

# Prognosis

Fletcher book

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# Prognosis and prognostic factors

Prognosis = to predict future as closely as possible

- Clinical course
- Natural history (asymptomatic diseases, underestimation of clinical symptoms)

Prognostic factors = tailored information on individual patient

- Patients characteristics

	Prognostic factors	Risk factors
Subjects	Prognostic study Sick people	Cohort study Healthy people
Outcomes	Onset of the disease	Disease consequences
Rates	↓	↑
Factors itself		

# Prognostic studies

## 1. Patients sample



Geographically or clinically defined population at the same stage of the disease (zero time\*)

→ GENERALIZABILITY

Population based study

Population sample based study

→ it is always important to make it clear who the study population is so that others can use the results appropriately

Patients characteristics

Patients setting

Way of sampling

# Prognostic studies

## 1. Patients sample



Zero time



All patients enter the study at the same point in the time of the disease course

→ GENERALIZABILITY

→ it is always important to make it clear who the study population is so that others can use the results appropriately

# Prognostic studies

## 1. Patients sample



Zero time



Onset of the symptoms, diagnosis (inception cohort), beginning of the treatment

→ GENERALIZABILITY

→ it is always important to make it clear who the study population is so that others can use the results appropriately

# Prognostic studies

## 2. Follow-up time

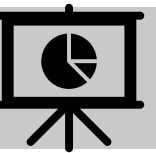


Too short

Too long



## 3. Outcomes



### The 5 Ds

Death

Disease

Discomfort

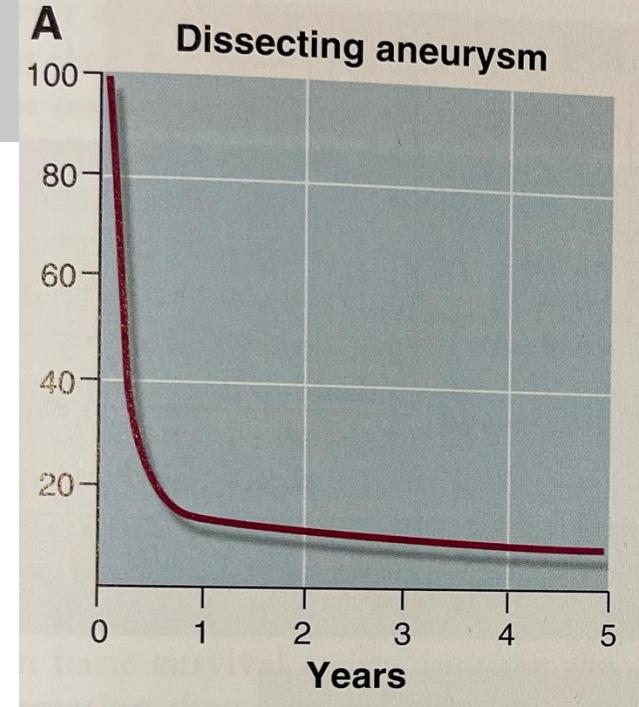
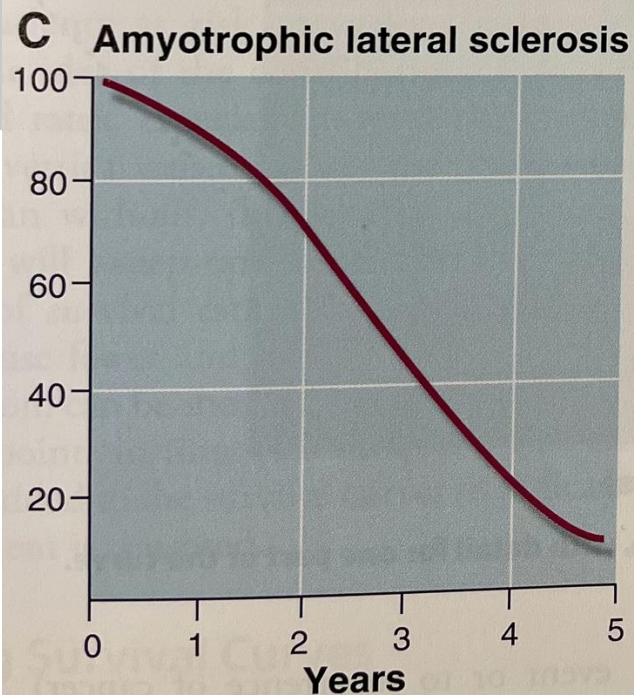
Disability

