Heterogeneity, contact patterns and modeling options

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### MMED 2016 http://www.ici3d.org/mmed/

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# The resilience of infectious disease

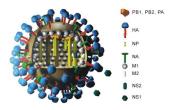
1967: It's time to close the book on infectious diseases







# Pathogen evolution





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# Human heterogeneity





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# Human heterogeneity





# Outline

Homogeneous disease models

The importance of heterogeneity

Effects of heterogeneity

Modeling approaches



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Homogeneous models assume everyone has the same:

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  - disease characteristics (e.g. susceptibility, tendency to transmit)

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mixing rate

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  - disease characteristics (e.g. susceptibility, tendency to transmit)

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- mixing rate
- probability of mixing with each person

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R<sub>0</sub> is the number of people who would be infected by an infectious individual *in a fully susceptible population.* 

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$$\blacktriangleright \ \mathcal{R}_0 = \beta / \gamma = \beta D = (cp)D$$

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c: Contact Rate

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- c: Contact Rate
- p: Probability of transmission (infectivity)

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- c: Contact Rate
- p: Probability of transmission (infectivity)
- D: Average duration of infection

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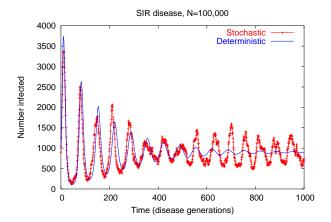
- c: Contact Rate
- p: Probability of transmission (infectivity)
- D: Average duration of infection
- A disease can invade a population if and only if  $\mathcal{R}_0 > 1$ .

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# Equilibrium

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# Equilibrium



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*R<sub>eff</sub>* is the number of people who would be infected by an infectious individual *in a general population*.

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• At equilibrium: 
$$\mathcal{R}_{eff} = \mathcal{R}_0 \frac{S}{N} = 1$$
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• Thus: 
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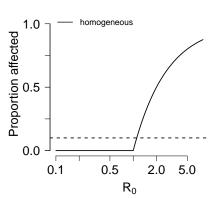
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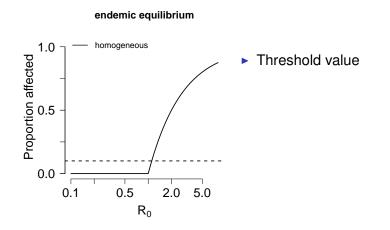
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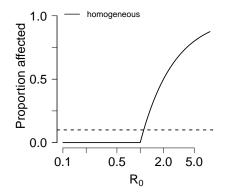
• Proportion 'affected' is  $V = 1 - S/N = 1 - 1/R_0$ .



endemic equilibrium

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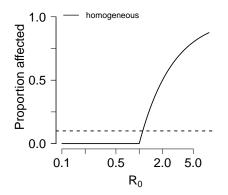


#### endemic equilibrium

- Threshold value
- Sharp response to changes in factors underlying transmission

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#### endemic equilibrium

- Threshold value
- Sharp response to changes in factors underlying transmission

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Works – sometimes

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► For diseases with no recovery, V is the disease prevalence

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- ► For diseases with no recovery, V is the disease prevalence
- For other diseases, the equilibrium value of P = I/N will be equal to V times the ratio of time spent sick to the time spent immune.

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- ► For diseases with no recovery, *V* is the disease prevalence
- For other diseases, the equilibrium value of P = I/N will be equal to V times the ratio of time spent sick to the time spent immune.

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Example: measles before vaccination

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Example: measles before vaccination

▶ *V* = 0.95

## Proportion affected and disease prevalence

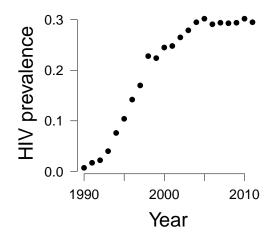
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Example: measles before vaccination

• 
$$\bar{P} = 0.95 \times (2 \text{wk}/60 \text{yr}).$$

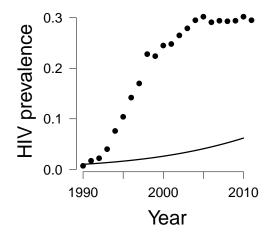
#### **Disease dynamics**



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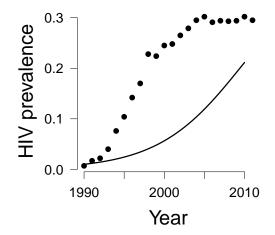


