

# Introduction to Causal Inference

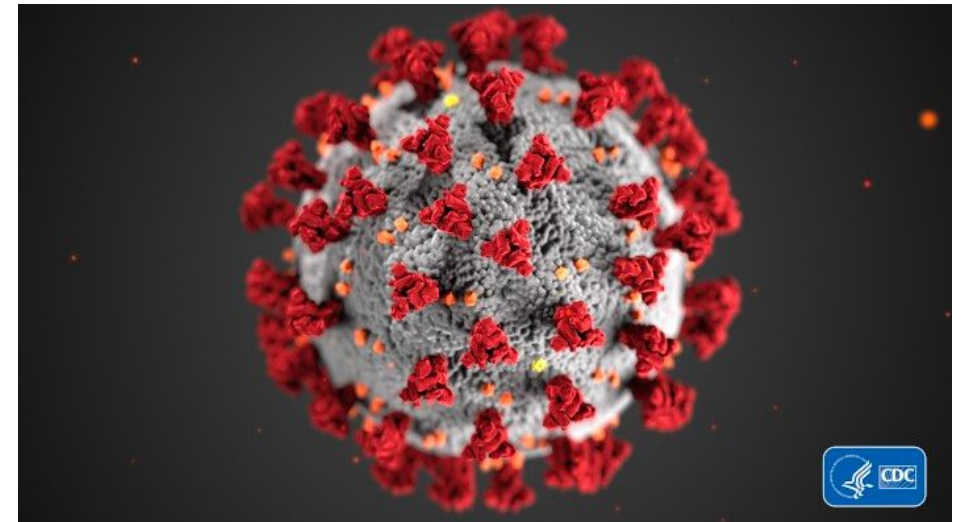
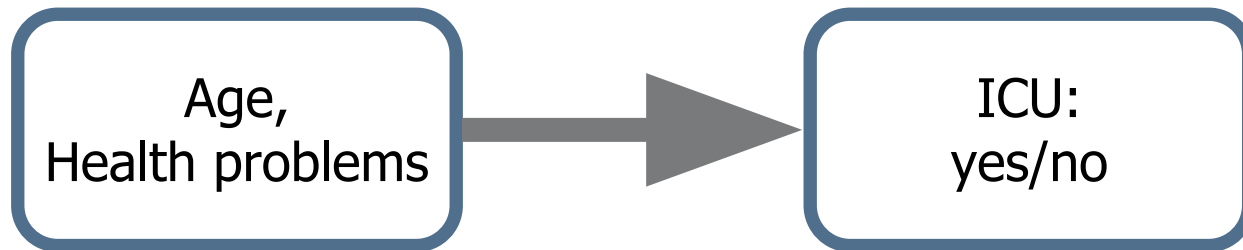
Sesiones de Estadística DAP-Cat

Aleix Ruiz de Villa

Main Objective of these 2 sessions

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## ICU ingress model due to covid



<https://www.cancer.org/es/noticias-recientes/preguntas-comunes-acerca-del-brote-del-nuevo-coronavirus.html>

## STATISTICS / CAUSAL INFERENCE

**Objective:** ¿do age and healthcare affect ICU ingress?

**Decision:** design public health strategies

**Model:** finding the correct model

## MACHINE LEARNING

**Objective:** Predict the risk

**Decision:** patient ranking

**Model:** finding the most accurate

## Machine Learning main assumption

**Past and Future behave the same**



<https://www.cancer.org/es/noticias-recientes/preguntas-comunes-acerca-del-brote-del-nuevo-coronavirus.html>

