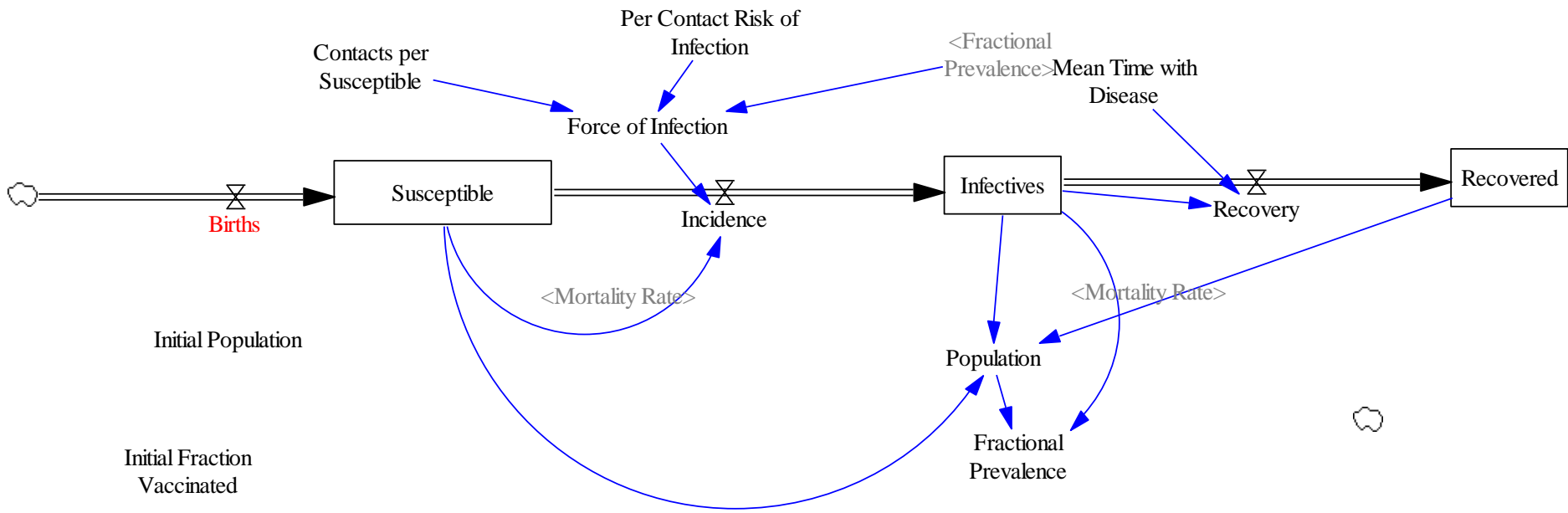


Infectious Disease Models 5 – Basic Epidemiological Quantities and Vaccination

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Shorthand for Key Quantities for Infectious Disease Models: Stocks

- I (or Y): Total number of infectives in population
 - This could be just one stock, or the sum of many stocks in the model (e.g. the sum of separate stocks for asymptomatic infectives and symptomatic infectives)
- N : Total size of population
 - This will typically be the sum of all the stocks of people
- S (or X): Number of susceptible individuals

Intuition Behind Common Terms

- I/N : The Fraction of population members (or, by assumption, contacts!) that are infective
 - Important: Simplest models assume that this is also the fraction of a given susceptible's contacts that are infective! Many sophisticated models relax this assumption
- $c(I/N)$: Number of *infectives* that come into contact with a susceptible in a given unit time
- $c(I/N)\beta$: “Force of infection”: *Likelihood a given susceptible will be infected per unit time*
 - The idea is that if a given susceptible comes into contact with $c(I/N)$ infectives per unit time, and if each such contact gives β likelihood of transmission of infection, then that susceptible has roughly a total likelihood of $c(I/N)\beta$ of getting infected per unit time (e.g. month)

A Critical Throttle on Infection Spread: Fraction Susceptible (f)

