SIRmv2) we obtain

$$(S_i + I_i)' = -\sigma_i S_i - \gamma I_i < 0.$$

Therefore S_i+I_i is a monotone non increasing function. Hence $\lim_{t\to\infty}(S_i+I_i)$ there exists. From (SIRmv1) the same considerations are true for function S_i . Consequently there exists $\lim_{t\to\infty}S_i(t)=:S_i(\infty)$. We deduce that there exists $\lim_{t\to\infty}I_i(t)=:I_i(\infty)$. If $I(\infty)>0$, we could choose t_0 large enough for that $t\geq t_0$ implies $I_i(t)>I_i(\infty)/2=:a>0$. Then $(S_i(t)+I_i(t))'\leq -\gamma I_i(t)\leq -\gamma a$. This inequality implies that $S_i(t)+I_i(t)\to -\infty$, when $t\to\infty$, which is a contradiction. Consequently $I_i(\infty)=0$.

From (SIRmv2) and (3) we obtain that

$$\begin{split} I_{i}(t) &= e^{-\gamma t} \left\{ I_{i}(0) + \int_{0}^{t} e^{\gamma s} S_{i}(s) \sum_{j=1}^{n} \beta_{ij} I_{j}(s) ds \right\} \\ &= e^{-\gamma t} \left\{ I_{i}(0) - \int_{0}^{t} e^{\gamma s} \left[S_{i}'(s) + \sigma_{i}(s) S_{i} \right] ds \right\} \\ &= e^{-\gamma t} \left\{ I_{i}(0) - \int_{0}^{t} e^{\gamma s} \sigma_{i}(s) S_{i}(s) ds - e^{\gamma t} S_{i}(t) + S_{i}(0) - \gamma \int_{0}^{t} e^{\gamma s} S_{i}(s) ds \right\} \\ &= e^{-\gamma t} \left(S_{i}(0) + I_{i}(0) \right) - S_{i}(t) + \gamma \int_{0}^{t} e^{\gamma (s-t)} S_{i}(s) ds - \int_{0}^{t} e^{\gamma s} \sigma_{i}(s) S_{i}(s) ds \right\} \end{split}$$

Taking the limit for $t \to \infty$ in previous identities and using Lemma 2.2 \Box

Conjecture 2.4.

$$\lim_{t \to \infty} \sigma_i(s) S_i(s) = 0.$$

3 Existence minimizers

In this Section, we prove that the optimal control problem (2) has a solution. That is, we prove that the hypothesis of the Filippov-Cesari Theorem are satisfied (see [Seierstad and Sydsæter, 1987]). In what follows, we will use the following notation

$$x = (S_1, \dots, S_n, I_1, \dots, I_n, R_1, \dots, R_n)$$

$$f_0(x_i, \sigma_i, t) = f_0(S_1, \dots, S_1, I_1, \dots, I_n, R_1, \dots, R_n; \sigma_1, \dots, \sigma_n; t) = \sum_{i=1}^n I_i(t).$$

For $i = 1, 2, \dots, n$, we denote by

$$f_i = (f_{0i}, f_{2i}, f_{3i}), \quad f = (f_1, \dots, f_n)$$

with

$$f_{1i} = \mu N_i - S_i(t) \sum_{j=1}^{n} \beta_i I_j(t) - \delta \sigma_i S_i(t) - dS_i(t)$$

$$f_{2i} = S_i(t) \sum_{j=1}^{n} \beta_i I_j(t) - (d + \alpha + \gamma) I_i(t)$$

$$f_{3i} = \gamma I_i(t) + \delta \sigma_i S_i(t) - dR_i(t)$$

and we define

$$N(x, U_{ad}, t) = \{(f_0 + \gamma, f) : \gamma \leq 0, \sigma \in U_{ad}\}.$$

Now, we will are in conditions to prove the following result.

Theorem 3.1. The optimal control problem (2) has a solution $\sigma^* \in U_{ad}$.

Proof. We will see that $N(x, U_{ad}, t)$ is a convex set, for every (x, t).

Let $(a_1,b_1),(a_2,b_2)\in N(x,U_{ad},t)$ be, then there exist $\gamma_1,r_2\leq 0$ and $\sigma_1,\sigma_2\in U_{ad}$ such that

$$(f_0(x, y_1, t) + \gamma_1; f(x, \sigma_2, t)) = (a_1, b_1)$$

and

$$(f_0(x, \sigma_2, t) + \gamma_2, f(x, \sigma_2, t)) = (a_2, b_2)$$

then

$$\lambda(a_1, b_1) + (1 - \lambda)(a_2, b_2) = (\lambda a_1 + (1 - \lambda)a_2, \lambda b_1 + (1 - \lambda)b_2)$$

$$= (\lambda (f_0(x, \sigma_1, t) + \gamma_1) + (1 - \lambda) (f_0(x_1, \sigma_2, t) + r_2), \lambda f(x_1, r_1, t) + (1 - \lambda) f(x_1 \sigma_2, t)).$$

