

# Optimal Vaccination Strategies in Interconnected Metropolitan Areas

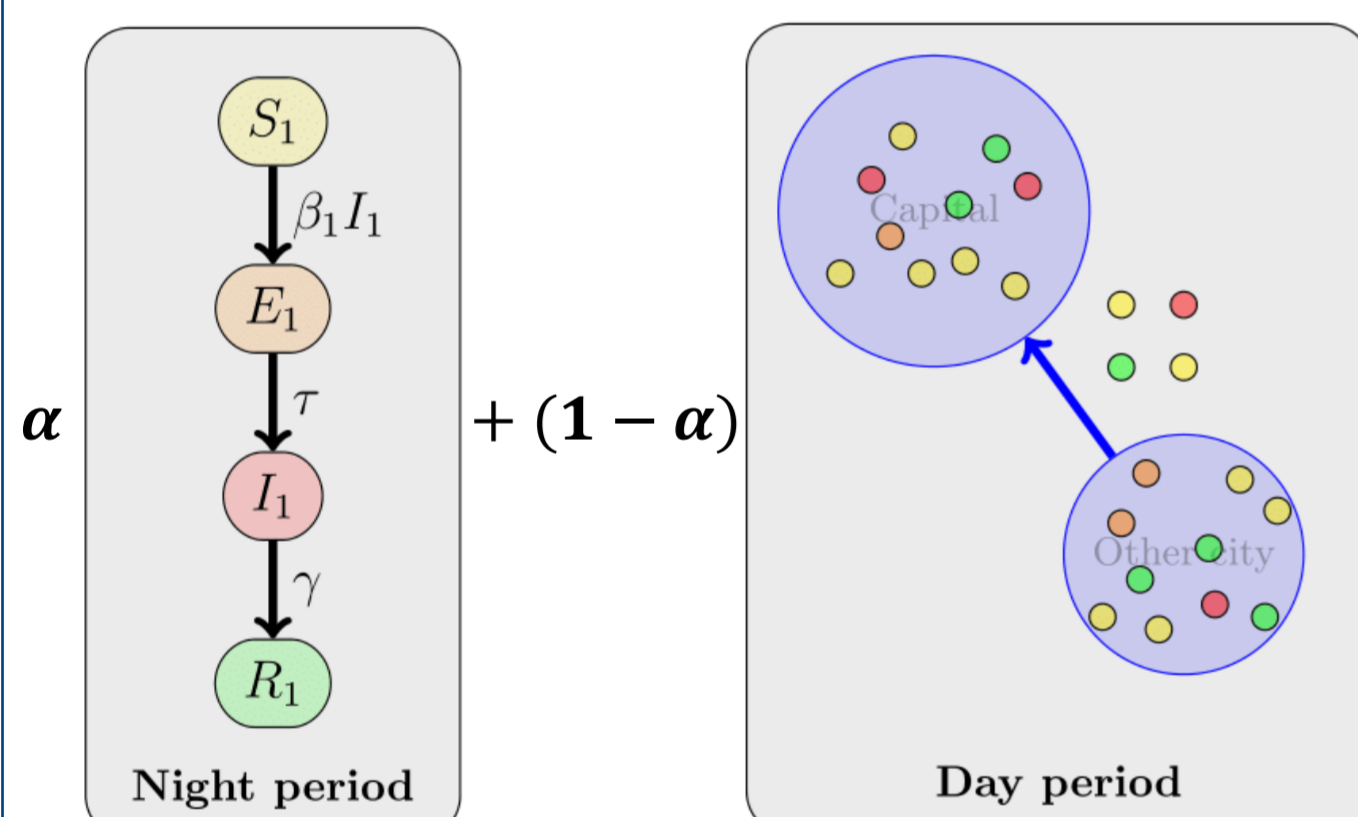
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## Highlights