- The fraction susceptible (here, S/N) is a key quantity limiting the spread of infection in a population
 - Recognizing its importance, we give this name f to the fraction of the population that is susceptible

Key Term: Flow Rate of New Infections

- This is the key form of the equation in many infectious disease models
- Total # of susceptibles infected per unit time
 - # of Susceptibles * “Likelihood” a given susceptible will be infected per unit time = $S * (\text{“Force of Infection”})$
 $= S(c(I/N)\beta)$
 - Note that this is a term that multiplies both S and I !
 - This is much different than the purely linear terms on which we have previously focused
 - “Likelihood” is actually a likelihood per unit time (e.g. can be >1 – indicating that mean time to infection is <1)

Another Useful View of this Flow

- Recall: Total # of susceptibles infected per unit time = # of Susceptibles * “Likelihood” a given susceptible will be infected per unit time = $S * (\text{“Force of Infection”}) = S(c(I/N)\beta)$
- The above can also be phrased as the following: $S(c(I/N)\beta) = I(c(S/N)\beta) = I(c * f * \beta) =$ # of Infectives * Mean # susceptibles infected per unit time by each infective
- This implies that as # of susceptibles falls \Rightarrow # of susceptibles surrounding each infective falls \Rightarrow the rate of new infections falls (“Less fuel for the fire” leads to a smaller burning rate)

Recall: The Importance of Susceptible Fraction

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Reasons

Critical Notions

- Contact rates & transmission probabilities
- Equilibria
 - Endemic
 - Disease-free
- R_0 , R_*
- Herd Immunity

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Infection

- Recall: For this model, a given infective infects $c(S/N)\beta$ others per time unit
 - This goes up as the number of susceptibles rises
- Questions
 - If the mean time a person is infective is μ , how many people does that infective infect before recovering?
 - With the same assumption, how many people would that infective infect if everyone else is susceptible?
 - Under what conditions would there be more infections after their recovery than before?

Fundamental Quantities

- We have just discovered the values of 2 famous epidemiological quantities for our model
 - Effective Reproductive Number: R_*
 - Basic Reproductive Number: R_0

Effective Reproductive Number: R_*

- Number of individuals infected by an 'index' infective in the current epidemiological context
- Depends on
 - Contact number
 - Transmission probability
 - Length of time infected
 - # (Fraction) of Susceptibles
- Affects
 - Whether infection spreads
 - If $R_* > 1$, # of cases will rise, If $R_* < 1$, # of cases will fall
 - Alternative formulation: Largest real eigenvalue ≤ 0
 - Endemic Rate

Basic Reproduction Number: R_0

- Number of individuals infected by an ‘index’ infective *in an otherwise disease-free equilibrium*
 - This is just R_* at disease-free equilibrium all (other) people in the population are susceptible other than the index infective
- Depends on
 - Contact number
 - Transmission probability
 - Length of time infected
- Affects
 - Whether infection spreads
 - If $R_0 > 1$, Epidemic Takes off, If $R_0 < 1$, Epidemic dies out
 - Alternative formulation: Largest real eigenvalue $< > 0$
 - Initial infection rise $\propto \exp(t^*(R_0-1)/D)$
 - Endemic Rate

Basic Reproductive Number R_0

- If contact patterns & infection duration remain unchanged and if fraction f of the population is **susceptible**, then mean # of individuals infected by an infective over the course of their infection is $f \cdot R_0$
- In endemic equilibrium: $\text{Inflow} = \text{Outflow} \Rightarrow (S/N) \cdot R_0 = 1$
 - Every infective infects a “replacement” infective to keep equilibrium
 - Just enough of the population is susceptible to allow this replacement
 - The higher the R_0 , the lower the fraction of susceptibles in equilibrium!
 - *Generally some* susceptibles remain: At some point in epidemic, susceptibles will get so low that can't spread

Our model

- Set
 - $c=10$ (people/month)
 - $\beta=0.04$ (4% chance of transmission per S-I contact)
 - $\mu=10$
 - Birth and death rate= 0
 - Initial infectives=1, other 1000 susceptible
- What is R_0 ?
- What should we expect to see ?

