# Lecture 10: SEIR Models

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# The Susceptible-Exposed-Infectious-Removed (SEIR) Model

The Susceptible-Exposed-Infectious-Removed (SEIR) model is obtained from SIR by introducing a compartment between Susceptible and Infected of population that has been exposed to the virus but are not yet contagious:

$$\begin{cases} \frac{dS}{dt} = -\beta \frac{SI}{N} , S(0) \\ \frac{dE}{dt} = \beta \frac{SI}{N} - \delta E , E(0) \\ \frac{dI}{dt} = \delta E - \alpha I , I(0) \\ \frac{dR}{dt} = \alpha I , R(0) \end{cases}$$
(SEIR Model)

where  $\delta \ge 0$  is the rate of transition from exposed to infected. Its reciprocal  $1/\delta$  represents the *average incubation period*.

If data is selected from the onset of infections, a natural initial condition is:  $R(0) = N \gg I(0), R(0) = 0$ . The initial exposed population E(0) may be set to I(0), or can be fine tuned to fit the data. Assumin E(0) = I(0) is known, the parameters that need to be calibrated are:  $\alpha, \beta, \gamma, \delta, N$ .

Numerical Results

# Simulations

#### Here are a few numerical results with the Euler's scheme:



Figure: Top left:  $\alpha = 0.2$ ,  $\beta = 0.8$ ,  $\delta = 0.3$ ; Top right:  $\alpha = 0.2$ ,  $\beta = 1.0$ ,  $\delta = 0.3$ ; Bottom left:  $\alpha = 1.0$ ,  $\beta = 1.5$ ,  $\delta = 1.0$ ; Bottom right:  $\alpha = 0.08$ ,  $\beta = 0.176$ ,  $\delta = 0.8$ 

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# Agent Based Simulation

Let  $T_0$  denote the Markov-Chain simulation time step. As in the case of SIR, the SEIR model can be implemented a piecewise linear differential system by replacing I(t) in  $\beta \frac{SI}{N}$  with the previous estimate  $I((p-1)T_0)$  for the interval  $t \in [(p-1)T_0, pT_0]$ . The dynamical system becomes:  $\begin{cases} \frac{dS(t)}{dt} = -\beta \frac{I((p-1)T_0)}{N}S(t) , S(0) \\ \frac{dE(t)}{dt} = \beta \frac{I((p-1)T_0)}{N}S(t) - \delta E(t) , E(0) \\ \frac{dI(t)}{dt} = \delta E(t) - \alpha I(t) , I(0) \\ \frac{dR(t)}{dt} = \alpha I(t) , R(0) \end{cases}$ for  $(p-1)T_0 \le t \le pT_0$ 

This defines the transition probability matrix:

$$\Pi = exp\left( \begin{bmatrix} T_0 - \frac{\beta}{N}I((p-1)T_0) & 0 & 0 & 0\\ \frac{\beta}{N}I((p-1)T_0) & -\delta & 0 & 0\\ 0 & \delta & -\alpha & 0\\ 0 & 0 & \alpha & 0 \end{bmatrix} \right)$$

# Agent Based Simulation (2)

The Markov Chain has the following form:



# Examples of Agent Based Simulations

Here are results of a large number of simulations  $(10^3)$  for  $T_0 = 0.01$ , with parameters  $\beta = 1$  and  $\alpha = 0.2$ ,  $\delta = 0.3$ , initial condition  $S_0 = 2000$ ,  $E_0 = I_0 = 23$ and  $R_0 = 0$ . The numerical solution is obtained with the Euler scheme and a stepsize h = 0.01. The shaded area has semiwidth of one std of simulations



Numerical Results

### Examples of Agent Based Simulations

Here are results of a large number of simulations  $(10^3)$  for  $T_0 = 1.00$ , with parameters  $\beta = 1$  and  $\alpha = 0.2$ ,  $\delta = 0.3$ , initial condition  $S_0 = 2000$ ,  $E_0 = I_0 = 23$ and  $R_0 = 0$ . The numerical solution is obtained with the Euler scheme and a stepsize h = 0.01. The shaded area has semiwidth of one std of simulations