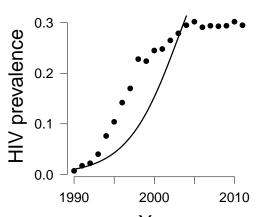
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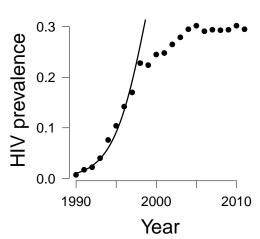




R0 = 4.00

Year

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R0 = 5.66

#### Outline

Homogeneous disease models

The importance of heterogeneity

Effects of heterogeneity

Modeling approaches



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Flavors of heterogeneity



Flavors of heterogeneity

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among hosts

Flavors of heterogeneity

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- among hosts
- spatial

#### Flavors of heterogeneity

- among hosts
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- demographic (discreteness of indviduals)

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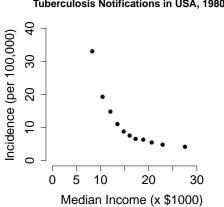
temporal

#### Flavors of heterogeneity

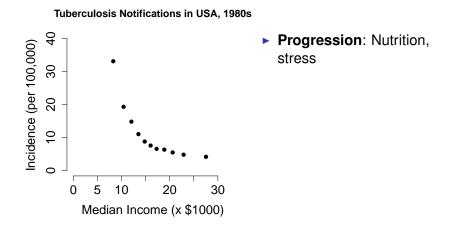
- among hosts
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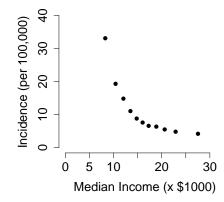
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- temporal
- others



**Tuberculosis Notifications in USA, 1980s** 



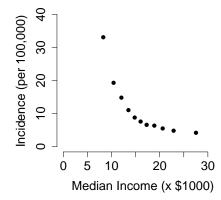


- Tuberculosis Notifications in USA, 1980s
- Progression: Nutrition, stress

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 Contact: Overcrowding, poor ventilation



- **Tuberculosis Notifications in USA, 1980s**
- Progression: Nutrition, stress

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- Contact: Overcrowding, poor ventilation
- Cure: Access to medical care

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STDs: Sexual mixing patterns, access to medical care

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STDs: Sexual mixing patterns, access to medical care

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Influenza: Crowding, nutrition

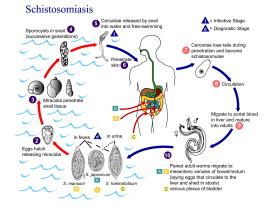
- STDs: Sexual mixing patterns, access to medical care
- Influenza: Crowding, nutrition
- Malaria: Attractiveness to biting insects, geographical location, immune status

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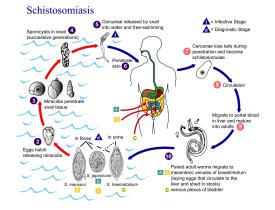
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Every disease!

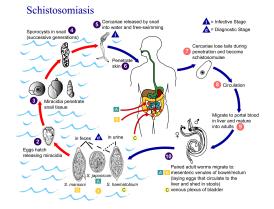


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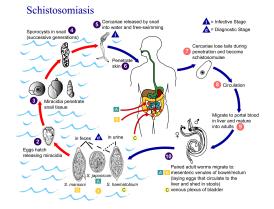


For schistosomiasis, the worldwide average  $\mathcal{R}_0 < 1$ 

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- Disease persists because of specific populations with  $\mathcal{R}_0 > 1$ .



- For schistosomiasis, the worldwide average  $\mathcal{R}_0 < 1$
- Disease persists because of specific populations with  $\mathcal{R}_0 > 1$ .
- This effect operates at many scales.