Dissatisfaction

# Description of prognosis

RATE	Definition	RATE	Definition
<b>5-year survival</b>	Percent of patients surviving 5 years from some point in the course of disease	<b>Response</b>	Percent of patients showing some evidence of improvement following an intervention
<b>Case fatality</b>	Percent of patients with a disease who die of it	<b>Remission</b>	Percent of patients entering a phase in which disease is no longer detectable
<b>Disease-specific mortality</b>	Number of people per 10 000 population dying of a disease	<b>Recurrence</b>	Percent of patients who have return of disease after a disease-free interval

10%  $\neq$  10%

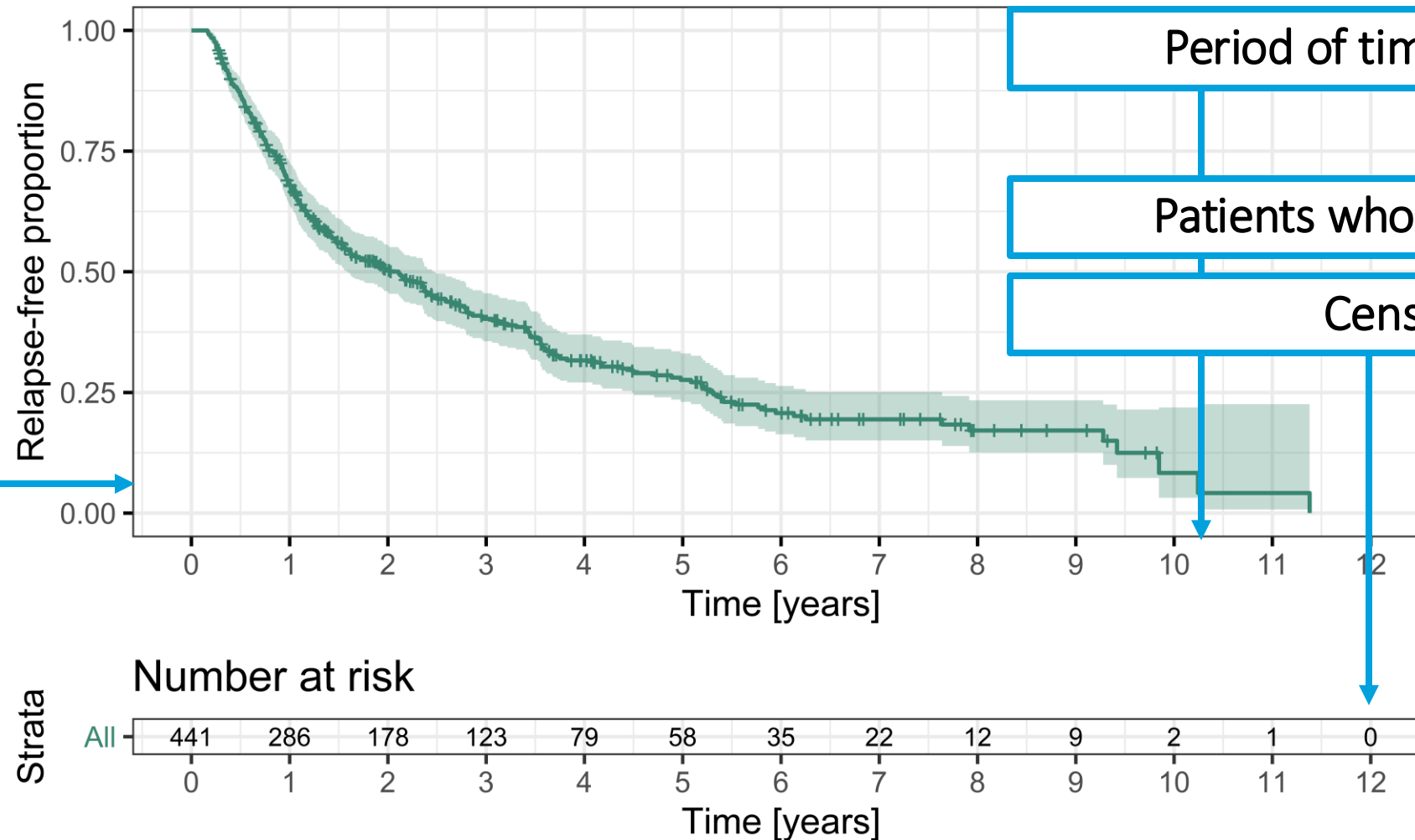


# Description of prognosis

## Survival analysis

Dichotomous outcome which can happen only once

Estimated probability of surviving



Period of time from the beginning

Patients who reached the outcome