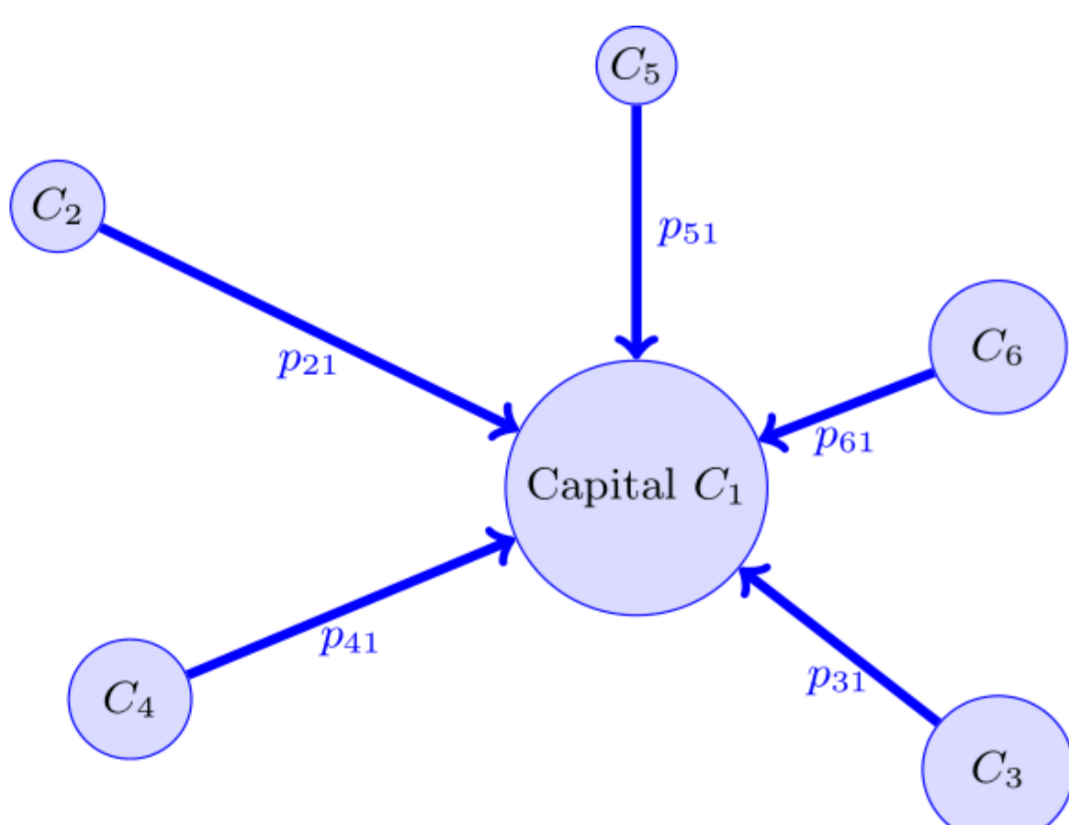
- Developed a mathematical model to track the **spread of an epidemic** in major metropolitan areas, including Rio de Janeiro, Paris, and New York.
- Implemented a centralized vaccination strategy in these regions by solving a **high-dimension optimal control problem with constraints**.
- Conducted numerical experiments that suggest a **higher vaccination rate in the capital** city can be beneficial, depending on the cost of the vaccine.

## Modelling strategy

- **Compartmental model** based on [1].
- The metropolitan area is divided in cities, which contains Susceptible, Exposed, Infectious and Recovered individuals as a usual SEIR model.
- A proportion  $p_{ij}$  of the city  $i$  works in the city  $j$  during the day.
- The individuals spend a proportion  $\alpha$  of their day in their home city and  $1 - \alpha$  in the capital working, yielding the following diagram



- We assume a **metropolitan structure**: there is a larger city, the capital, that attracts most of the workers from other cities.
- Then, a parcel  $p_{k1}$  of each city works in the **capital**, leading to the following graph:



- The **force of infection** is  $\alpha\beta_k I_k$  in the night period and

$$(1 - \alpha)(p_{k1}\beta_1 I_1^{eff} + p_{kk}\beta_k I_k^{eff})$$

during the working hours for each non-capital city, where  $I_k^{eff}$  is the effective number of infectious individuals in city  $k$  during the day.