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MATH 420: SEIR Calibration

Numerical Results

### Examples of Agent Based Simulations

Here are results of a large number of simulations  $(10^3)$  for  $T_0 = 0.001$ , with parameters  $\beta = 1$  and  $\alpha = 0.2$ ,  $\delta = 0.3$ , initial condition  $S_0 = 2000$ ,  $E_0 = I_0 = 23$ and  $R_0 = 0$ . The numerical solution is obtained with the Euler scheme and a stepsize h = 0.01. The shaded area has semiwidth of one std of simulations



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MATH 420: SEIR Calibration

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Numerical Results

# SEIR Model Calibration

Recall the model:

$$\begin{array}{rcl} \frac{dS}{dt} &=& -\beta \frac{SI}{N} \ , \ S(0) \\ \frac{dE}{dt} &=& \beta \frac{SI}{N} - \delta E \ , \ E(0) \\ \frac{dI}{dt} &=& \delta E - \alpha I \ , \ I(0) \\ \frac{dR}{dt} &=& \alpha I \ , \ R(0) \end{array}$$

with the sub-compartments  $X(t) = (1 - \gamma)R(t)$  for "recovered" and  $Y(t) = \gamma R(t)$  for deaths.

Before making useful predictions (testing), the model has to be calibrated. We use the same calibration strategy as in the SIR model.

### Optimization based on Forward Model Simulation Pre-processing steps

**Step 1:**  $t_0$ . First detect the onset of infections, and reset the time origin to match this starting time  $t_0$ .

**Step 2:** I(t). The time series of cumulative detected infections  $\{V(0), \dots, V(T_{T_{max}})\}$  should be turned into a rate of infections. The daily rate may not be relevant to infection transmissions. Instead use a time window to convert the cumulative count into a rate:

$$I(t) = V(t + t_0 + \tau_0) - V(t + t_0 - \tau_0)$$
,  $t = 0, 1, 2, \cdots, T_{max}$ 

where  $\tau_0 > 0$  is chosed so that  $\tau_0$  accounts for average infection period. For Covid-19 it is somewhere between 5 days and 10 days. A possible value is  $\tau_0 = 7$ . Thus I(t) measures the number of infections in a 2-week period centered around t.

**Step 3:**  $Y_{measured}$ . Align the time series of cumulative death with the onset time  $t_0$ :

$$Y_{measured}(t)=Y(t+t_0)$$
 ,  $t=0,1,2,\cdots,T_{max}$  .

# Optimization based on Forward Model Simulation The Meta Loop

A natural choice for initial conditions is given by: S(0) = N and R(0) = 0, where we assumed  $E(0) + I(0) \ll N$ . Another choise is E(0) = I(0) since we do not know the undetected number of infections at time 0 (perhaps E(0) > I(0) is closer to the truth):

$$\begin{cases} \frac{dS}{dt} &= -\beta S \frac{I}{N} , \quad S(0) = N \\ \frac{dE}{dt} &= \beta \frac{SI}{N} - \delta E , \quad E(0) \\ \frac{dI}{dt} &= \delta E - \alpha I , \quad I(0) \\ \frac{dR}{dt} &= \alpha I , \quad R(0) = 0 , \quad Y = \gamma R \end{cases}$$

At this point, the parameters that need to be estimated are:  $\{\alpha, \beta, \delta, \gamma, \rho\}$ . The Forward Model based calibration works like this:

1. Construct a search set  $\Omega$  of the "free" parameters  $(\alpha,\beta,\delta)$ 

2. For each  $(\alpha, \beta, \delta) \in \Omega$ : (i) run a SIR simulator using the Euler scheme that produces  $(S_{sim}, E_{sim}, I_{sim}, R_{sim})$ . (ii) Fit  $\hat{\gamma}$  to match the observed time series  $Y_{measured}$ . (iii) Fit  $\hat{\rho}$  that accounts the undercounting of actual infections in the observed time series *I*. (iv) Compute the value of the objective function  $J_p(\alpha, \beta, \delta, \hat{\gamma}, \hat{\rho})$ . 3. Select  $(\hat{\alpha}, \hat{\beta}, \hat{\delta}, \hat{\gamma}, \hat{\rho})$  that minimize *J*.