Censored patients

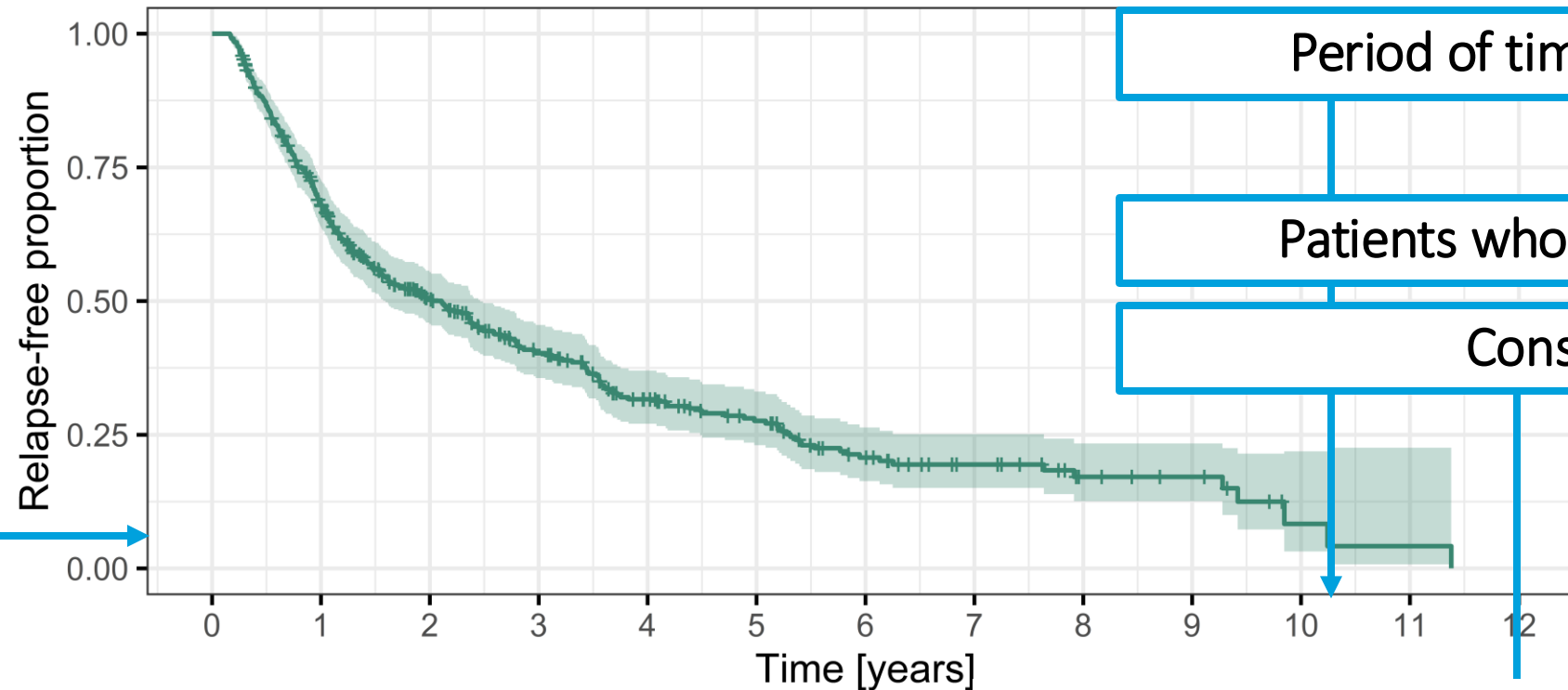


# Description of prognosis

## Survival analysis

Dichotomous outcome which can happen only once

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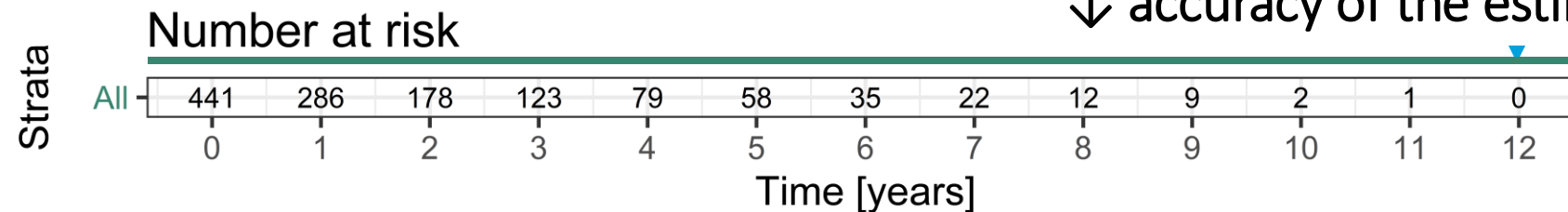


Period of time from the beginning

Patients who reached the outcome

Censored patients

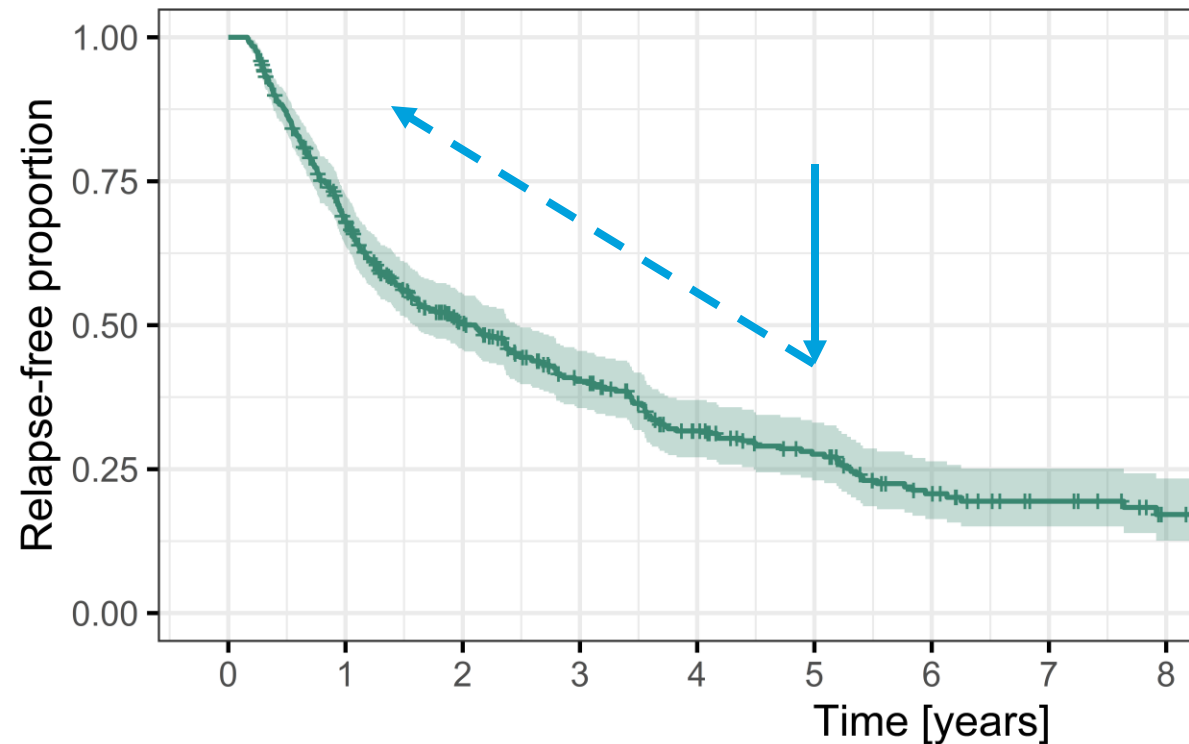
↓ accuracy of the estimated probability of survival