#### Outline

Homogeneous disease models

The importance of heterogeneity

Effects of heterogeneity

Modeling approaches



Assume p = στ has a susceptibility component and a transmission component:

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• 
$$\mathcal{R}_0 = \sigma \tau c D$$

Assume p = στ has a susceptibility component and a transmission component:

- $\mathcal{R}_0 = \sigma \tau c D$
- $\mathcal{R}_{eff} = \sigma \tau c DS / N$

•  $\tau D$  applies to infectious individuals  $\rightarrow \tau_I D_I$ 

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•  $\sigma$  applies to susceptible individuals  $\rightarrow \sigma_S$ 

- $\tau D$  applies to infectious individuals  $\rightarrow \tau_I D_I$
- $\sigma$  applies to susceptible individuals  $\rightarrow \sigma_S$

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• c is complicated  $\rightarrow c_S c_I / \bar{c}$ 



 Imagine a disease spread by people who differ only in their effective mixing rates

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- Imagine a disease spread by people who differ only in their effective mixing rates
- If the disease has just started spreading in the population, how do c<sub>S</sub> and c<sub>l</sub> compare to c̄?

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•  $c_S \approx \overline{c}; c_l > \overline{c}.$ 

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If the disease is very widespread in the population?

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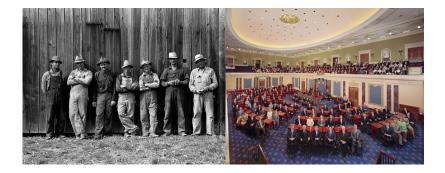
•  $c_S \approx \bar{c}; c_l > \bar{c}.$ 

If the disease is very widespread in the population?

►  $C_S < \overline{C}; C_I \rightarrow \overline{C}.$ 







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What happens when a peanut farmer is elected to the US Senate?



What happens when a peanut farmer is elected to the US Senate?

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The average IQ goes up in both places!

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When the disease invades:

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  - $\blacktriangleright$  The susceptible population  $\approx$  the general population

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- When the disease invades:
  - The susceptible population  $\approx$  the general population
  - The infectious population is likely to have higher values of c, D and/or \(\tau\)

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- When the disease invades:
  - The susceptible population  $\approx$  the general population
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*R*<sub>0</sub> is typically greater than you would expect from a homogeneous model

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As disease prevalence goes up:



- As disease prevalence goes up:
  - Susceptible pool is the most resistant, or least exposed group

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- As disease prevalence goes up:
  - Susceptible pool is the most resistant, or least exposed group

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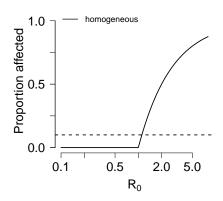
 Infectious pool moves looks more like the general population.

- As disease prevalence goes up:
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- Infectious pool moves looks more like the general population.
- ▶ → lower proportion affected for a given value of  $\mathcal{R}_0$ .

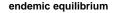
### Homogeneous endemic curve

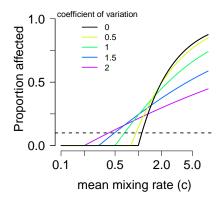


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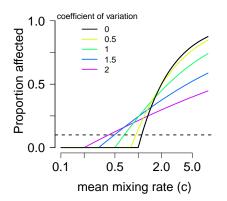
endemic equilibrium

### Heterogeneous endemic curves





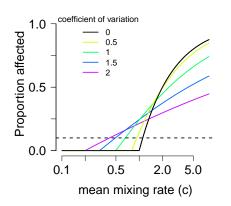
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#### endemic equilibrium

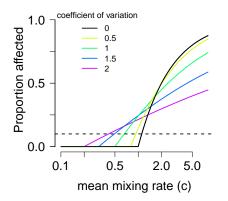


#### endemic equilibrium

### Heterogeneity has a double-edged effect

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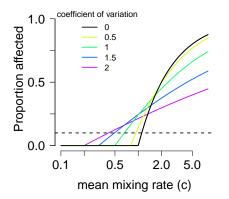


#### endemic equilibrium

- Heterogeneity has a double-edged effect
  - Effects of disease are *lower* for a given value of R<sub>0</sub>.