Now, we consider the second component

$$\lambda f(x, \sigma_1, t) + (j - \lambda) f(x, \sigma_2, t)$$

and from the linearity of f with respect σ , we have

$$\lambda f(x, \sigma_1, t) + (1 - \lambda) f(x, \sigma_2, t) = f(x, \lambda \sigma_1 + (1 - \lambda) \sigma_2, t)$$

Moreover, $\overline{\sigma} = \lambda \sigma_1 + (1 - \lambda)\sigma_2 \in U_{ad}$. In fact, $\lambda \sigma_1 + (1 - \lambda)\sigma_2$ is measurable, $\lambda \sigma_1 + (1 - \lambda)\sigma_2 \geqslant 0$ and if we consider $\sigma_1 = (\sigma_{11}, \ldots, \sigma_{1n})$ and $\sigma_2 = (\sigma_{21}, \ldots, \sigma_{2n})$ in U_{ad} , then we have

$$\lambda \sigma_1 + (1 - \lambda)\sigma_2 = (\lambda \sigma_{11} + (1 - \lambda)\sigma_{21}, \dots \lambda \sigma_{1n} + (1 - \lambda)\sigma_{2n})$$

$$\lambda \sigma_{11} + (1 - \lambda)\sigma_{21} + \dots + \lambda \sigma_{nr+} + (1 - \lambda)\sigma_{2n} =$$

$$\lambda (\sigma_{11} + \dots + \sigma_{1n}) + (1 - \lambda)(\sigma_{21} + \dots + \tau_{2n}) = \sigma_{Tot}.$$

Next, we will see that there exists $\gamma \leq 0$ such that

$$f_0(x, \overline{\sigma}, t) + \gamma = \lambda a_1 + (1 - \lambda)a_2.$$

We note that f_0 is constant with respect to the control variable, then

$$f_0(x, \lambda \sigma_1 + (1 - \lambda)\sigma_2, t) = \lambda f_0(x_1, \sigma_1, t) + (1 - \lambda)f_0(x, \sigma_2, t)$$

If we define $\gamma = \lambda \gamma_1 + (1 - \lambda) \gamma_2 \leq 0$, we have that

$$f_{0}(x_{1}\lambda\sigma_{1} + (1-\lambda)\sigma_{2}, t) + \gamma = [\lambda f_{0}(x_{1}\sigma_{1}, t) + (1-\lambda)f_{0}(x, \sigma_{2}, t)] + (\lambda\gamma_{1} + (1-\lambda)\gamma_{2}]$$

$$= \lambda f_{0}(x_{1}\sigma_{1}, t) + \lambda\gamma_{1} + (1-\lambda)f_{0}(x, \sigma_{2}t) + (1-\lambda)\gamma_{2}$$

$$= \lambda (f_{0}(x, \sigma_{1}, t) + \gamma_{1}) + (1-\lambda)(f_{0}(x, \sigma_{2}, t) + \gamma_{2})$$

$$= \lambda a_{1} + (1-\lambda)a_{2}.$$

Therefore, it was proved that there exists $\gamma = \lambda \gamma_1 + (1 - \lambda)\gamma_2 \leq 0$ and there exists $\overline{\sigma} = \lambda \sigma_1 + (1 - \lambda)\sigma_2 \in U_{ad}$ such that

$$(\lambda a_1 + (1 - \lambda)a_2, \lambda b_1 + (1 - \lambda)b_2) = (f_0(x, \overline{\sigma}, t) + \gamma, f(x, \overline{\sigma}, t)),$$

i.e.

$$(\lambda a_1 + (1 - \lambda)a_2, \lambda b_1 + (1 - \lambda)b_2) \in N(x, U_{ad}, t)$$

and $N(x, U_{ad}, t)$ is a convex set, for all fixed (x, t).

Moreover, U_{ad} is a compact set, since $0 \leqslant \sigma_i \leqslant \sigma_{\text{Tot}}$, $\forall i = 1, ..., n$. Finally, taking into account that the number of susceptible, infected and removed individuals are bounded by the total quantity of individuals, we have that $||x(t)|| \leqslant b$. Therefore, we have verified the hypothesis of Filippov-Cesari Existence Theorem and the thesis holds.

4 Optimality system

In this Section, we obtain the optimality system, which is derived of the Pontryagin Maximum Principle. We define the Lagrangian by

$$L = \sum_{i=1}^{n} I_i(t)$$

and the Hamiltonian by

$$H = L + \sum_{i=1}^{n} \left[\lambda_{1i} \frac{dS_i}{dt} + \lambda_{2i} \frac{dI_i}{dt} + \lambda_{3i} \frac{dR_i}{dt} \right]$$

where λ_{1i} , λ_{2i} and λ_{3i} are the adjoint variables to be determined suitably.