# Description of prognosis

## Survival analysis

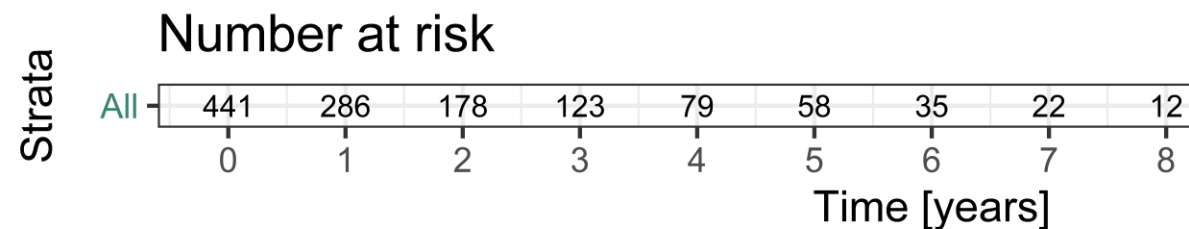
Dichotomous outcome which can happen only once



Estimated propability of surviving

The propability of surviving to any point is estimated form cumulative probility of surviving in each of the time intervals that precede it.

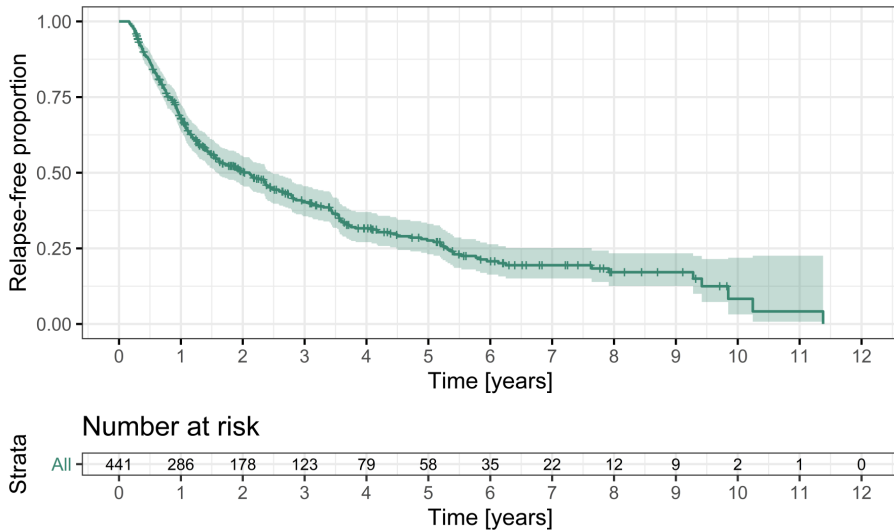
Ratio of the number of patients surviving to the number of patients at risk at the same moment



