

# Example 1: Fitting the parameters to the model using simulated data

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## Background

A generalized SEIR model is used to simulate an epidemic breakout. Seven different states are considered in the following in a similar fashion as by [1], which is derived from SEIQR models [2] and has several similarities with mathematical models for SARS transmission, e.g. [3] :

1. Susceptibles cases  $S(t)$
2. Insusceptible cases  $P(t)$
3. Exposed cases  $E(t)$
4. Infectious cases  $I(t)$
5. Quarantined cases  $Q(t)$
6. Recovered cases  $R(t)$
7. Dead cases  $D(t)$

The parameters are as follows:

- alpha: protection rate
- beta: infection rate
- gamma: Inverse of the average latent time
- delta: rate at which infectious people enter in quarantine
- lambda: time-dependant recovery rate
- kappa: time-dependant mortality rate

The population is assumed constant, i.e. the births and natural death are not modelled. The cure rate and mortality rate are here time-dependent but they need some empirical coefficients to tune the time-dependency of these parameters.

The generalized SEIR model [1] is:

$$\frac{dS(t)}{dt} = -\alpha S(t) - \beta \frac{S(t)I(t)}{N_{pop}}$$

$$\frac{dE(t)}{dt} = -\gamma E(t) + \beta \frac{S(t)I(t)}{N_{pop}}$$

$$\frac{dI(t)}{dt} = \gamma E(t) - \delta I(t)$$

$$\frac{dQ(t)}{dt} = \delta I(t) - \lambda(t)Q(t) - \kappa(t)Q(t)$$

$$\frac{dR(t)}{dt} = \lambda(t)Q(t)$$

$$\frac{dD(t)}{dt} = \kappa(t)Q(t)$$

$$\frac{dP(t)}{dt} = \alpha S(t)$$

## Recovery and mortality rates

The content presented hereafter is **not** proposed by [1]. Therefore, it can significantly differ from their work. In the following, the mortality rate  $\kappa(t)$  is modelled as

$$\kappa(t) = \frac{\kappa_0}{\exp(\kappa_1(t - \tau_\kappa)) + \exp(-\kappa_1(t - \tau_\kappa))}$$

or as

$$\kappa(t) = \kappa_0 \exp(-[\kappa_1(t - \tau_\kappa)]^2)$$

or as

$$\kappa(t) = \kappa_0 + \exp(-\kappa_1(t + \tau_\kappa))$$

where  $\kappa_0$ ,  $\kappa_1$  and  $\tau_\kappa$  are parameters to be empirically determined. The parameters  $\kappa_0$  and  $\kappa_1$  have the dimension of the inverse of a time and  $\tau_\kappa$  has the dimension of a time.

The recovery rate  $\lambda(t)$  is either modelled as

$$\lambda(t) = \frac{\lambda_0}{1 + \exp(-\lambda_1(t - \tau_\lambda))}$$

or as

$$\lambda(t) = \lambda_0 + \exp(-\lambda_1(t + \tau_\lambda))$$

where  $\lambda_0$ ,  $\lambda_1$  and  $\tau_\lambda$  are parameters to be empirically determined. The parameters  $\lambda_0$  and  $\lambda_1$  have the dimension of the inverse of a time and  $\tau_\lambda$  has the dimension of a time.

The choice of best approximation for  $\lambda(t)$  and  $\kappa(t)$  is done automatically inside the function `fit_SEIRQDP` based on a preliminary assessment of the recovery rate and the mortality rate. The idea behind these functions is that the mortality rate should become close to zero (or a constant value  $\kappa_0$ ) at time increases while the recovery rate converges toward a constant value  $\lambda_0$ .

