

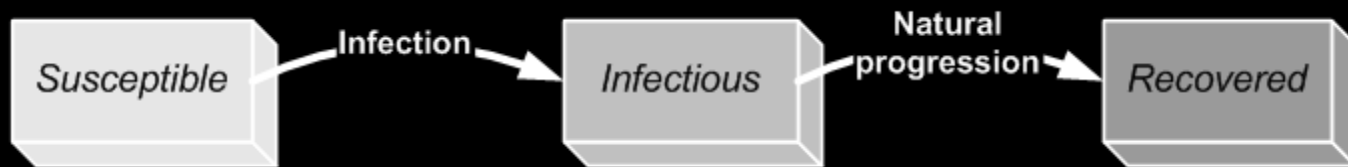
Markov chain and state transition models

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Moving between disease states in a Reed-Frost model

- The transition from one disease state to another is a linear process:

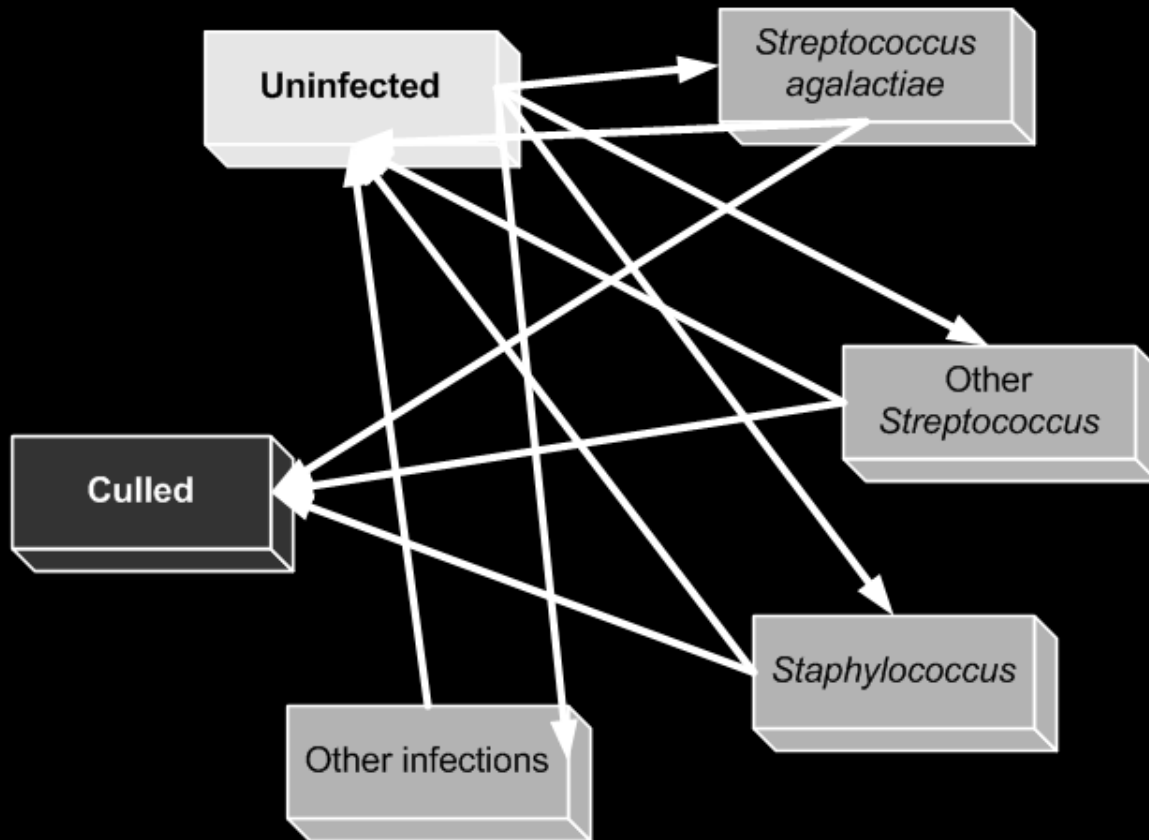


- The number of individuals in each state during a particular time period is dependent only on the number of individuals in each state during the previous time period
- “Lags” can be introduced to alter the timing, but this basic principle still applies