5 at risk, 1 died  $\rightarrow$  4 : 5

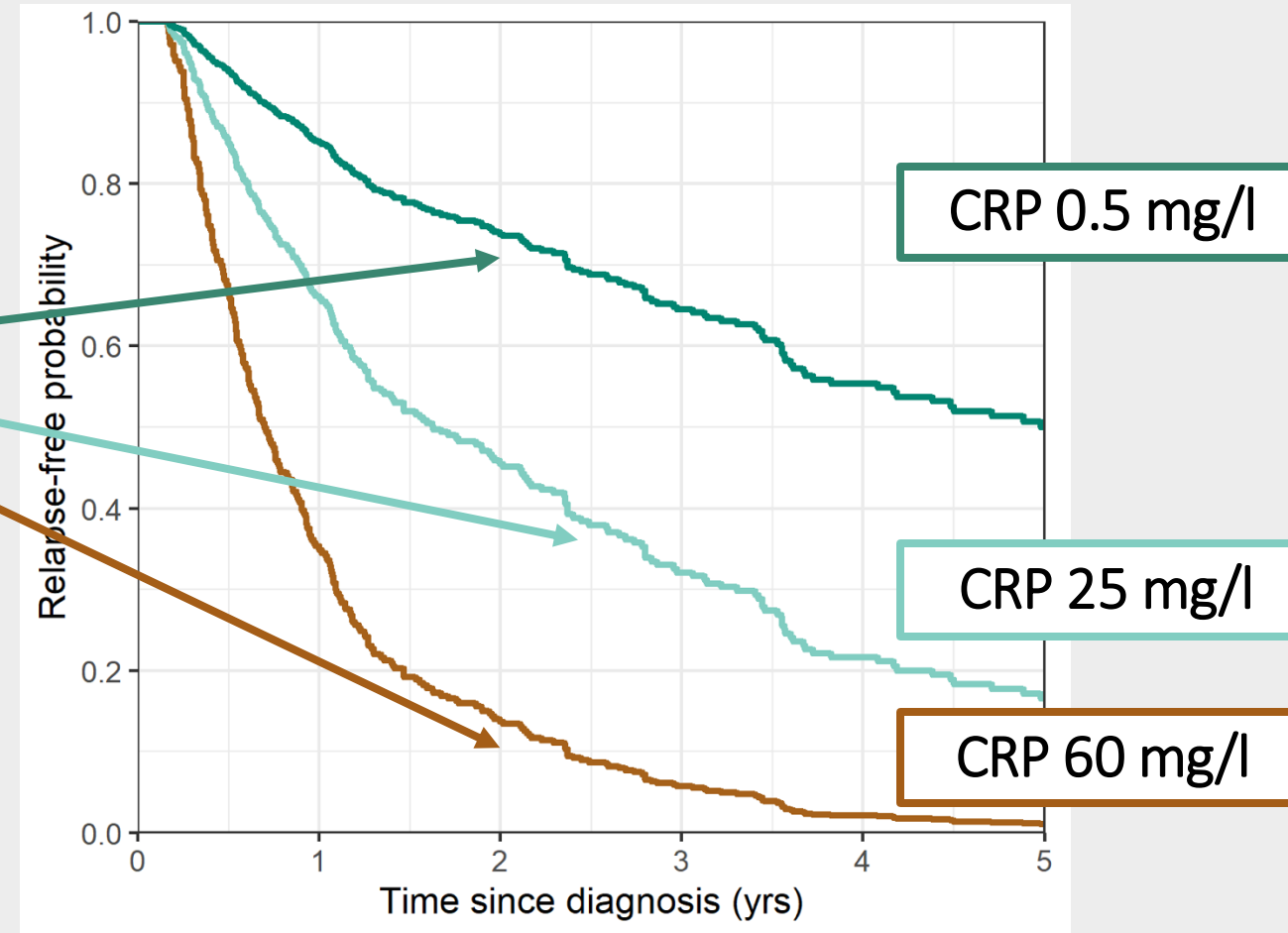
# Prognostic factors

## Prognosis of a group



Newly diagnosed paediatric CD patients starting treatment with azathioprin at the time of diagnosis

