# Estimation and statistical considerations

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### Overview

- Statistical background
- Study design issues





## Prevalence vs incidence

#### Cross sectional

- Prevalence plus life tables
- Sullivan's method

#### Longitudinal – uses transitions

- Incidence
- Remission

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- Case fatality



# Longitudinal methods

 Double decrement - Non-reversible processes only Multi-state methods – Multi-state life tables Increment decrement life tables – Multi-state models Microsimulation

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## Multi-state models

- Model discrete state of health
- Discrete time
  - Small fixed intervals
  - Simplifies algebra
- Continuous time
  - Time dependent transition intensities

### Allows covariates





#### Multi state models







# Modelling framework (discrete)

- Probability of being in state k at a longitudinal time point given you were in a state j initially
- Modelled using multivariate logistic model
- Regression smoothed transitions
- Markov assumption (history of process not important)





Modelling framework (discrete) Individual *i* in state *j* at time  $x_i$ and state k after time h then  $p_{ik} = \Pr(Y_{t+h} = k \mid Y_t = j)$  $\log\left(\frac{p_{jk}(x_i)}{p_{il}(x_i)}\right) = a_{jk} + b_{jk}x_i$ 





Life expectancies (discrete)  $e^{jk}_{z_0} = \sum_{y=1}^{\infty} P(X_{(x_i+h)} = k \mid X_{x_i} = j, z_0)$ 

Life expectancy of state jk is the sum of the probability of transitions given covariates

$$e^{j} = \pi_{z_{0}}^{1} e_{z_{0}}^{j1} + \pi_{z_{0}}^{2} e_{z_{0}}^{j2}$$

Total life expectancy for each state j is the combination of the state based life expectancies weighted to the proportions of the population in each state

![](_page_8_Picture_4.jpeg)

![](_page_8_Picture_5.jpeg)

# Continuous time

- Similar methodology, but intensities not probabilities modelled
- Covariates are modelled on intensities
- Matrix with transition probabilities for elapsed time t is given by exp(tQ)
- Hidden transitions are modelled
- Where  ${\boldsymbol{\mathsf{Q}}}$  is the transition intensity matrix

	<b>~</b> 1 <i>2</i>	113
$q_{21}$ -	$-\{q_{21}+q_{23}\}$	$q_{_{23}}$
	0	0

Integration used for life expectancies

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![](_page_9_Picture_8.jpeg)

# Microsimulation

- Simulate trajectories from estimated coefficients (any multistate model)
- Creates life histories for individuals

![](_page_10_Picture_3.jpeg)

![](_page_10_Picture_4.jpeg)

# Study design issues

- Longitudinal data
- States measured similarly at each stage
- Fixed or variable time intervals
- Irreversible or reversible
- Main limitation number of transitions
- Sampling method important for variance estimation

![](_page_11_Picture_7.jpeg)

# Considerations

- Time intervals
- Number of transitions
- Study design

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- Sampling scheme
- Size of study
- Model assumptions

![](_page_12_Picture_7.jpeg)