- The parameter  $\beta$  represents the infection rate, which depends on the **city's density**. The parameter  $\tau$  is the inverse of the **infectious period**, and  $\gamma$  is the inverse of the **recovery period**.

## Basic reproduction number

- We calculate the  $R_0$  using the spectral radius of the **next-generation matrix**, which reflects the rate of new infections and the duration of infectiousness.
- Despite not being able to obtain a closed-form expression, we establish a **general bound**:

$$\min_i v_i \leq R_0 \leq \max_i v_i,$$

in which  $v_i = \alpha R_0^i + (1 - \alpha) \sum_{j=1}^K p_{ij} R_0^j$ .

- Another bound uses the **assumption of a metropolitan area**. Assuming  $\alpha \geq 0.5$ , we observed numerically that this bound is tighter in 80% of the times.

## Optimal control problem

- We include **vaccination** as a control strategy to curb the epidemic: susceptible, exposed and recovered individuals receive a vaccine at a rate  $u_i$  depending on the city.
- The model aims to minimize a cost functional that balances the **number of vaccinated and hospitalized individuals** at the final time  $T$ .
- We consider **capacity and logistic restrictions**: a weekly cap of available vaccines and an instantaneous cap of vaccinated individuals.
- The final model is a **mixed control-state and pure-state** constrained optimal control problem:

$$\min_u \sum_{i=1}^K c_v n_i V_i(T) + c_h \int_0^T r_h n_i I_i dt,$$

$$\text{s.a.} \sum_{j=1}^K V_j(t) n_j \leq D(t), \text{ a.e. } t \in [0, T]$$

$$u_i(t) \cdot (S_i(t) + E_i(t) + R_i(t)) \leq D_i, \text{ a.e. } t \in [0, T]$$

$$u_i(t) \geq 0, \text{ a.e. } t \in [0, T]$$

$$\frac{dS_i}{dt} = -\alpha\beta_i S_i I_i - (1 - \alpha) S_i \sum_{j=1}^K \beta_j p_{ij} I_j^{eff} - u_i S_i$$

$$\frac{dE_i}{dt} = \alpha\beta_i S_i I_i + (1 - \alpha) S_i \sum_{j=1}^K \beta_j p_{ij} I_j^{eff} - \tau E_i - u_i E_i$$

$$\frac{dI_i}{dt} = \tau E_i - \gamma I_i$$

$$\frac{dR_i}{dt} = \gamma I_i - u_i R_i$$

$$\frac{dV_i}{dt} = u_i \cdot (S_i + E_i + R_i)$$

## Theoretical results

- We proved **existence of optimal solution** for our problem as an application of Cesari's paper [2].
- We derived the **necessary conditions** based on the work of Boccia, De Pinho and Vinter [3].
- The pure-state constraint poses a challenge due to the **corresponding multiplier being a measure**.
- This allowed us to analyze the general behavior of the solution. We could verify that **the solution attains the bounds**, that is,