### Optimization based on Forward Model Simulation Details: The objective function

The objective function  $J_p$  can be chosen to measure a norm of residuals  $I - \rho I_{sim}$  and  $Y_{measured} - \gamma R_{sim}$ . To do so, you need to choose and fix meta-parameters p,  $c_I$  and  $c_Y$ . For  $1 \le p < \infty$  define

$$J_{p}(\alpha,\beta,\delta,\gamma,\rho) = c_{I} \sum_{t=0}^{T_{max}} |I(t) - \rho I_{sim}(t)|^{p} + c_{Y} \sum_{t=0}^{T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)|^{p}$$

For  $p = \infty$  define

$$J_{\infty}(\alpha,\beta,\delta,\gamma,\rho) = c_{I} \max_{0 \le t \le T_{max}} |I(t) - \rho I_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le t \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{measured}(t) - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)| + c_{Y} \max_{0 \le T_{max}} |Y_{max} - \gamma R_{sim}(t)|$$

Typical choices:  $p = 1, 2, \infty$  and  $(c_I, c_Y) = (0, 1)$  (if the cumulative detected infections are unreliable) or  $(c_I, c_Y) = (1, 1)$  (if detected infections is a reliable measure).

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#### The Calibration Algorithm for SEIR Models Algorithm (Meta-Algorithm for SEIR Calibration)

Inputs: Time series  $\{V(0), \dots, V(T)\}$ ,  $\{Y(0), \dots, Y(T)\}$ . Parameters:  $V_{min}$  (default,  $\overline{V_{min}} = 5$ ),  $\tau_0$  (default,  $\tau_0 = 7$ ), N = Population, E(0) (default E(0) = I(0)),  $p \in [1, \infty]$  (default, p = 2),  $c_I, c_Y > 0$ . Search set  $\Omega$ .

**1** Detect the onset of the infection  $t_0$  as the first time so that  $V(t_0) \ge V_{min}$ , reset the time origin, and create the time series of infection rates  $I(t) = V(t + t_0 + \tau_0) - V(t + t_0 - \tau_0)$ , and aligned cumulative death  $Y_{measured}(t) = Y(t + t_0), 0 \le t \le T_{max}$ .

2 For each  $(\alpha, \beta, \delta) \in \Omega$  repeat:

Simulate a SEIR model with parameters (α, β, δ) and initial condition S<sub>sim</sub>(0) = N, E<sub>sim</sub>(0) = E(0), I<sub>sim</sub>(0) = I(0), R<sub>sim</sub>(0) = 0, and obtain daily time series (S<sub>sim</sub>, E<sub>sim</sub>, I<sub>sim</sub>, R<sub>sim</sub>).

2 Solve 
$$\hat{\gamma} = \operatorname{argmin}_{\gamma} \| Y_{\text{measured}} - \gamma R_{\text{sim}} \|_{p}$$
.

3 Solve 
$$\hat{\rho} = \operatorname{argmin}_{\rho} \|I - \rho I_{sim}\|_{\rho}$$

**3** Compute the objective function  $J = J_p(\alpha, \beta, \delta, \hat{\gamma}, \hat{\rho})$ .

**③** Determine the minimum and the minimizer of J. <u>Outputs</u>: Estimated  $\hat{\alpha}, \hat{\beta}, \hat{\delta}, \hat{\gamma}, \hat{\rho}$  and minimum value of the objective function  $J_{min}$ .

# Numerical results (1)

Analysis of DC data for 2020



Numerical Results

# Numerical results (2) Analysis of DC data for 2020. p = 1, $c_l = 0$ , $c_Y = 1$



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Numerical Results

# Numerical results (3) Analysis of DC data for 2020. p = 2, $c_l = 0$ , $c_Y = 1$



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Numerical Results

# Numerical results (4) Analysis of DC data for 2020. $p = \infty$ , $c_l = 0$ , $c_Y = 1$



Numerical Results

# Numerical results (5) Analysis of DC data for 2020. p = 1, $c_l = 1$ , $c_Y = 1$



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Numerical Results

# Numerical results (6) Analysis of DC data for 2020. p = 2, $c_l = 1$ , $c_Y = 1$



Numerical Results

# Numerical results (7) Analysis of DC data for 2020. $p = \infty$ , $c_l = 1$ , $c_Y = 1$