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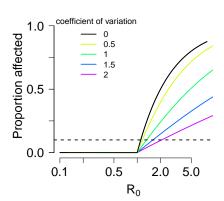
#### endemic equilibrium

- Heterogeneity has a double-edged effect
  - Effects of disease are *lower* for a given value of R<sub>0</sub>.
  - But R<sub>0</sub> is *higher* for given mean values of factors underlying transmission

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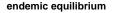
### Heterogeneous endemic curves

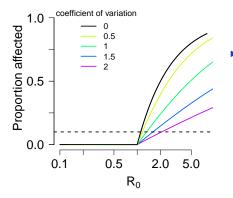


#### endemic equilibrium

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## Heterogeneous endemic curves





 Heterogeneity makes the endemic curve flatter

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Diseases that invade have high values of R<sub>0</sub>

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• Diseases that invade have high values of  $\mathcal{R}_0$ 

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•  $\mathcal{R}_{eff}$  must be 1 at equilibrium

- Diseases that invade have high values of R<sub>0</sub>
- $\mathcal{R}_{eff}$  must be 1 at equilibrium
  - Potentially infectious contacts are wasted

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Those not affected less susceptible than average

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- Those not affected less susceptible than average
- Infectious pool less infectious

### Spatial and network models

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 Individual-level, or spatial, heterogeneity also usually increases wasted contacts

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Infectious people meet:

 Individual-level, or spatial, heterogeneity also usually increases wasted contacts

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- Infectious people meet:
  - people with similar social backgrounds

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- Infectious people meet:
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  - people with similar behaviours

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  - people with similar social backgrounds
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  - people who are nearby geographically or in the contact network

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More wasted contacts further flatten the endemic curve

### Outline

Homogeneous disease models

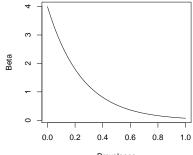
The importance of heterogeneity

Effects of heterogeneity

Modeling approaches

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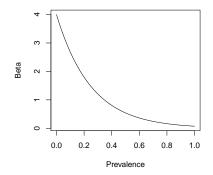
# Phenomenological



Prevalence

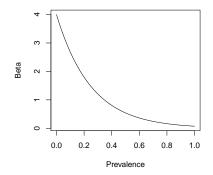
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### Phenomenological



• You can simply *make*  $\beta$  go down as prevalence goes up

### Phenomenological



- You can simply make  $\beta$  go down as prevalence goes up
  - Need to choose a functional form

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Divide the population into groups.



### Divide the population into groups.

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cities and villages

### Divide the population into groups.

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- cities and villages
- rich and poor

#### Divide the population into groups.

- cities and villages
- rich and poor
- high and low sexual activity

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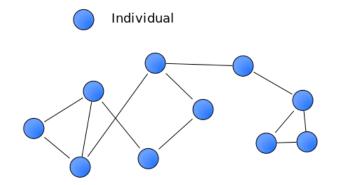
age, gender

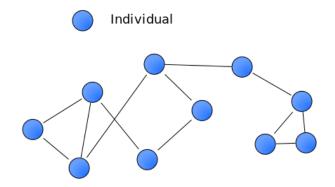
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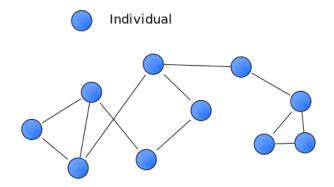
- age, gender
- ▶ ...





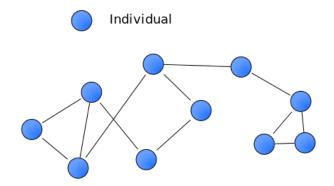
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Allow many possibilities:



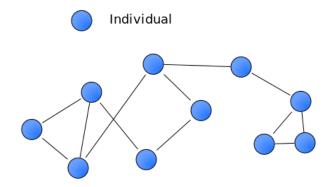
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- Allow many possibilities:
  - vary individual characteristics



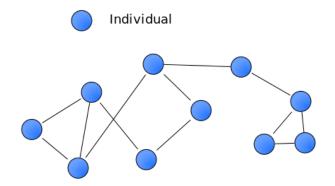
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- Allow many possibilities:
  - vary individual characteristics
  - add a network of interactions



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- Allow many possibilities:
  - vary individual characteristics
  - add a network of interactions
  - let the network change



- Allow many possibilities:
  - vary individual characteristics
  - add a network of interactions
  - let the network change
- Individual-based approaches require stochastic models

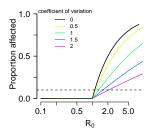
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# Summary



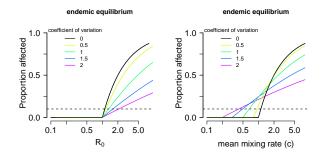
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#### endemic equilibrium



### Summary





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