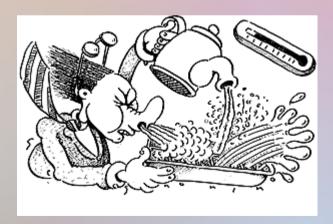
STOCHASTIC DYNAMIC MODEL OF INFECTIOUS DISEASE SPREAD

- A simplified approach

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Overview

- 1. A few words about Epidemic Models, Dynamic Behavior and Stochasticity.
- 2. Theoretical basis and assumptions of the model:
 - SEIR model and multi-compartmental formulation (age structure)
 - Seasonality time-dependent transmission rate
 - Stochasticity event-driven approach
 - Other assumptions

Overview (cont.)

- 3. Mathematical formulation.
- 4. Modeling approach tau-leap method.
- 5. Parameters initialization.
- 6. Results.
- 7. Extensions for the model future work.

1. Epidemic Models, Dynamic Behavior and Stochasticity

- SIR model (Kermack and McKendrick, 1927):
 - → Susceptible: previously unexposed to the pathogen
 - Infected : currently colonized by the pathogen
 - → Recovered: have successfully cleared the infection

$$S \longrightarrow \mathbb{I} \longrightarrow \mathbb{R}$$

Generalized SIR:
$$\frac{dS}{dt} = \mu - \beta SI - \mu S$$
(demography included,
$$\frac{dI}{dt} = \beta SI - \gamma I - \mu I$$
(lifelong immunity)
$$\frac{dR}{dt} = \gamma I - \mu R$$

$$\gamma = \text{removal (recovery) rate }_{4}$$

1. Epidemic Models, Dynamic Behavior and Stochasticity (cont.)

- Other Epidemic models :
 - SI (infection-induced mortality)
 - SIS (without immunity)
 - SIRS (waning immunity)
 - SEIR (latent period)

Dynamic behavior: S(t), I(t), R(t), β(t)

1. Epidemic Models, Dynamic Behavior and Stochasticity (cont.)

• Stochasticity: to include randomness in the model in the form of probabilities.

Events of disease transmission, recovery, birth, death (natural) do not happen at a constant rate, but probabilities rule their appearance.

Role of chance most important when the population size and the number of infectious individuals are relatively small.

- SEIR model and multi-compartmental formulation (age structure).
 - Susceptibles (X)
 - Exposed (W)

 $S \longrightarrow \mathbb{E} \longrightarrow \mathbb{I} \longrightarrow \mathbb{R}$

- Infectious (Y)
- Recovereds (Z)
- Multi-compartmental model: age structure is introduced to the model to depict the higher risk some age groups may experience.
 - ✓ 0-5 , 6-9 , 10-19, 20+ years old

• Seasonality – time-dependent transmission rate.

Assuming the disease occurs more often during specific times of the year (ex. December, January, February):

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\beta(t) = \beta o * (1 + b_1 * term(t)),
where \ term(t) = +1 \ for \ t \in (Dec, Jan, Feb),
term(t) = -1 \ for \ t \notin (Dec, Jan, Feb)
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Stochasticity – event-driven approach

"Demographic stochasticity is defined as fluctuations in population processes that arise form the random number of events at the level of the individual" (M.Keeling, P.Rohani, 2008).

• Event driven methods require explicit consideration of events. In the presented model, there are 8 events.

We therefore calculate the probability of:

• Birth

Recovery

Death of infected

- Exposure
- Death of recovered
- Death of susceptibles

Infection

Death of exposed

- Other assumptions
 - Lifetime immunity
 - The Exposed group is not infectious
 - No vaccination
 - Only those at 20+ age group give births
 - > Only those at 20+ age group suffer (natural) deaths
 - > 6-9 age group has the highest value of transmission coefficient (assortative mixing at school combined with probable lack of previous exposure)

3. Mathematical formulation

• SEIR model with four (4) age classes:

We are now dealing with (integer) numbers of S,E,I,R.