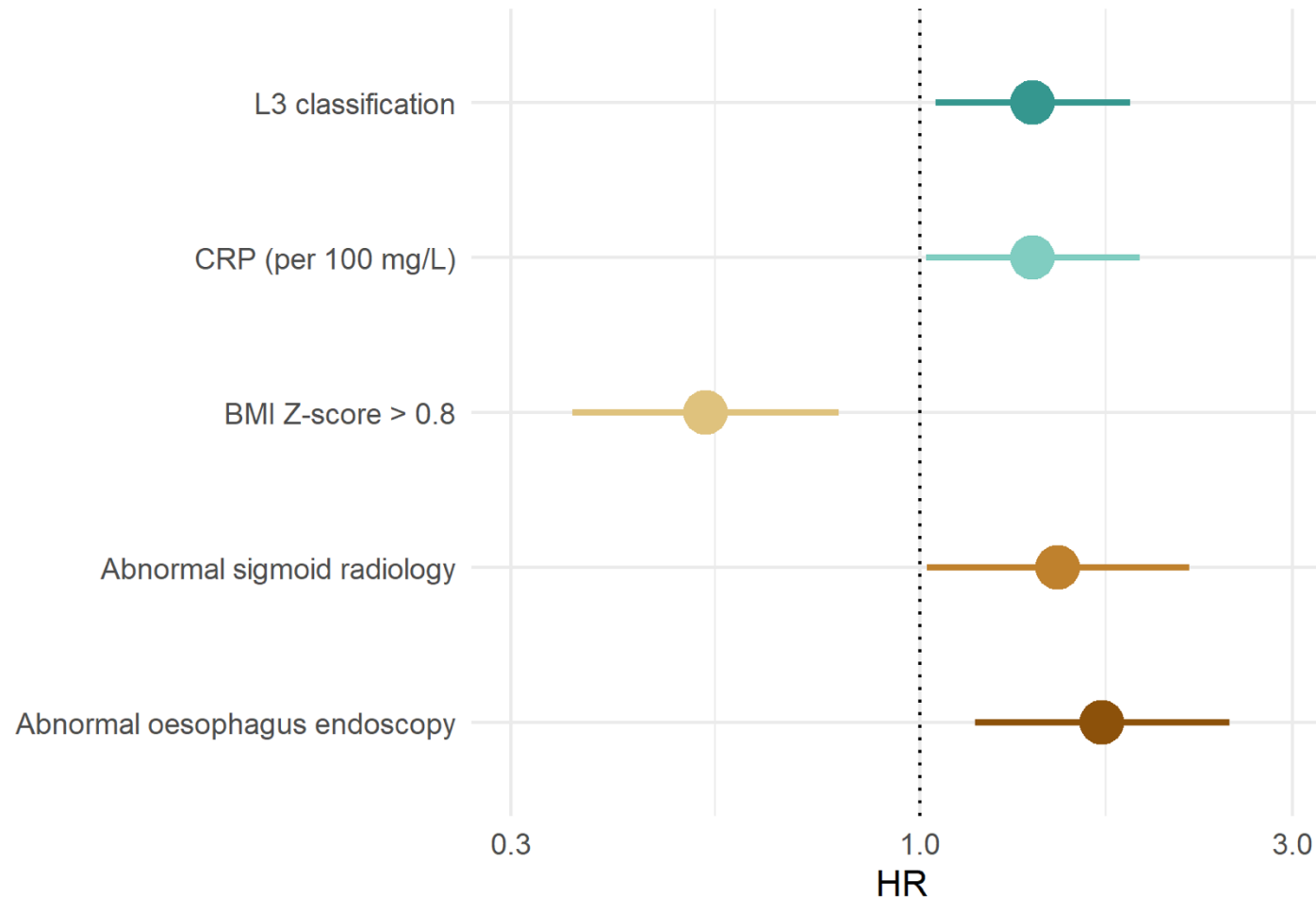
## Prognosis in patients with different characteristics



CRP level at diagnosis = prognostic factor

# Prognostic factors

Effects of one prognostic factor relative to the effect of another can be summarized from data in a time to event analysis by Hazard ratio (HR)



# Study designs

## Case series

### Clinical prediction rule

- combination of different variables
- fix point in a time
- training and testing set

## Cohort studies

- sampling bias
- migration bias
- measurement bias (equality, unawareness, clear rules)
- bias from non-differential misclassification
- bias from missing data (imputation)
  - Recognize the bias and consider its implication on clinical question

# Study designs

## Case series

### Clinical prediction rule

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## Cohort studies

- sampling bias
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- bias from non-differential misclassification
- bias from missing data (imputation)
  - Sensitivity analysis

# Conclusion

- Prognosis
- Prognostic factors
- Clinical course
- Nature history
- Zero time
- Inception cohort
- Event
- Survival analysis
- Kaplan-Mayer analysis
- Censored
- Hazard ratios
- Case series
- Case report
- Clinical prediction rule
- Training set
- Test set
- Validation
- Prognostic stratification
- Sampling bias
- Migration bias
- Dropouts
- Measurement bias
- Multiple imputation
- Sensitivity analysis
- Best-case / worst-case analysis