## Numerical solutions

I also decided to re-write the system of ODEs in a matrix form for the sake of clarity:

$$\frac{dY}{dt} = A * Y + F$$

where

$$Y = [S, E, I, Q, R, D, P]^T$$

$$A = \begin{bmatrix} -\alpha & 0 & 0 & 0 & 0 & 0 & 0 \\ 0 & -\gamma & 0 & 0 & 0 & 0 & 0 \\ 0 & \gamma & -\delta & 0 & 0 & 0 & 0 \\ 0 & 0 & \delta & -\kappa(t) - \lambda(t) & 0 & 0 & 0 \\ 0 & 0 & 0 & \lambda(t) & 0 & 0 & 0 \\ 0 & 0 & 0 & \kappa(t) & 0 & 0 & 0 \\ \alpha & 0 & 0 & 0 & 0 & 0 & 0 \end{bmatrix}$$

$$F = S(t) \cdot I(t) \cdot \begin{bmatrix} -\frac{\beta}{N_{\text{pop}}} \\ \frac{\beta}{N_{\text{pop}}} \\ 0 \\ 0 \\ 0 \\ 0 \\ 0 \end{bmatrix}$$

The equation  $\frac{dY}{dt} = A * Y + F$  is then solved using the classic 4th order Runge-Kutta method.

## Important note

The initial number of exposed  $E_0$  and infectious  $I_0$  is not known. In the present version, these values are not estimated by least-square fit although the choice of  $E_0$  and  $I_0$  could affect the goodness of the fit and the estimated parameters.

## References

- [1] Peng, L., Yang, W., Zhang, D., Zhuge, C., & Hong, L. (2020). Epidemic analysis of COVID-19 in China by dynamical modeling. *arXiv preprint arXiv:2002.06563*.
- [2] Feng, Z., & Thieme, H. R. (1995). Recurrent outbreaks of childhood diseases revisited: the impact of isolation. *Mathematical Biosciences*, 128(1-2), 93-130.
- [3] Lipsitch, M., Cohen, T., Cooper, B., Robins, J. M., Ma, S., James, L., ... & Fisman, D. (2003). Transmission dynamics and control of severe acute respiratory syndrome. *Science*, 300(5627), 1966-1970.

## Example

### Initialisation

Case of an imaginary epidemic outbreak that took place on 2010-01-01. The simulation time is set to 6 months.

```
clearvars;close all;clc;

% Time definition
dt = 0.1; % time step
time1 = datetime(2010,01,01,0,0,0):dt:datetime(2010,06,01,0,0,0);
N = numel(time1);
t = [0:N-1].*dt;
```