SEIR
$$\rightarrow$$
(X,W,Y,Z)

(for simplicity, probability symbols are excluded from the equations)

$$\frac{dX_{i}}{dt} = v_{i} N_{4} - \sum_{j=1}^{c_{a}} \frac{\beta_{ij}(t) Y_{j} X_{i}}{N} - \mu_{i} X_{i}$$

$$\frac{dW_{i}}{dt} = \sum_{j=1}^{c_{a}} \frac{\beta_{ij}(t) Y_{j} X_{i}}{N} - \mu_{i} W_{i} - \sigma W_{i}$$

$$\frac{dY_{i}}{dt} = \sigma W_{i} - \mu_{i} Y_{i} - \gamma Y_{i}$$

$$\frac{dZ_{i}}{dt} = \gamma Y_{i} - \mu_{i} Z_{i}$$

$$where v_{1} = \mu_{4} \neq 0$$

3. Mathematical formulation

SEIR model with four (4) age classes (cont.) :

A fraction of each class moves to the age class above:

$$Q_{1} = Q_{1} - Q_{1}/6$$

$$Q_{2} = Q_{2} + Q_{1}/6 - Q_{2}/4$$

$$Q_{3} = Q_{3} + Q_{2}/4 - Q_{3}/10$$

$$Q_{4} = Q_{4} + Q_{3}/10$$
where $Q_{i} \in (X_{i}, W_{i}, Y_{i}, Z_{i})$

since age groups are 0-5, 6-9, 10-19, 20+ years

3. Mathematical formulation

• SEIR model with four (4) age classes (cont.):

Additionally, the transmission coefficient is now a transmission matrix, where bij is the transmission coefficient for Xi's meeting Yi's.

$$\beta_o = \begin{bmatrix} b_{11} & b_{12} & b_{13} & b_{14} \\ b_{21} & b_{22} & b_{23} & b_{24} \\ b_{31} & b_{32} & b_{33} & b_{34} \\ b_{41} & b_{42} & b_{43} & b_{44} \end{bmatrix}$$

Reminding

$$\beta(t) = \begin{cases} \beta o * (1+b_1), t \in (Dec, Jan, Feb) \\ \beta o * (1-b_1), t \notin (Dec, Jan, Feb) \end{cases}$$

4. Modeling approach – tau-leap method

- Gillespie's "τ-leap method" (Gillespie, 2001)
 - → Discrete-time simulation.
 - \rightarrow 1. Time increment δt small and fixed
 - → 2. Mi number of (ex.) transmission events by time t
 - → 3. Define $\delta M_i = Mi(t+\delta t) Mi(t)$ (i = events of the model, here transmission events), then

$$P(\delta M_T = 1 | X, Y) = \frac{\beta XY}{N} \delta t$$
, $N = total population$

defines the transition probability for transmission events occurring in the time interval δt .

4. Modeling approach – tau-leap method (cont.)

- Gillespie's "τ-leap method" (Gillespie, 2001) (cont.)
 - \rightarrow 4. For δ t small, δ MT is approximately Poisson :

$$\delta M_T \approx Poisson(\frac{\beta XY}{N}\delta t)$$

→ 5. Variables can be updated:

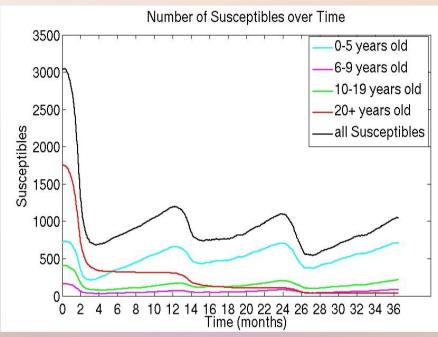
$$X(t+\delta t) = X(t) - \delta M_T \qquad Y(t+\delta t) = Y(t) + \delta M_T$$

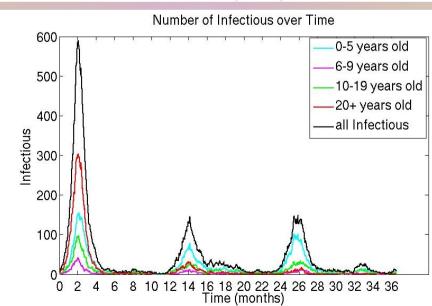
- \rightarrow 6. Time is updated, $t = t + \delta t$. Return to 4.
- For all 8 events in the presented model we assume Poisson distributions with appropriate parameters.

