Modeling Epidemics: Introduction

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First Models

- Preliminary goal: Model the spread of an infectious (contagious) illness through a population.
- Simplifying assumptions:
 - The total population *N* is constant in time.
 - A newly infected person becomes infectious the next day and remains infectious forever.
 - Each infectious person is equally likely (probability *p*) to infect each noninfectious person on a given day.
- Let *l*(*t*) be the number of infectious people at the start of day *t*.

Stochastic Model

- Number the people from 1 to *N*.
- Let x_n(t) be the infectious status (1 if infectious, 0 if not) of person n at the start of day t.
- We can simulate a possible spread of the illness with the following program ("rand"= random number):

```
for t=1:T-1 for n=1:N let x(n,t+1)=x(n,t) for m=1:N if x(m,t)=1 and rand<p, then let x(n,t+1)=1 end end end
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Simulation Results

- Notice that $I(t) = \sum_{n=1}^{N} x_n(t)$.
- Here are the results of a simulation with $p = 10^{-4}$, N = 1000, and I(1) = 10:



Simulation Results

 And here are the results of three different simulations with p = 10⁻⁴, N = 1000, and I(1) = 10:



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Simulation Results

 Finally, here are the results of three different simulations with p = 10⁻⁴, N = 1000, and I(1) = 1:



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Expected (Average) Daily Outcome

- Let's determine the expected number of people infected on a day *t* that starts with I(t) infectious people and N I(t) who are susceptible to infection.
- A susceptible person *n* has probability 1 − *p* of NOT being infected on day *t* by a given infectious person *m*. Therefore, person *n* has probability (1 − *p*)^{*l*(*t*)} of NOT being infected on day *t*.
- The expected number of people who are infected on day t is then [1 - (1 - p)^{l(t)}][N - l(t)], so

 $E[I(t+1)] = I(t) + [1 - (1 - p)^{I(t)}][N - I(t)]$

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Deterministic Models

• If both I(t) and N - I(t) are large enough, it may be reasonable to approximate I(t + 1) by its expected value, resulting in a deterministic model:

 $I(t+1) = I(t) + [1 - (1 - p)^{I(t)}][N - I(t)]$ (1)

• If pl(t) is small, we can approximate $(1 - p)^{l(t)}$ by 1 - pl(t), yielding a simpler model:

$$I(t+1) = I(t) + \rho I(t)[N - I(t)]$$
(2)

• For these models, given *l*(1) we can compute *l*(2), *l*(3),

Deterministic versus Stochastic

- These deterministic models are much more efficient to compute (1 calculation versus N^2 for the stochastic model). Their predictions may be just as reasonable as any particular simulation of the stochastic model.
- The stochastic model can give some idea of the uncertainty of its predictions via multiple simulations; the deterministic models we've written down say nothing about their uncertainty.

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Continuous-Time Model

- The models we have discussed so far are called discrete-time models; time *t* takes on only integer values.
- We can approximate these models by continuous-time processes; approximating model (2), we get

$$l'(t) = \rho l(t)[N - l(t)]$$
 (3)

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• We can write down an exact solution to this differential equation:

$$I(t) = \frac{NI(0)}{I(0) + [N - I(0)]e^{-pNt}}$$

Fitting the Model to Data

- The solution *I*(*t*) of model (3) has three parameters:
 N, *p*, and *I*(0). Suppose we know *N* but not the other two parameters. Given a set of data points [*t_j*, *l_j*], we can ask which values of *p* and *I*(0) best fit the data.
- [A more fundamental (but more difficult) question is whether the model can adequately fit the data at all; are there ANY parameters of the model that fit the data reasonably well?]
- We could try to minimize the sum of the squares of the residuals *I_j* - *I*(*t_j*). However, this would be a NONlinear least squares problem, because *I*(*t*) is not a linear function of *p* or *I*(0).

Way 1 to use Linear Least Squares

 If the data is given at consecutive values of *t*, say *t_j* = *j*, then one approach is to use model (2) and write

$$I(t + 1) - I(t) = \rho I(t)[N - I(t)].$$

The right-hand side is a linear function of the parameter p, and linear least squares yields the value of p that minimizes the sum of the squares of the residuals $I_{j+1} - I_j - pI_j(N - I_j)$.

• This doesn't resolve the question of which value of I(0) to use. If we let $t_0 = 0$ for the first data point, then we could let $I(0) = I_0$. However, this might not be the best choice of I(0) in order to make the residuals $I_j - I(t_j)$ small.

Way 2 to use Linear Least Squares

 Going back to the solution of model (3), we can make a transformation of variables so that the transformed solution does depend linearly on its parameters. First we divide both sides into N and simplify:

 $N/I(t) = 1 + [N/I(0) - 1]e^{-pNt}$

• Next subtract 1 and take the logarithm:

 $\log[N/I(t) - 1] = \log[N/I(0) - 1] - \rho Nt$

• Let $Z(t) = \log[N/I(t) - 1]$; then the model becomes Z(t) = Z(0) - pNt. This is a linear function of the parameters pN and Z(0). One can transform the data to pairs (t_j, Z_j) , use linear least squares to determine values for pN and Z(0), and then solve for p and I(0).

Caveat

- Both ways of using linear least squares transform the model or its solution into a linear relationship between two quantities that can be computed from the data points (*t_j*, *l_j*); in the second way, the model predicts that *Z_j* is a linear function of *t_j*.
- Rather than simply accept the result of the least squares fit, one should graph the predicted relationship (e.g., *Z_j* versus *t_j*) and see if it actually looks linear. This gives some idea of how valid the model is.
- Regardless of how one determines values for *p* and *l*(0), one should also check directly how well the resulting *l*(*t*) fits the data.

More Sophisticated Models

- Let's re-examine the assumptions behind our first models and discuss how to make them more realistic.
- We assumed a fixed population size *N* that was isolated from other sources of the hypothetical illness we modeled.
- We assumed that a single number *p* represents the probability of an infectious person infecting a susceptible person on each day, for each such pair of people.
- A more realistic model would allow *p* to depend on a number of factors.

Modeling the Infection Probability p

- In real life, the infection probability *p* depends on the pair of people. However, introducing an independent probability *p_{mn}* for each pair of people *m* and *n* results in way too many parameters.
- Also, *p* depends on time; for example, day of week.
- Perhaps most importantly, *p* depends on how long the infectious person has had the illness. Typically it peaks a certain amount of time after infection, then decreases to 0.
- To keep the number of parameters manageable, we need to have a model for how *p* depends on these factors.

Compartmental Models

- Many models divide the population into a relatively small number of categories ("compartments") and keep track of the number of people in each compartment.
- Our first deterministic models had two compartments: "susceptible" and "infectuous". We'll call the continuous time model (3) the SI model.
- A widely studied model is the SIR model, which introduces a third compartment: "recovered". People in this category are no longer infectious.
- Other possible compartments can take into account more stages in the progression of the illness, different behavior patterns, different biological characteristics, etc.

Fitting to Data, Revisited

- In our earlier discussion, we assumed that the number of infectious people at a given time could be measured. But how would we ever know this number?
- The number of infectious people is often inferred from data on new diagnoses of the illness. However:
 - Not all people who get the illness see a doctor.
 - Diagnosis may come well after a person becomes infectious.
 - Data is not always reported (e.g., to CDC) promptly or reliably.
- A common problem in modeling is to relate the quantities of interest to the available data.