Moving between disease states in life

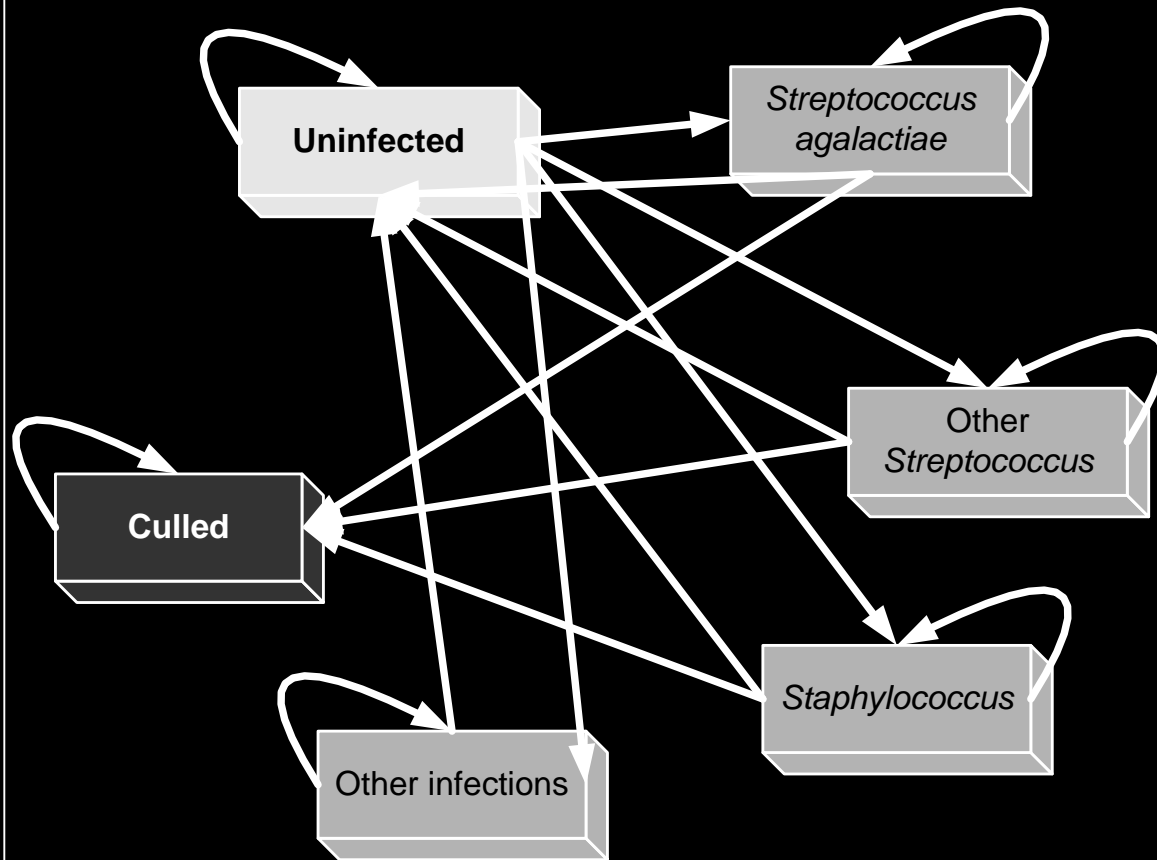
- In more complex (e.g., real) systems, the transition between disease states is not necessarily linear
 - Chronic conditions
- The Reed-Frost framework does not account for the effects of interventions
 - Culling
 - Vaccination

Example: A state transition diagram for a model of mastitis in cattle (I)



- The arrows represent all of the possible (or at least all of the important) transitions between states
- As in Reed-Frost and the differential calculus models, an individual can have only one state per time period

Example: A state transition diagram for a model of mastitis in cattle (II)



- Individuals can also remain in a state for more than one time period
- It is more convenient to illustrate a state transition model as a table, or a *matrix*

Example: A state transition matrix for a model of foot-and-mouth disease

FROM	TO			
	<i>Susceptible</i>	<i>Infectious</i>	<i>Immune</i>	<i>Removed</i>
<i>Susceptible</i>	Remaining susceptible	Infection	Effective vaccination	Preemptive culling
<i>Infectious</i>	--	--	Natural immunity	Slaughter of affected herds
<i>Immune</i>	Waning immunity	--	Remaining immune	--
<i>Removed</i>	--	--	--	Once removed, an individual remains removed

- Each cell shows the pathway or pathways by which individuals transition between states
- These pathways can be expressed as probabilities

Example: Representing an *SLIR* model in a state transition matrix



	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	$1 - P(\text{infection})$	$P(\text{infection})$	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- The value in each cell represents the probability that an individual will move from $State_{Row}$ to $State_{Column}$

Example: A state transition matrix for mastitis

	<i>Uninfected</i>	<i>Strep ag.</i>	<i>Strep. other</i>	<i>Staph.</i>	<i>Other</i>	<i>Culled</i>
<i>Uninfected</i>	0.52	0.03	0.02	0.12	0.02	0.29
<i>Strep. ag.</i>	0.7	0.01	0	0	0	0.29
<i>Strep. other</i>	0.7	0	0.01	0	0	0.29
<i>Staph.</i>	0.1	0	0	0.4	0	0.5
<i>Other</i>	0.71	0	0	0	0	0.29
<i>Culled</i>	0	0	0	0	0	1

- Note that the sum across each row is 1: every “possible” transition is accounted for

Applying a state transition matrix in a disease model: *SLIR*

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- Let's start with a population of 100 susceptible individuals

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0

- What does the model predict for time period 1?