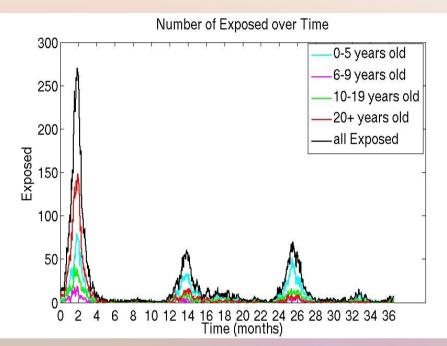
5. Parameters initialization

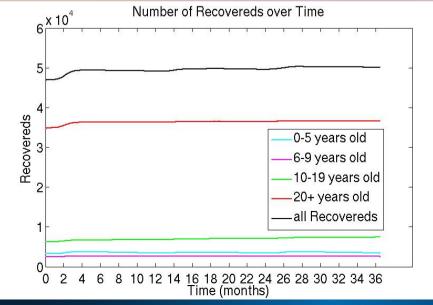
- $\sigma = 1/4$, meaning 4 days average latent (incubation) period
- $\gamma = 1/10$, meaning 10 days average infectious period
- $\mu_4 = v_1 = 0.02$ average birth and death rates per year
- $b_0 \approx 0.5 \pm 0.25$ average values in the transmission matrix, \pm seasonality (b1)
- N=50000 total population
- $\delta t = \tau = 1 \text{ day}$
- time simulated = 3 years
- Starting point 5 infectious and 5 exposed in every age group
- Life expectancy 75 years