$$u_i(S_i + E_i + R_i) \in \{0, D_i\}.$$

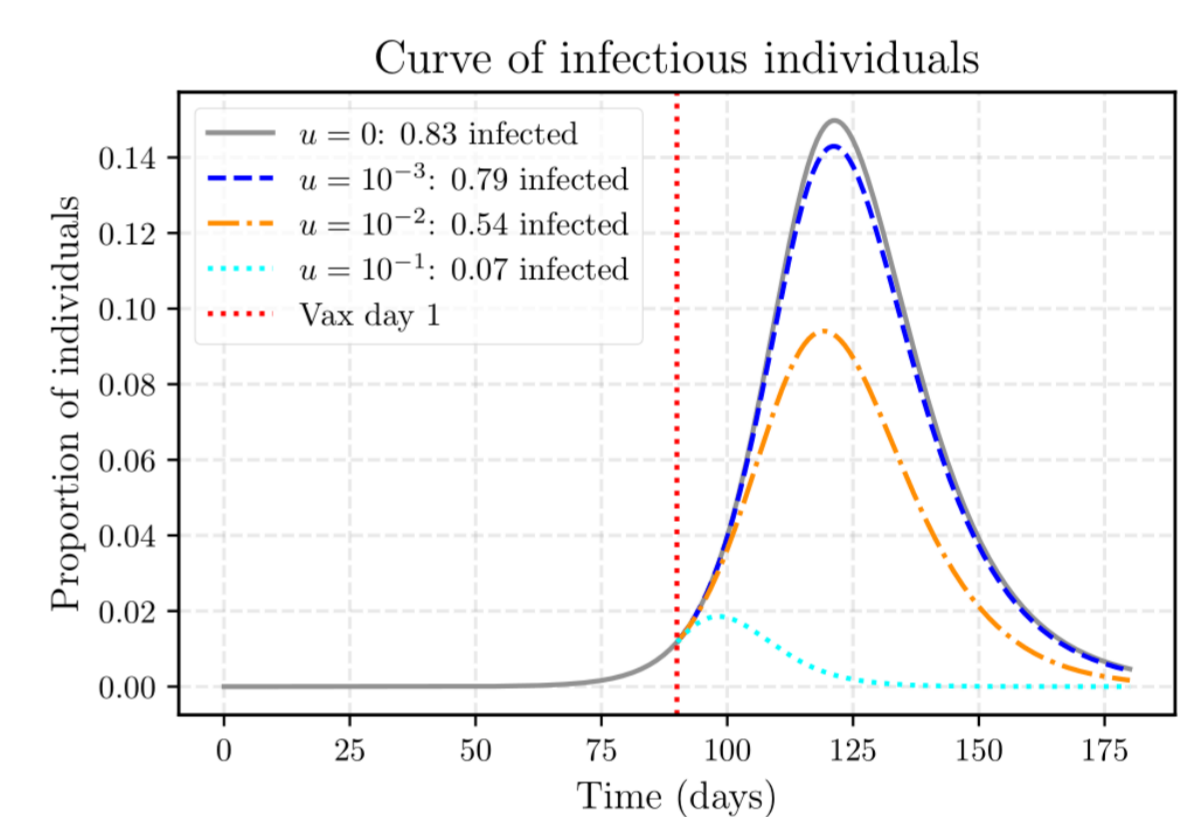
- This result exhibits characteristics like the **Bang Bang solution**, but with variable bounds.

## Numerical experiments

- We noticed that  $R_0$  is **most influenced by the value of  $\beta_1$** , the infection rate of the capital. So, decreasing it is more relevant.
- The parameters  $\alpha$  and  $p_{ij}$  do not change the behavior of the epidemic but **increase it in more interconnected regions**.
- The metropolitan area assumption resulted in worse upper and lower bounds in only 4% of the time compared to the general bound. We highlight the percentage of times the upper (UB) and lower (LB) bounds were better.