Thresholds

- R_*
 - Too low # susceptibles $\Rightarrow R^* < 1$: # of infectives declining
 - Too high # susceptibles $\Rightarrow R^* > 1$: # of infectives rising
- R_0
 - $R_0 > 1$: Infection is introduced from outside will cause outbreak
 - $R_0 < 1$: “Herd immunity”: infection is introduced from outside will die out (may spread to small number before disappearing, but in unsustainable way)
 - **This is what we try to achieve by control programs, vaccination, etc.**
- Outflow from susceptibles (infections) is determined by the # of Infectives

Equilibrium Behaviour

- With Births & Deaths, the system can approach an “endemic equilibrium” where the infection stays circulating in the population – but in balance
- The balance is such that (simultaneously)
 - The rate of new infections = The rate of immigration
 - Otherwise # of susceptibles would be changing!
 - The rate of new infections = the rate of recovery
 - Otherwise # of infectives would be changing!

Equilibria

- Disease free
 - No infectives in population
 - Entire population is susceptible
- Endemic
 - Steady-state equilibrium produced by spread of illness
 - Assumption is often that children get exposed when young
- The stability of these equilibria (whether the system departs from them when perturbed) depends on the parameter values
 - For the disease-free equilibrium on R_0

Vaccination

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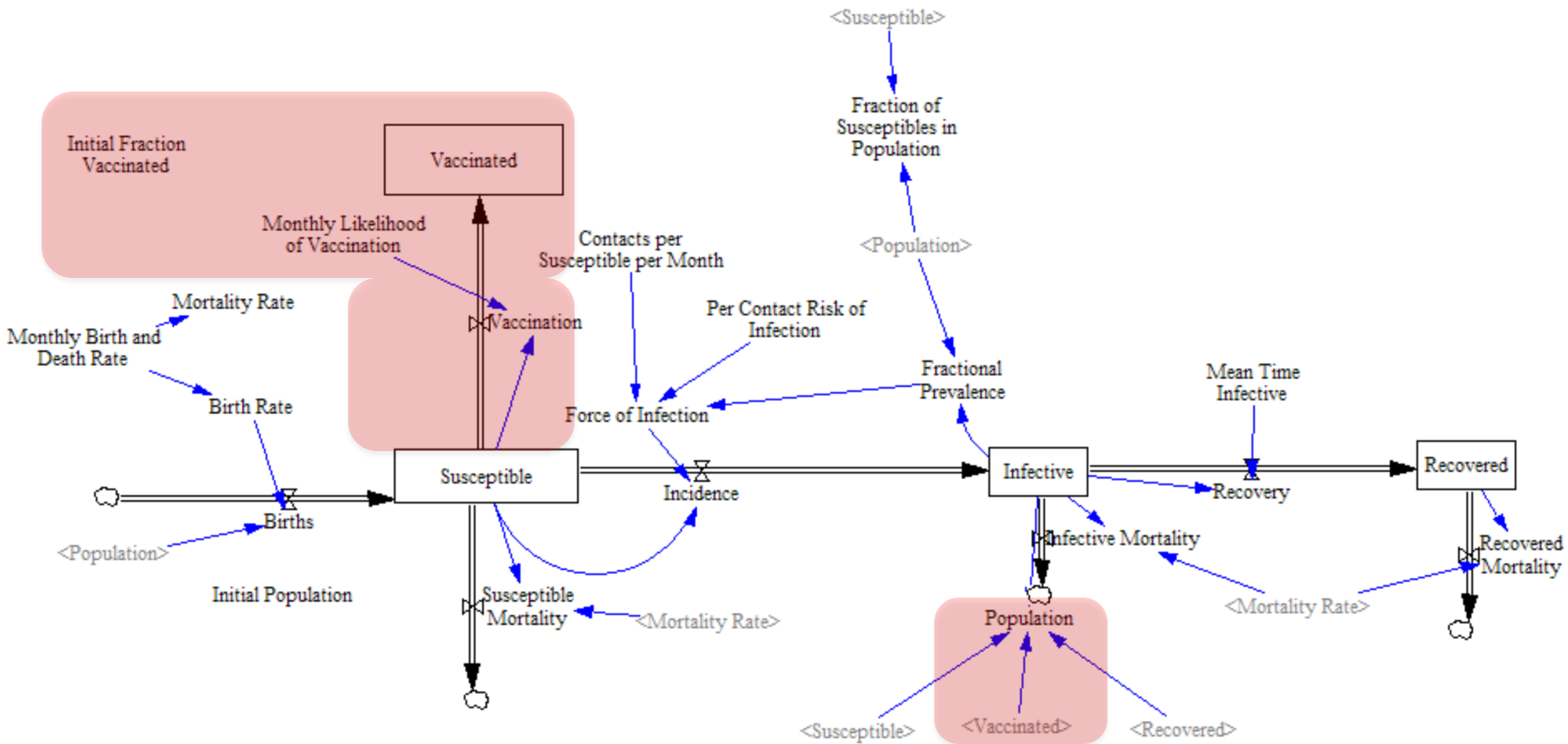
Adding Vaccination Stock