### Generate the data

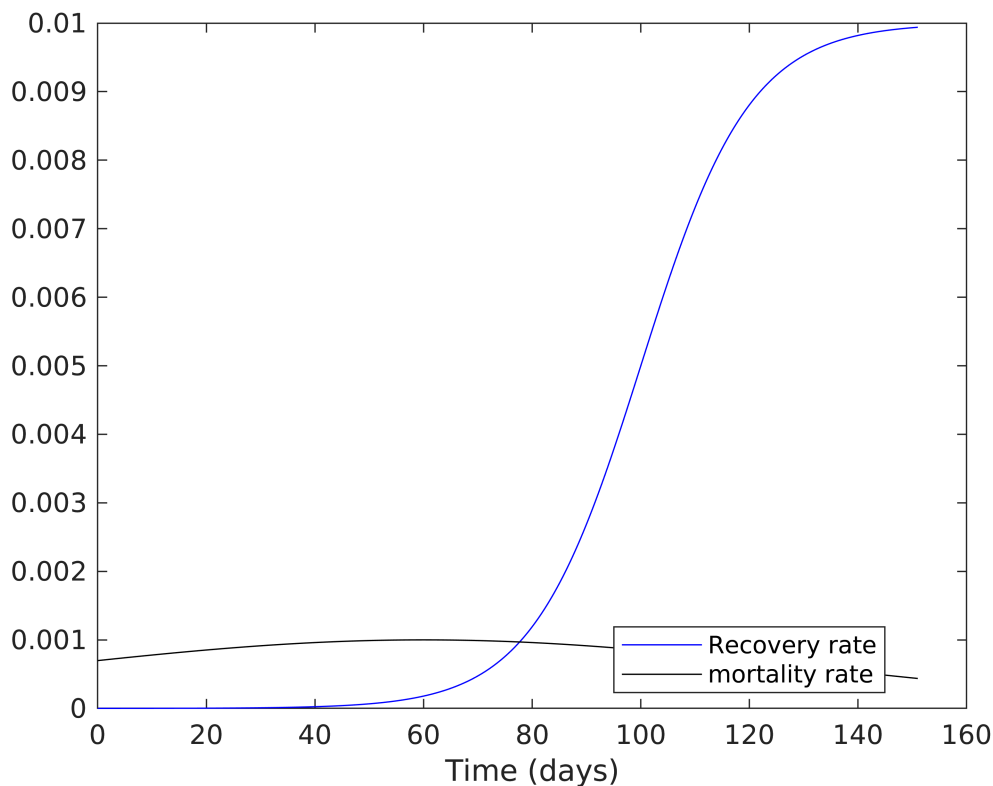
```
Npop= 60e6; % population (60 millions)
Q0 = 200; % Initial number of infectious that have bee quanrantined
I0 = Q0; % Initial number of infectious cases non-quarantined
E0 = 0; % Initial number of exposed
R0 = 10; % Initial number of recovereds
D0 = 10; % Initial number of deads
alpha = 0.08; % protection rate
beta = 0.9; % infection rate
gamma= 0.2; % inverse of average latent time
delta= 0.5; % rate at which infectious people enter in quarantine
Lambda = [0.01 0.1 100]; % cure rate (time dependant)
Kappa = [0.001 0.01,60]; % mortality rate (time dependant)

% Choice of a particular form for lambda(t)
lambdaFun0 = @(a,t) a(1)./(1+exp(-a(2)*(t-a(3))));
kappaFun0 = @(a,t) a(1).*exp(-(a(2)*(t-a(3))).^2);

[S,E,I,Q,R,D,P] = SEIQRDP(alpha,beta,gamma,delta,Lambda,Kappa,Npop,E0,I0,Q0,R0,D0,t,lan

figure
```

```
plot(t,lambdaFun0(Lambda,t),'b',t,kappaFun0(Kappa,t),'k')
legend('Recovery rate','mortality rate','location','best');
xlabel('Time (days)')
```



```
set(gcf,'color','w')
```

## Fit the data

The fitting is done using the time histories of the number of quarantined  $Q(t)$ , recovered  $R(t)$  and deads  $D(t)$  only. The number of exposed, susceptible, insusceptible and infectious is computed in the model but not used as target.

```
lambdaGuess = [0.01 0.5 0.1];
kappaGuess = [0.01 0.1,10];
alphaGuess = 0.05;
betaGuess = 0.7;
deltaGuess = 0.2;
gammaGuess = 0.3;
guess = [alphaGuess,betaGuess,deltaGuess,gammaGuess,lambdaGuess,kappaGuess]; % initial

[alpha1,beta1,gamma1,delta1,Lambda1,Kappa1,lambdaFun,kappaFun0] = fit_SEIQRDP(Q,R,D,Npop,
[S1,E1,I1,Q1,R1,D1,P1] = ...
    SEIQRDP(alpha1,beta1,gamma1,delta1,Lambda1,Kappa1,Npop,E0,I0,Q0,R0,D0,t,lambdaFun,k
```