## Machine Learning limitation

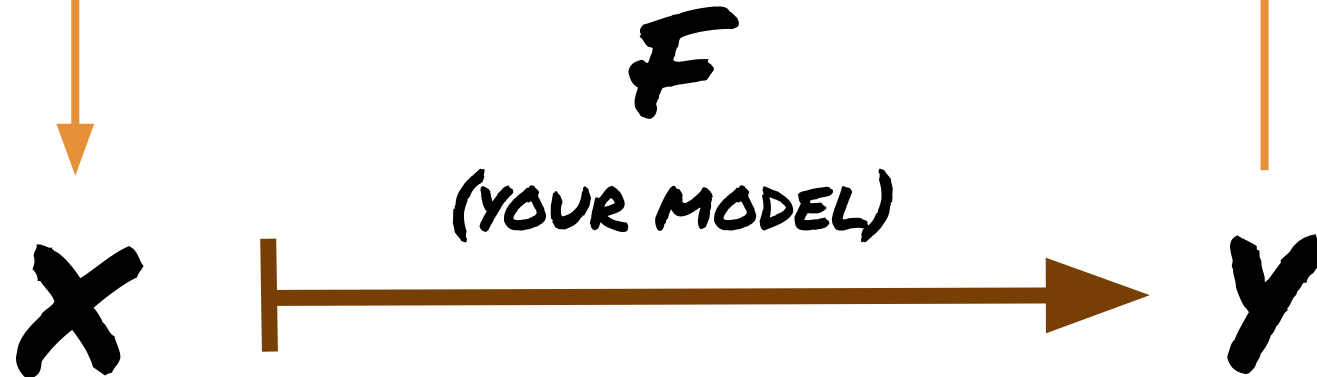
**What if we behave in a different manner?**



<https://www.alpinerecoverylodge.com/intervention-assistance/>

USE **CAUSALITY** WHEN YOUR  
ACTIONS MAKE AN IMPACT HERE

USE **MACHINE LEARNING**  
WHEN YOUR ACTIONS DEPEND ON THIS



CAUSALITY = RCTS + CAUSAL INFERENCE

# Introduction to Causal Inference







# Observational Examples

NEWS

• LIVE TV

INDIA  
TODAY

APP

HOME

 MY FEED

INDIA

WORLD

BUSINESS

TECH

MOVIES

SPORTS

SCIENCE

HEALTH

VIDEOS

News / Lifestyle / Health / Moderate drinking during pregnancy doesn't harm baby

## Moderate drinking during pregnancy doesn't harm baby



# Observational Examples

The logo for MailOnline, featuring the word "Mail" in a large, bold, black serif font, followed by "Online" in a smaller, blue, sans-serif font.

[Home](#) [News](#) [U.S.](#) | [Sport](#) | [TV&Showbiz](#) | [Australia](#) | [Femail](#) | [Health](#) | [Science](#) | [Mone](#)  
[Latest Headlines](#) | [Covid-19](#) | [Royal Family](#) | [Prince Harry](#) | [Meghan Markle](#) | [World News](#) | [Headlines](#) | [Mos](#)

**Processed meat 'is to blame for one in 30 deaths': Scientists say a rasher of cheap bacon a day is harmful**

## Frameworks

- Potential Outcomes: Biostatistics & Econometrics
- Directed Acyclic Graphs: Computer Science

Both are equivalent, but each one suitable for different things

# Simpson's paradox

—

# Simpson's Paradox

Treatment	Size	Number	Recovered
A	Small	87	81
B	Small	270	234
A	Large	263	192
B	Large	80	55



## Hospital's Main Problem

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*Which of the two treatments should they take?*

# Simpson's Paradox

Size	Treatment A	Treatment B
Recovery	78% (273/350)	<b>83%</b> (289/350)



# Simpson's Paradox

	Treatment A	Treatment B
Large	<b>93%</b> (81/87)	87% (234/270)
Small	<b>73%</b> (192/263)	69% (55/80)