	UB	LB
49%	X	X
46%	X	
1%		X

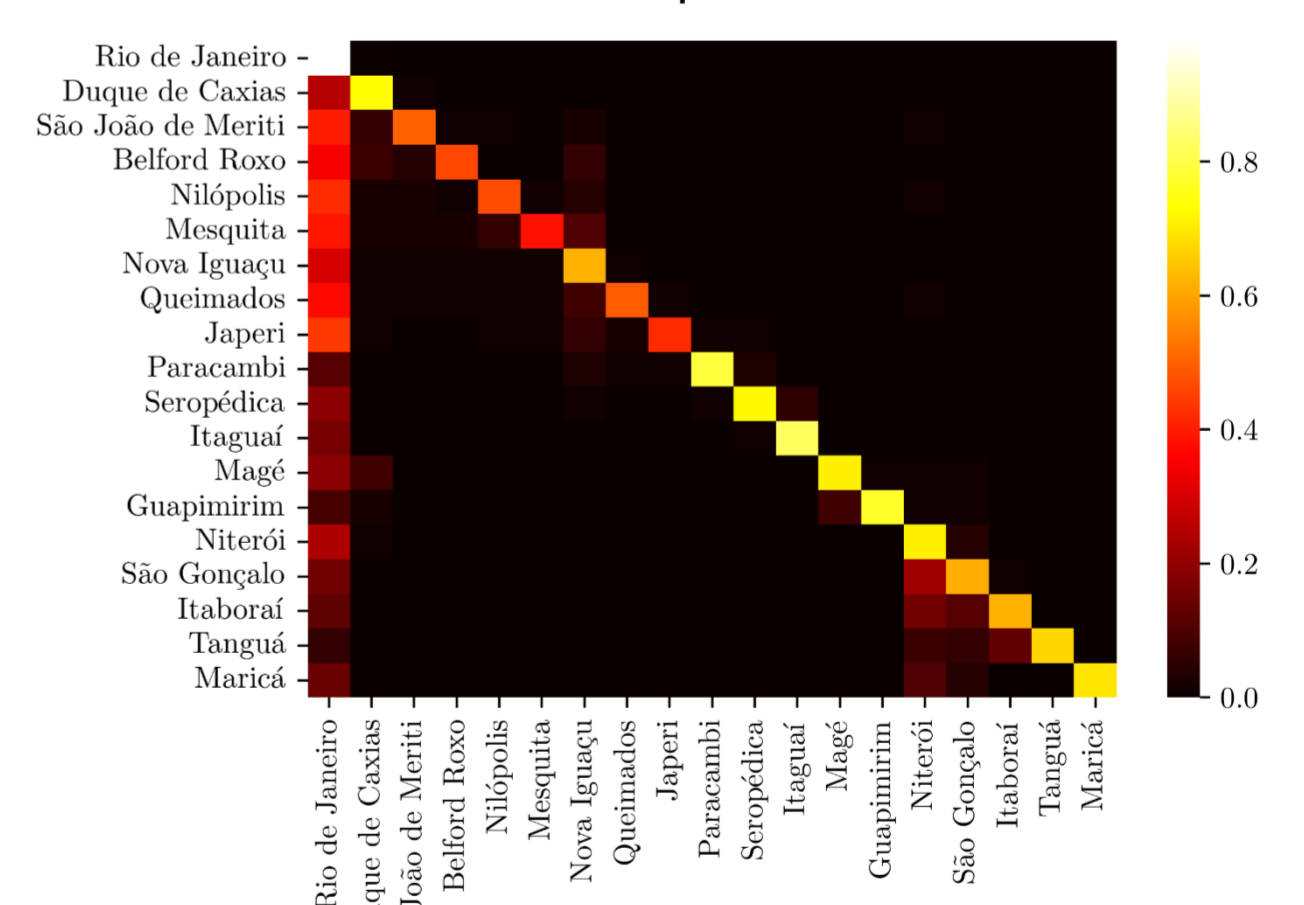
- When including vaccination, we can verify its **relevance in reducing the number of infections** in the metropolitan region:



- We verified that the optimizer prefers to **vaccinate the capital** when the susceptible population still plays a role.

## The case of Rio de Janeiro

- The transition matrix of working place in Rio de Janeiro shows our assumption:



## Discussion and conclusions

- **Optimal control** is a powerful tool with a robust mathematical foundation that can provide solutions to **real-world problems**.
- A comprehensive understanding of the model can be achieved by studying the problem **theoretically and numerically**.
- There is a **gap in the field** regarding second-order and sufficient conditions for optimality in control-affine problems with constraints. This presents an opportunity for **further research** and development.
- Our study found that a **higher vaccination rate in the capital** accelerate the control of the epidemic. However, the interplay of various factors in this process is complex.

## References

- 1) L. G. Nonato, P. Peixoto, T. Pereira, C. Sagastizábal, and P. J. S. Silva. Robot Dance: A mathematical optimization platform for intervention against COVID-19 in a complex network. EURO Journal on Computational Optimization 10, 2022.
- 2) L. Cesari. Existence Theorems for Optimal Solutions in Pontryagin and Lagrange Problems. Journal of the SIAM, Control, 1965.
- 3) A. Boccia, M. D. R. de Pinho, and R. B. Vinter. Optimal Control Problems with Mixed and Pure State Constraints. SIAM Journal on Control and Optimization, 2016.