## **Stochastic Simulation**

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# Project - 10

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## Stochastic simulation of epidemic models

This mini-project concerns stochastic simulation of epidemic models. There are multiple advantages of considering stochastic models over their deterministic counterparts in epidemiology. First, the insurgence of a contagion when two or more individuals are in contact seems to be better described as a probabilistic rather than a purely deterministic process. Second, it is difficult in deterministic models to admit the possibility of the sudden extinction of the epidemic or the possibility of a minor epidemic outbreak. On the other hand, stochastic models can naturally estimate quantities such as the probability of extinction or the probability that the epidemic lasts longer than a certain time.

## 1 Introduction and background

Let us consider a closed population of size N partitioned into three non-overlapping classes: S (susceptible), I (infectious) and R (removed). Assume that individuals in the I-class are at the same time infected and infectious, i.e., they are able to transmit the disease to susceptible individuals. Contagious contacts only can happen when an individual from the S-class meets an infectious individual. Infected individuals recover after an exponentially distributed random time and gain immunity to the disease, thus becoming individuals of the R-class. In the model considered in this project we assume that immunity lasts forever; that is, once individuals enter the R-class, they are no longer susceptible to the infection. Since we are considering a closed population in this simple SIR model, the sum S + I + R is constant in time and equals the total number N of individuals in the population. Therefore, we just need to keep track of the sizes of S and I classes. Denote by  $(s,i) \in \mathbb{N}^2$  the number of susceptible (s) and infected (i) people at a given time. The (stochastic) SIR model can be then modeled by the following discrete-state, continuous time Markov process  $\{X(t) = (S(t), I(t)) \in \{0, 1, \ldots, N\}^2, t > 0\}$  with transition probabilities:

$$\mathbb{P}(X(t+dt) = (s,i) + (-1,1)|X(t) = (s,i)) = \beta sidt + o(dt) 
\mathbb{P}(X(t+dt) = (s,i) + (0,-1)|X(t) = (s,i)) = \gamma idt + o(dt),$$
(1)

where  $\beta, \gamma \in \mathbb{R}^+$ .

When  $N, S(t), I(t) \ge 1$ , are large, so that  $(S(t), I(t)) + (-1, 1) \approx (S(t), I(t))$  and  $(S(t), I(t)) + (0, -1) \approx (S(t), I(t))$ , i.e., a single reaction changes very little the state, one can obtain the

so-called mean field approximation of SIR which can be written in differential form as

$$\frac{\mathrm{d}S}{\mathrm{d}t} = -\beta S(t)I(t),$$

$$\frac{\mathrm{d}I}{\mathrm{d}t} = \beta S(t)I(t) - \gamma I(t),$$
(2)

with  $(S(0), I(0)) = (S_0, I_0)$ , where we assume here  $S(t), I(t) \in \mathbb{R}^+$   $\forall t$ . A better approximation is given by the following diffusion approximation of SIR:

$$dS = -\beta S(t)I(t)dt - \sqrt{\beta S(t)I(t)}dW_S$$

$$dI = (\beta S(t)I(t) - \gamma I(t))dt + \sqrt{\beta S(t)I(t)}dW_S - \sqrt{\gamma I(t)}dW_I$$
(3)

with  $(S(0), I(0)) = (S_0, I_0)$ , and where  $W_S, W_I$  are two independent standard Wiener processes.

A more realistic epidemic model, hereafter named SIR-d, takes into account demographic effects, i.e., the death and birth of the population. Its mean field approximation is given by

$$\frac{\mathrm{d}S}{\mathrm{d}t} = m(S(t) + I(t) + R(t)) - mS(t) - \beta S(t)I(t),$$

$$\frac{\mathrm{d}I}{\mathrm{d}t} = \beta S(t)I(t) - \gamma I(t) - (m+v)I(t),$$

$$\frac{\mathrm{d}R}{\mathrm{d}t} = \gamma I(t) - mR(t),$$
(4)

where m is the host death/birth rate and v is the pathogen-induced mortality rate. The term that describes the birth of susceptible hosts, m(S + I + R), ensures that deaths due to non-pathogen-related causes are balanced, and the total population (S + I + R) remains constant over time, as long as there is no death due to the epidemic (expressed by -vI). Thus, at any given time t, one of the following 6 processes  $a_i$  can happen:  $(a_1)$  host birth,  $(a_2)$  death of susceptible host,  $(a_3)$  death of infected host,  $(a_4)$  death of recovered host,  $(a_5)$  infection,  $(a_6)$  Recovery.

#### Goals of the project

- (a) We begin by comparing the dynamics generated by (1), (2) and (3). To that end, consider a small population of N=100 individuals and initial condition  $(S_0, I_0)=(99,1)$ . Implement both the stochastic SIR (1), its mean field (2) and diffusion (3) approximations for a time T=10. In particular, simulate exactly the stochastic SIR, and use suitable time discretizations for the mean field and diffusion models. Plot the resulting trajectories. Run your simulations for  $\beta=0.02$  and  $\gamma=0.4$ .
- (b) In the stochastic SIR model, the disease is considered extinct whenever I(t) = 0 for some  $t \in [0, T]$ . Using the same setting as in the previous point, use the SIR model (1) and a Monte Carlo approach to estimate the probability of extinction of the disease at time T, that is, estimate  $\mathbb{P}(I(t) = 0)$  for some  $t \leq T$ . Report your values for T = 1, 2, 10. Choose appropriately the sample size to guarantee a relative error of 5%.

- (c) Propose and implement a variance reduction technique for your previous estimate. Discuss your results. Test now your proposed method to estimate the probability of extinction at T=2, starting from the initial condition  $(S_0, I_0)=(95, 5)$ . Comment the results and suggest possible improvements, if needed.
- (d) Derive the formulation of the stochastic SIR-d process associated to (4). **Hint:** What is the rate of each  $(a_i)$ , i = 1, 2, ..., 6?
- (e) Simulate then the SIR-d process and compare to its mean field approximation (4). Set  $T=10, \, \beta=0.02, \, \gamma=0.4$  and experiment with different values of  $m\in[10^{-4},10^{-3}]$  and  $v\in[10^{-3},10^{-2}]$ . Compare your results.
- (f) In a deterministic model an epidemic will go extinct (i.e.,  $I(t) \to 0$  as  $t \to \infty$ ) if the basic reproduction number,  $R_0$  of the infection, given by

$$R_0 = \frac{\beta(S(0) + I(0) + R(0))}{m + v + \gamma},$$

is less than one. Consider the stochastic SIR-d model with demographic effects. Is it possible that the epidemic will be extinct at a time T for  $R_0 = 1.01, 1.05, 1.1, 1.5$ ? Consider a population of N = 100 individuals with  $R(0) = 0, S(0) = 95, I(0) = 5, m = 10^{-4}, v = 10^{-2}$  and  $\beta$  chosen to obtain the previous values of  $R_0$  and estimate the probability of extinction for T = 1, 2, 10.

#### References

- [1] Priscilla E Greenwood and Luis F Gordillo. Stochastic epidemic modeling. In *Mathematical and statistical estimation approaches in epidemiology*, pages 31–52. Springer, 2009.
- [2] Alvaro Moraes. Simulation and statistical inference of stochastic reaction networks with applications to epidemic models, Chapter 2. PhD thesis, King Abdullah University of Science and Technology, 2015.