- Add a
 - “Vaccinated” stock
 - A constant called “Monthly Likelihood of Vaccination”
 - “Vaccination” flow between the “Susceptible” and “Vaccinated” stocks
 - The rate is the stock times the constant above
- Set initial population to be divided between 2 stocks
 - Susceptible
 - Vaccinated
- Incorporate “Vaccinated” in population calculation

Additional Settings

- $c = 10$
- $\text{Beta} = .04$
- Duration of infection = 10
- Birth & Death Rate = 0

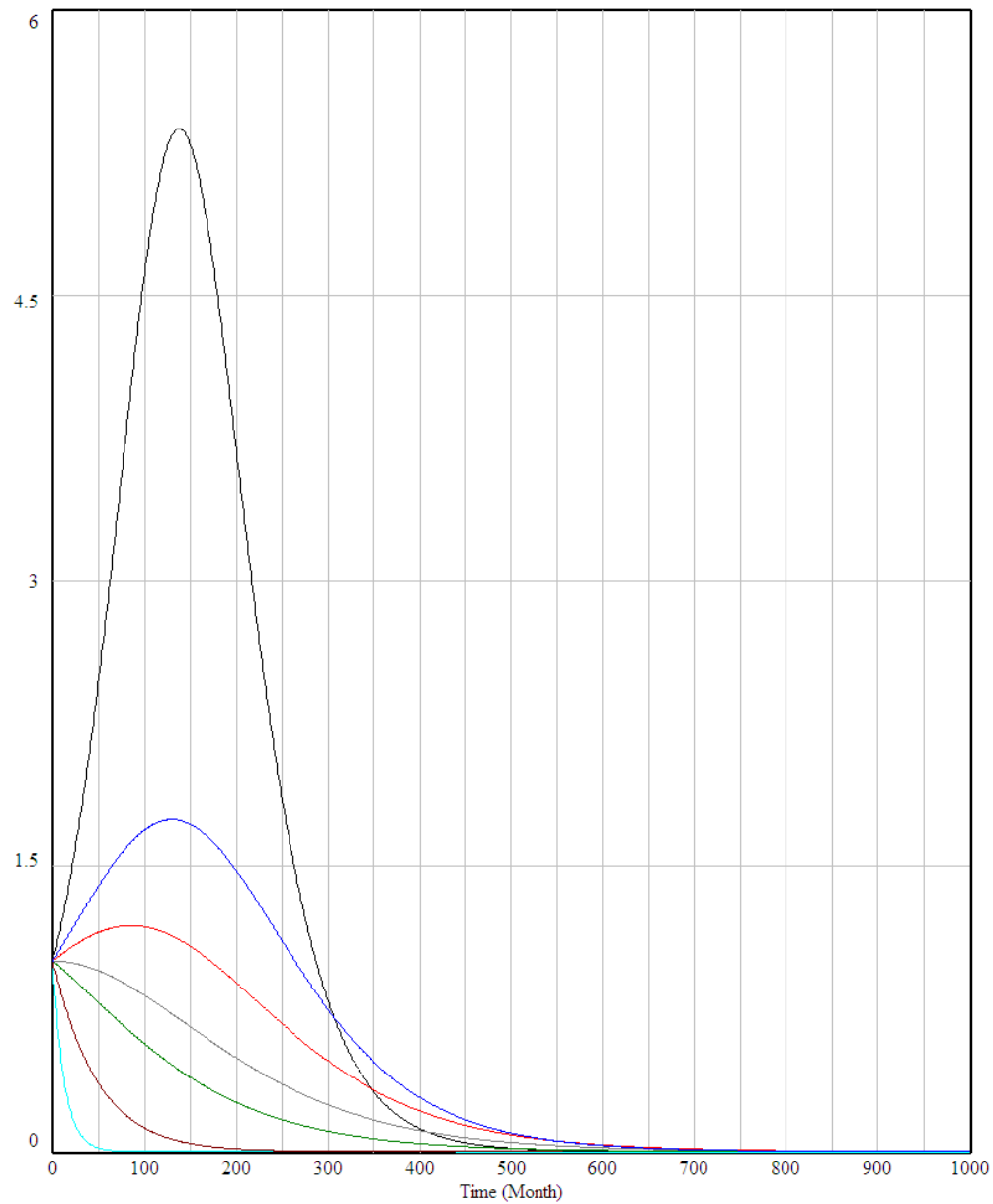
Adding Stock



Experiment with Different Initial Vaccinated Fractions

- Fractions = 0.25, 0.50, 0.6, 0.7, 0.8

Infectives



Infectives : No Immigration Test Fraction Vaccinated= .73
Infectives : No Immigration Test Fraction Vaccinated= .74
Infectives : No Immigration Test Fraction Vaccinated= .76
Infectives : No Immigration Test Fraction Vaccinated= .75
Infectives : No Immigration Test Fraction Vaccinated= .7
Infectives : No Immigration Test Fraction Vaccinated= .8
Infectives : No Immigration Test Fraction Vaccinated= .95

Recall: Thresholds

- R^*
 - Too low # susceptibles $\Rightarrow R^* < 1$: # of infectives declining
 - Too high # susceptibles $\Rightarrow R^* > 1$: # of infectives rising
- Outflow from susceptibles (infections) is determined by the # of Infectives
- Delays:
 - For a while after infectives start declining, they still deplete susceptibles sufficiently for susceptibles to decline
 - For a while after infectives start rising, the # of infections is insufficient for susceptibles to decline