6. Results

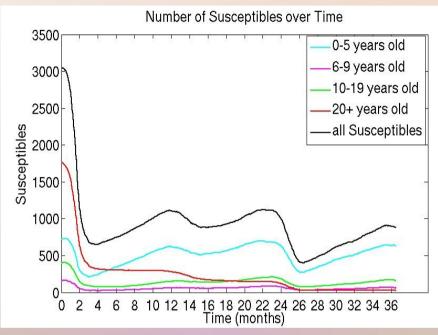


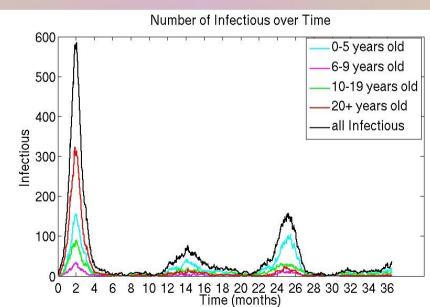


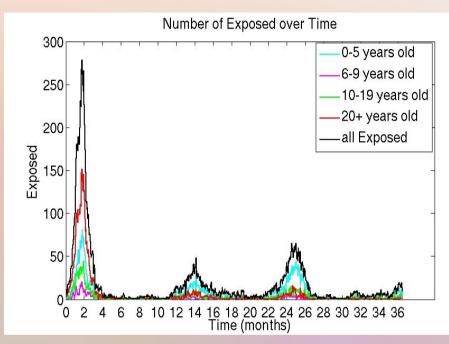


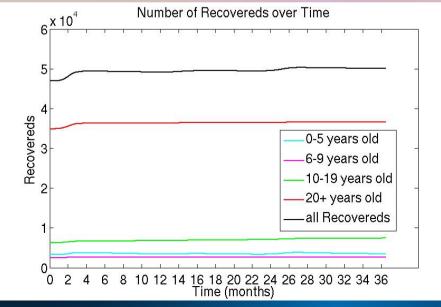


6. Results (cont.)

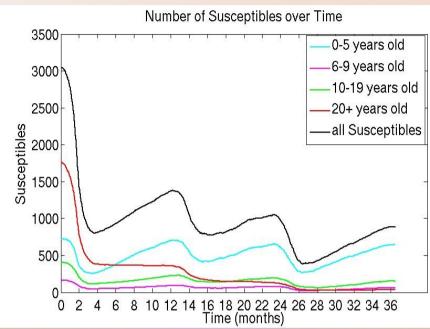


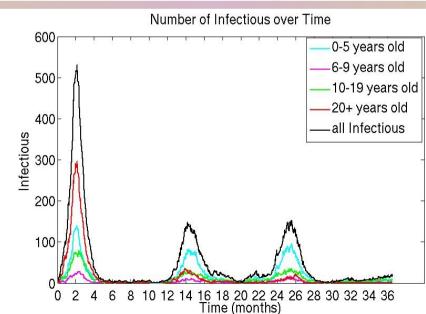


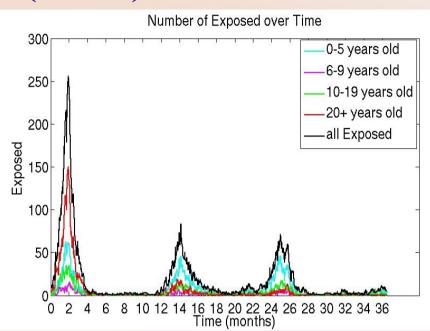


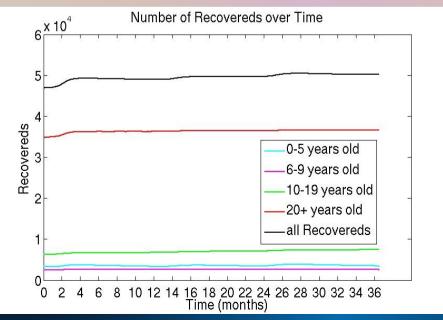


6. Results (cont.)

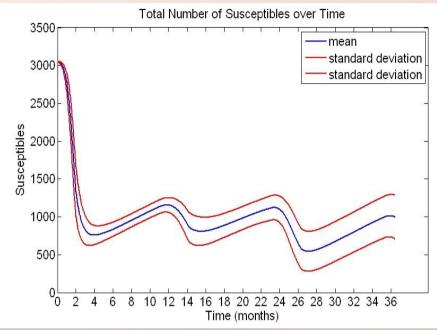


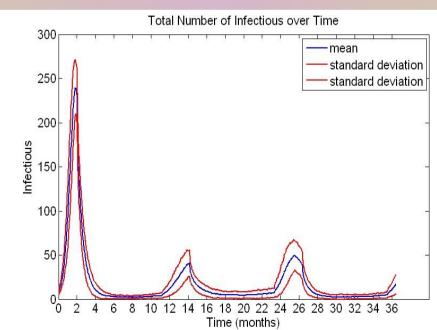


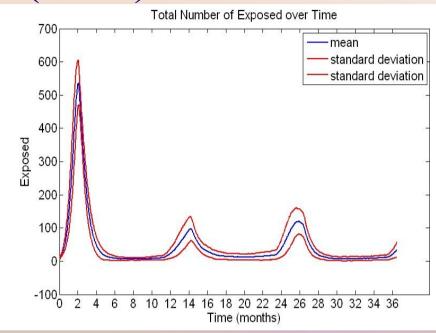


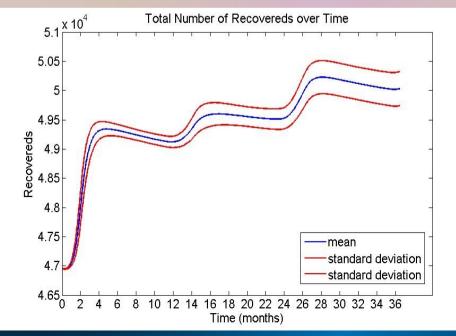


6. Results (cont.)









7. Extensions for the model – Future work

- Ro and Sensitivity analysis (identifying which parameters are important for the prediction imprecision)
- Spatial model
- Controlling the disease Vaccination
- Waning or totally absent immunity

Thank you!

Questions?