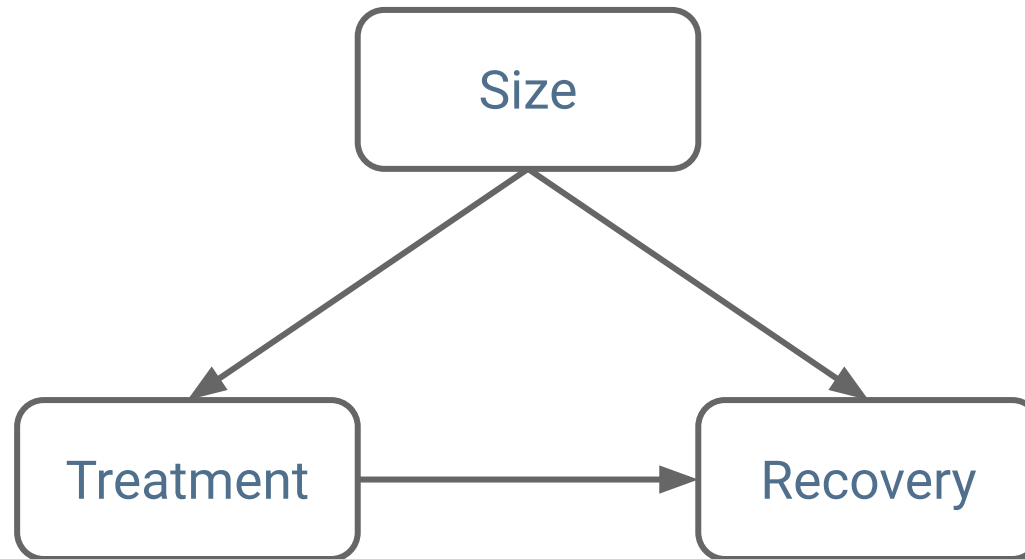




# Analysis of Simpson's paradox

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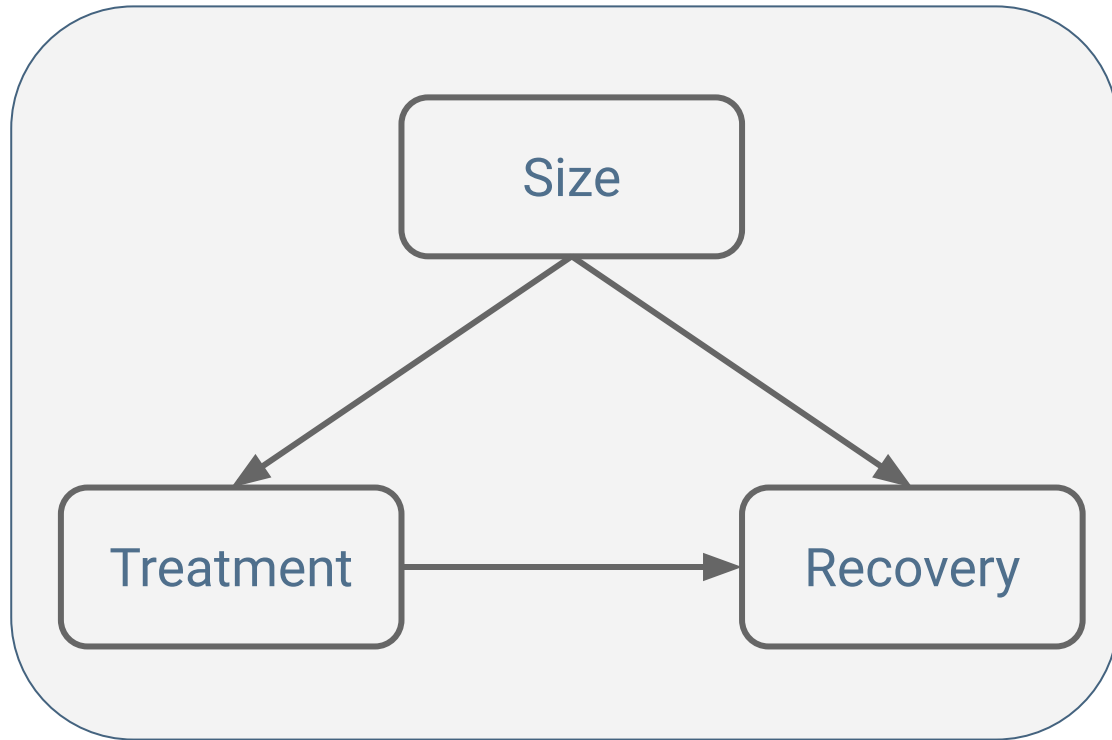
# Graph