Effective Reproductive Number: R_*

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 - Recognizing its importance, we give this name f to the fraction of the population that is susceptible
- If contact patterns & infection duration remain unchanged and, then mean # of individuals infected by an infective over the course of their infection is $f * R_0$

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Critical Immunization Threshold

- Consider an index infective arriving in a “worst case” scenario when no one else in the population is infective or recovered from the illness
 - In this case, that infective is most “efficient” in spreading
- The goal of vaccination is keep the fraction susceptible low enough that infection cannot establish itself even in this worst case
 - We do this by administering vaccines that makes a person (often temporarily) immune to infection
- We say that a population whose f is low enough that it is resistant to establishment of infection exhibits “herd immunity”

Critical Immunization Threshold

- Vaccination seeks to lower f such that $f \cdot R_0 < 1$
- *Worst case: Suppose we have a population that is divided into immunized (vaccinated) and susceptible*
 - Let q_c be the critical fraction immunized to stop infection
 - *Then $f = 1 - q_c$, $f \cdot R_0 < 1 \Rightarrow (1 - q_c) \cdot R_0 < 1 \Rightarrow q_c > 1 - (1/R_0)$*
- So if $R_0 = 4$ (as in our example), $q_c = 0.75$ (i.e. 75% of population must be immunized – just as we saw!)