Theorem 4.1. Let (I^*, σ^*) be a optimal solution for the optimal control problem (2), where $I^*(t) = (I_1^*(t), \dots, I_n^*(t))$ and $\sigma^*(t) = (\sigma_1^*(t), \dots, \sigma_n^*(t))$. Then there exist adjoint variables λ_{1i} , λ_{2i} and λ_{3i} for $i = 1, \dots, n$, that satisfy

$$\frac{d\lambda_{1i}}{dt} = -\lambda_{1i} \left(\sum_{j=1}^{n} \beta_{ij} I_j(t) - \delta \sigma_i - d \right) - \lambda_{2i} \left(\sum_{j=1}^{n} \beta_{ij} I_j(t) \right) - \lambda_{3i} \delta \sigma_i$$
 (7)

$$\frac{d\lambda_{2i}}{dt} = -1 - \lambda_{2i} \left(d + \alpha + \gamma \right) - \lambda_{3i} \gamma \tag{8}$$

$$\frac{d\lambda_{3i}}{dt} = \lambda_{3i}d\tag{9}$$

with transversality conditions

$$\lambda_{1i}(T) = \lambda_{2i}(T) = \lambda_{3i}(T) = 0 \tag{10}$$

Furthermore, the optimality equation is given by

$$(\lambda_{1i} - \lambda_{1i})\delta S_i^* = 0. (11)$$

Proof. We know, from the Pontryagin Maximum Principle (see [?]), that if (I^*, σ^*) is a optimal solution for the optimal control problem (2), then there exists a vectorial function $\lambda(t) = (\lambda_1(t), \dots, \lambda_n(t))$ satisfying the following equalities

$$\left\{ \begin{array}{l} \frac{d\lambda}{dt} = -\frac{\partial H(t,I^*(t)),\sigma^*(t),\lambda(t)}{\partial I} \\ \frac{\partial H(t,I^*(t)),\sigma^*(t),\lambda(t)}{\partial \sigma} = 0 \\ \frac{dI}{dt} = -\frac{\partial H(t,I^*(t)),\sigma^*(t),\lambda(t)}{\partial \partial \lambda}. \end{array} \right.$$

Here, taking into account that

$$H = \sum_{i=1}^{n} I_i(t) + \sum_{i=1}^{n} \left[\lambda_{1i} \frac{dS_i}{dt} + \lambda_{2i} \frac{dI_i}{dt} + \lambda_{3i} \frac{dR_i}{dt} \right]$$

we have

$$\begin{split} \frac{d\lambda_{1i}}{dt} &= -\frac{dH}{dS_i} = -\frac{dI_i}{dS_i} - \left[\lambda_{1i}\frac{\partial}{\partial S_i}(\frac{dS_i}{dt}) + \lambda_{2i}\frac{\partial}{\partial S_i}(\frac{dI_i}{dt}) + \lambda_{3i}\frac{\partial}{\partial S_i}(\frac{dR_i}{dt})\right] \\ \frac{d\lambda_{2i}}{dt} &= -\frac{dH}{dI_i} = -\frac{dI_i}{dI_i} - \left[\lambda_{1i}\frac{\partial}{\partial I_i}(\frac{dS_i}{dt}) + \lambda_{2i}\frac{\partial}{\partial I_i}(\frac{dI_i}{dt}) + \lambda_{3i}\frac{\partial}{\partial I_i}(\frac{dR_i}{dt})\right] \\ \frac{d\lambda_{3i}}{dt} &= -\frac{dH}{dR_i} = -\frac{dI_i}{dR_i} - \left[\lambda_{1i}\frac{\partial}{\partial R_i}(\frac{dS_i}{dt}) + \lambda_{2i}\frac{\partial}{\partial R_i}(\frac{dI_i}{dt}) + \lambda_{3i}\frac{\partial}{\partial R_i}(\frac{dR_i}{dt})\right] \end{split}$$

that is, we obtain the equations (7), (8) and (9), with the transversality conditions (10). Moreover, the optimality equation is given by

$$\frac{\partial H(t, I^*(t)), \sigma^*(t), \lambda(t))}{\partial \sigma} = 0$$

i.e.

$$(\lambda_{1i} - \lambda_{1i})\delta S_i^* = 0.$$

and the thesis holds.

Remark 4.2. We note that the optimal control σ^* is not explicit in the optimality equation (11).

5 Numerical results

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References

[Filippov, 1988] Filippov, A. F. (1988). Differential Equations With Discontinuous Righthand Sides: Control Systems. Springer Science & Business Media.

 $[{\rm Meyer,\,2000}]\,$ Meyer, C. D. (2000). Matrix Analysis and Applied Linear Algebra. SIAM.

[Seierstad and Sydsæter, 1987] Seierstad, A. and Sydsæter, K. (1987). Optimal Control Theory With Economic Applications, volume 1. Elsevier Science.