# Size Distribution

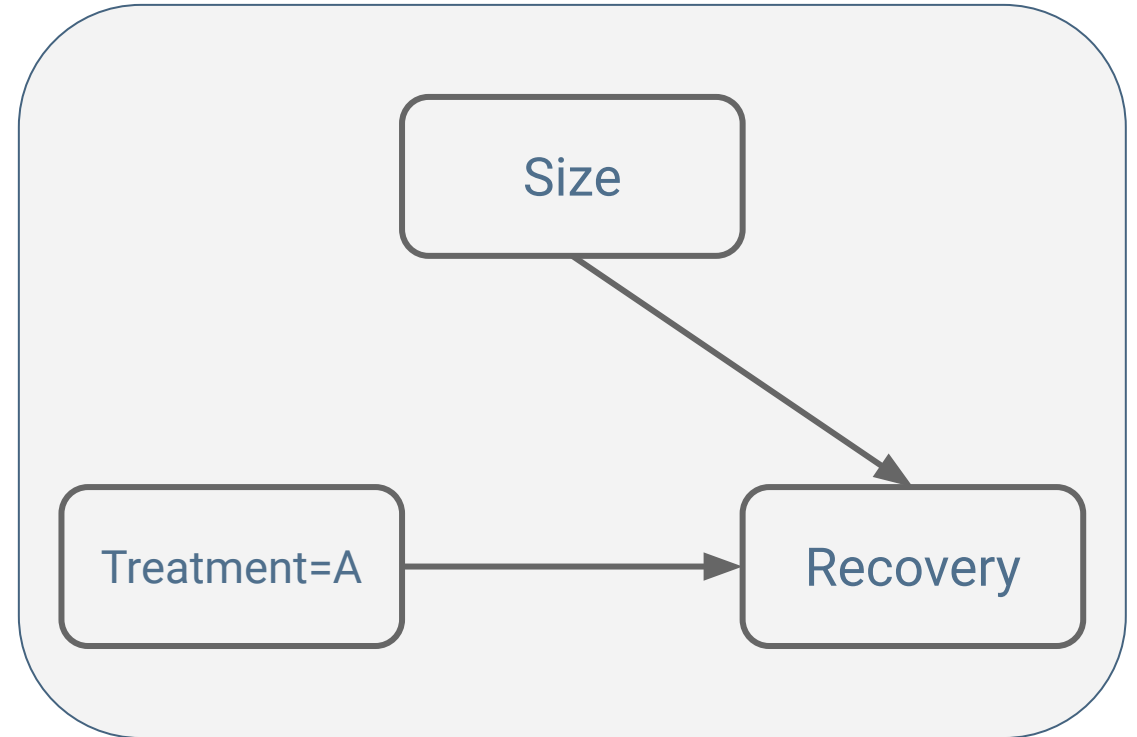


# What is an intervention?



The data you have

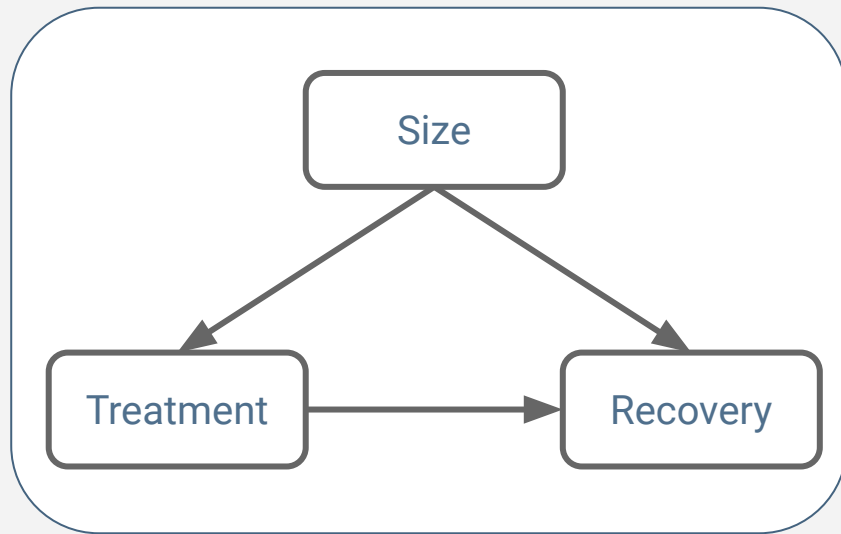
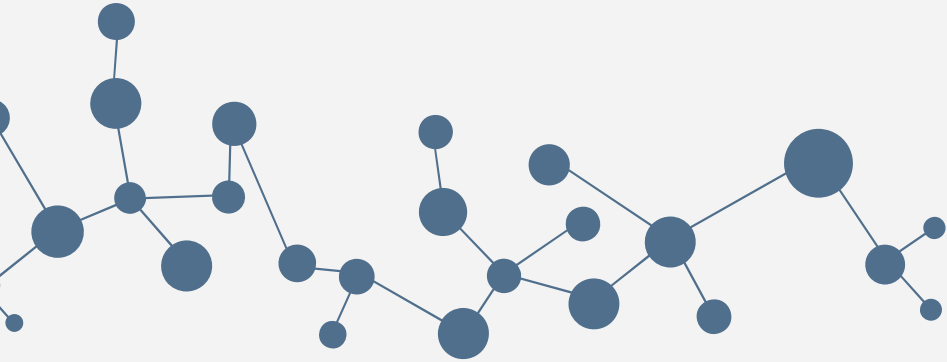
***Distribution  $P$***