Susceptible individuals in time period 1 (SLIR)

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- $S_1 = (S_0 \times P_{SS}) + (L_0 \times P_{LS}) + (I_0 \times P_{IS}) + (R_0 \times P_{RS})$
- $S_1 = (100 \times 0.7) + (0 \times 0) + (0 \times 0) + (0 \times 0)$
- $S_1 = 70$

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0
1	70			

Latent individuals in time period 1 (SLIR)

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- $L_1 = (S_0 \times P_{SL}) + (L_0 \times P_{LL}) + (I_0 \times P_{IL}) + (R_0 \times P_{RL})$
- $L_1 = (100 \times 0.3) + (0 \times 0) + (0 \times 0) + (0 \times 0)$
- $L_1 = 30$

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0
1	70	30	0	0

Susceptible individuals in time period 2 (SLIR)

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- $S_2 = (S_2 \times P_{SS}) + (L_2 \times P_{LS}) + (I_2 \times P_{IS}) + (R_2 \times P_{RS})$
- $S_2 = (70 \times 0.7) + (30 \times 0) + (0 \times 0) + (0 \times 0)$
- $S_2 = 49$

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0
1	70	30	0	0
2	49			

Latent individuals in time period 2 (SLIR)

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- $L_2 = (S_2 \times P_{SL}) + (L_2 \times P_{LL}) + (I_2 \times P_{IL}) + (R_2 \times P_{RL})$
- $L_2 = (70 \times 0.3) + (30 \times 0) + (0 \times 0) + (0 \times 0)$
- $L_2 = 21$

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0
1	70	30	0	0
2	49	21		

Infectious individuals in time period 2 (SLIR)

	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
<i>Susceptible</i>	0.7	0.3	0	0
<i>Latent</i>	0	0	1	0
<i>Infectious</i>	0	0	0	1
<i>Recovered</i>	0	0	0	1

- $I_2 = (S_2 \times P_{SI}) + (L_2 \times P_{LI}) + (I_2 \times P_{II}) + (R_2 \times P_{RI})$
- $I_2 = (70 \times 0) + (30 \times 1) + (0 \times 0) + (0 \times 0)$
- $I_2 = 21$

	<i>Number of individuals in each state</i>			
<i>Time period</i>	<i>Susceptible</i>	<i>Latent</i>	<i>Infectious</i>	<i>Recovered</i>
0	100	0	0	0
1	70	30	0	0
2	49	21	30	0

A more complex state transition model: Mastitis

	<i>Uninfected</i>	<i>Strep ag.</i>	<i>Strep. other</i>	<i>Staph.</i>	<i>Other</i>	<i>Culled</i>
<i>Uninfected</i>	0.52	0.03	0.02	0.12	0.02	0.29
<i>Strep. ag.</i>	0.7	0.01	0	0	0	0.29
<i>Strep. other</i>	0.7	0	0.01	0	0	0.29
<i>Staph.</i>	0.1	0	0	0.4	0	0.5
<i>Other</i>	0.71	0	0	0	0	0.29
<i>Culled</i>	0	0	0	0	0	1

$$U_1 = (803 \times 0.52) + (52 \times 0.7) + (52 \times 0.7) + (79 \times 0.1) + (14 \times 0.71) + (0 \times 0) = 508$$

$$Sa_1 = (803 \times 0.03) + (52 \times 0.01) + (52 \times 0) + (79 \times 0) + (14 \times 0) + (0 \times 0) = 25$$

<i>Time period</i>	<i>Uninfected</i>	<i>Strep ag.</i>	<i>Strep. other</i>	<i>Staph.</i>	<i>Other</i>	<i>Culled</i>
0	803	52	52	79	14	0
1	508	25	17	128	16	305
<i>etc.</i>						