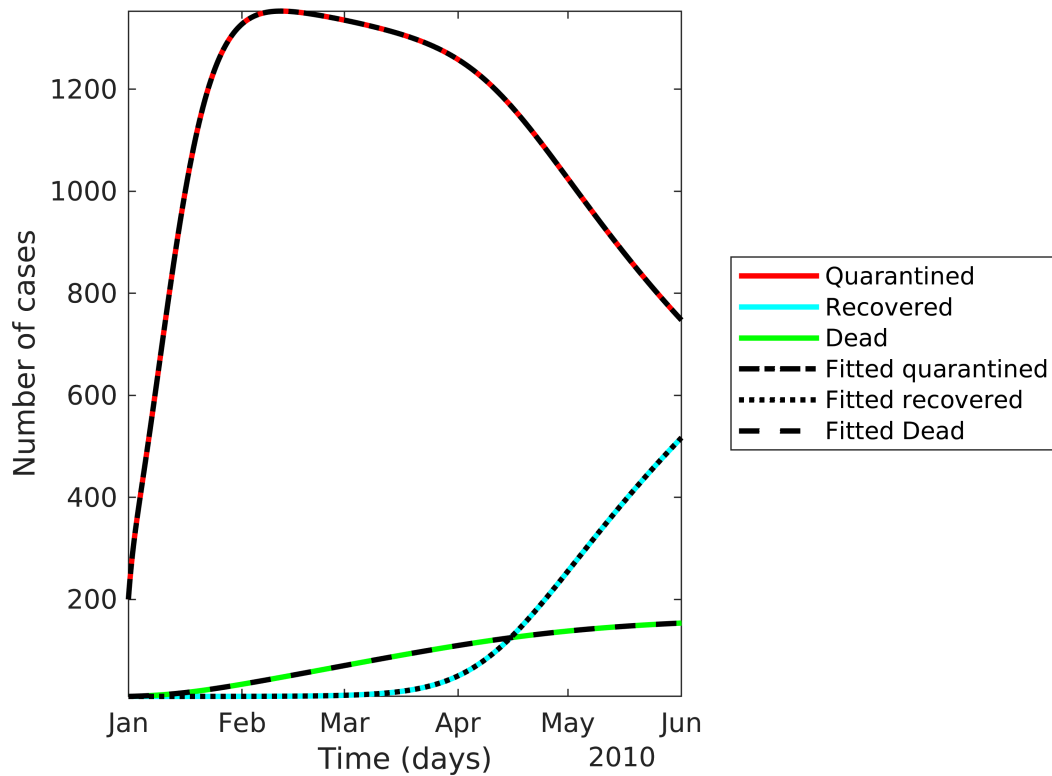
## Comparison between fitted and generated time histories

```
figure
```

```

clf;close all;
plot(time1,Q,'r',time1,R,'c',time1,D,'g','linewidth',2);
hold on
plot(time1,Q1,'k-.',time1,R1,'k:',time1,D1,'k--','linewidth',2);
% ylim([0,1.1*Npop])
ylabel('Number of cases')
xlabel('Time (days)')
leg = {'Quarantined','Recovered','Dead','Fitted quarantined','Fitted recovered','Fitted Dead'};
legend(leg{:},'location','eastoutside')
set(gcf,'color','w')
axis tight

```



### Case where the recovered (R) and quarantined (Q) data are not available separately

The number of quarantined  $Q(t)$  and recovered cases  $R(t)$  is unknown, However,  $Q(t) + R(t)$  is known.

```

guess = [alphaGuess,betaGuess,deltaGuess,gammaGuess,lambdaGuess,kappaGuess]; % initial
[alpha1,beta1,gamma1,delta1,Lambda1,Kappa1,lambdaFun,kappaFun] = ...
    fit_SEIQRDP(Q+R,[],D,Npop,E0,I0,time1,guess,'Display','off');

```

Warning: No data available for "Recovered"

```

[S1,E1,I1,Q1,R1,D1,P1] =...
    SEIQRDP(alpha1,beta1,gamma1,delta1,Lambda1,Kappa1,...
    Npop,E0,I0,Q0,R0,D0,t,lambdaFun,kappaFun);

```

```

figure
clf;close all;
plot(time1,Q+R,'r',time1,D,'g','linewidth',2);
hold on
plot(time1,Q1+R1,'k-.',time1,D1,'k--','linewidth',2);
% ylim([0,1.1*Npop])
ylabel('Number of cases')
xlabel('Time (days)')
leg = {'Tested positive minus the deceased cases','Deceased cases','Fitted Tested posit
legend(leg{:},'location','southoutside')
set(gcf,'color','w')
axis tight

```