The data you would like to have

***Distribution  $P^A = P^{do(T:=A)}$***

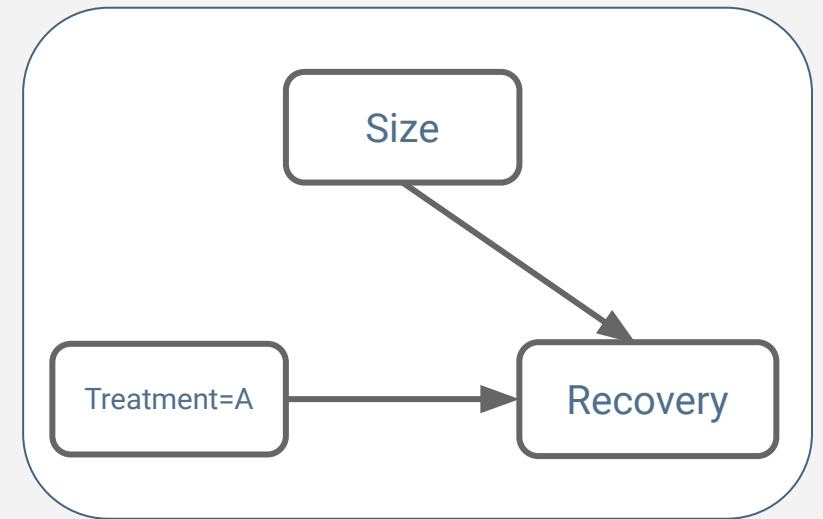
# Main objective of causal inference



*Use observational data*



*To infer about interventional data*



# Average Treatment Effect

## Conditional Probability

$$P(R=1 | T=A) = P(R=1 | T=A, S=Small) * P(S=Small|T=A) + P(R=1 | T=A, S=Large) * P(S=Large|T=A)$$

$$= 93\% * 25\% + 73\% * 75\% = 78\%$$

## Adjustment

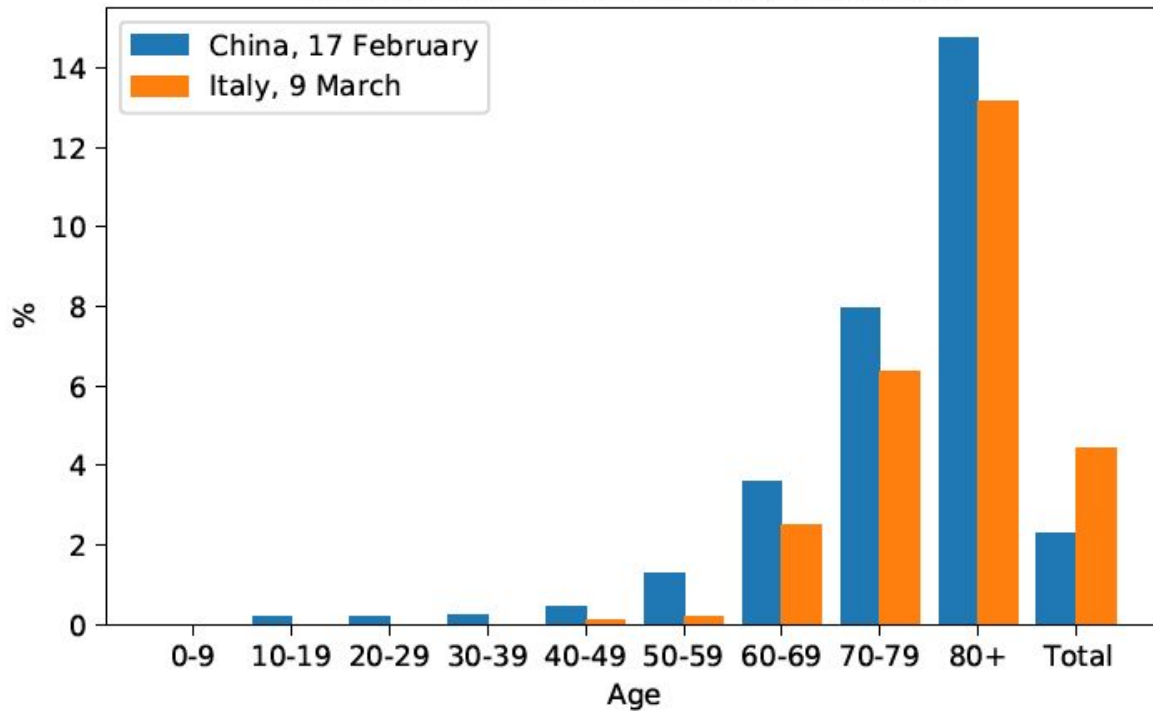
$$P(R=1 | do(T=A)) = P(R=1 | T=A, S=Small) * P(S=Small) + P(R=1 | T=A, S=Large) * P(S=Large)$$