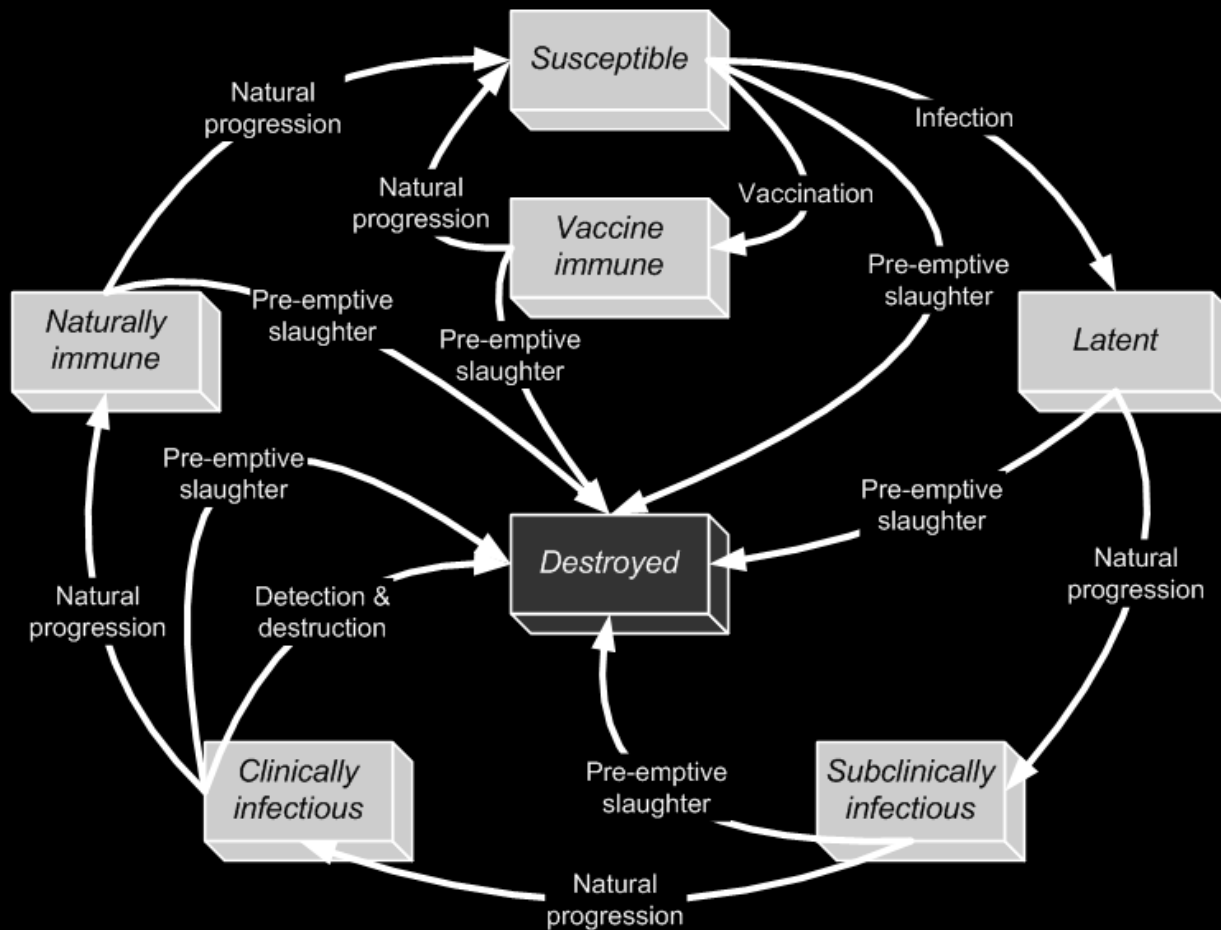
An aside: Markov chain vs. state transition models

- A Markov chain is a special case of a state transition model
 - In a Markov chain model, the transition probabilities are fixed
 - Markov chain models can be useful for modeling endemic diseases (as in the mastitis example)
- In more complex state transition models (“modified Markov chains”), the transition probabilities can change over time
 - For example, the probability of infection may depend on the number of cases in the previous time period
 - Does this look familiar?

Monte Carlo state transition models

- We've seen that Monte Carlo methods can be used to allow probabilities in a model to vary
- There are a lot of probabilities in a state transition matrix
- There is a great deal of potential for the use of Monte Carlo methods in a state transition model

Preview of coming attractions



- *NAADSM* is a framework for sophisticated state transition models which make extensive use of Monte Carlo methods

Recommended reading

- Carpenter, T.E., 1988. Microcomputer programs for Markov and modified Markov chain disease models. *Preventive Veterinary Medicine* 5: 169–179. (*A very nice demonstration of state transition modeling with a spreadsheet.*)
- Miller, W.M., 1976. A state-transition model of epidemic foot-and-mouth disease. In Ellis, P.R., Shaw, A.P.M., and Stephens, A.J., eds. *New Techniques in Veterinary Epidemiology and Economics, Proceedings of a Symposium*, University of Reading, England: ISVEE I.
<http://www.sciquest.org.nz> (*An early application of a state transition model for animal disease.*)

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- Harvey, N., Reeves, A., Schoenbaum, M.A., Zagmutt-Vergara, F.J., Dubé, C., Hill, A.E., Corso, B.A., McNab, W.B., Cartwright, C.I., and Salman, M.D., 2007. The *North American Animal Disease Spread Model*: A simulation model to assist decision making in evaluating animal disease incursions. *Preventive Veterinary Medicine*, in press.
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