$$= 93\% * 51\% + 73\% * 49\% = 83\%$$

# Covid Example

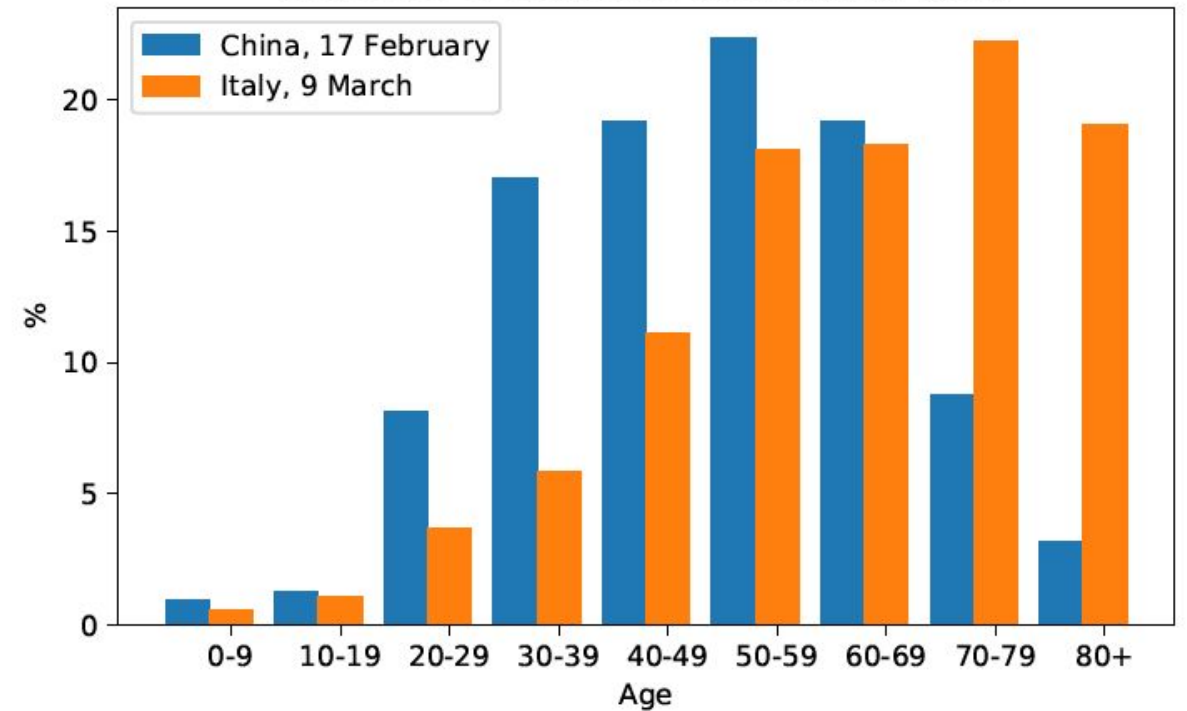


Case fatality rates (CFRs) by age group



(a)

Proportion of confirmed cases by age group



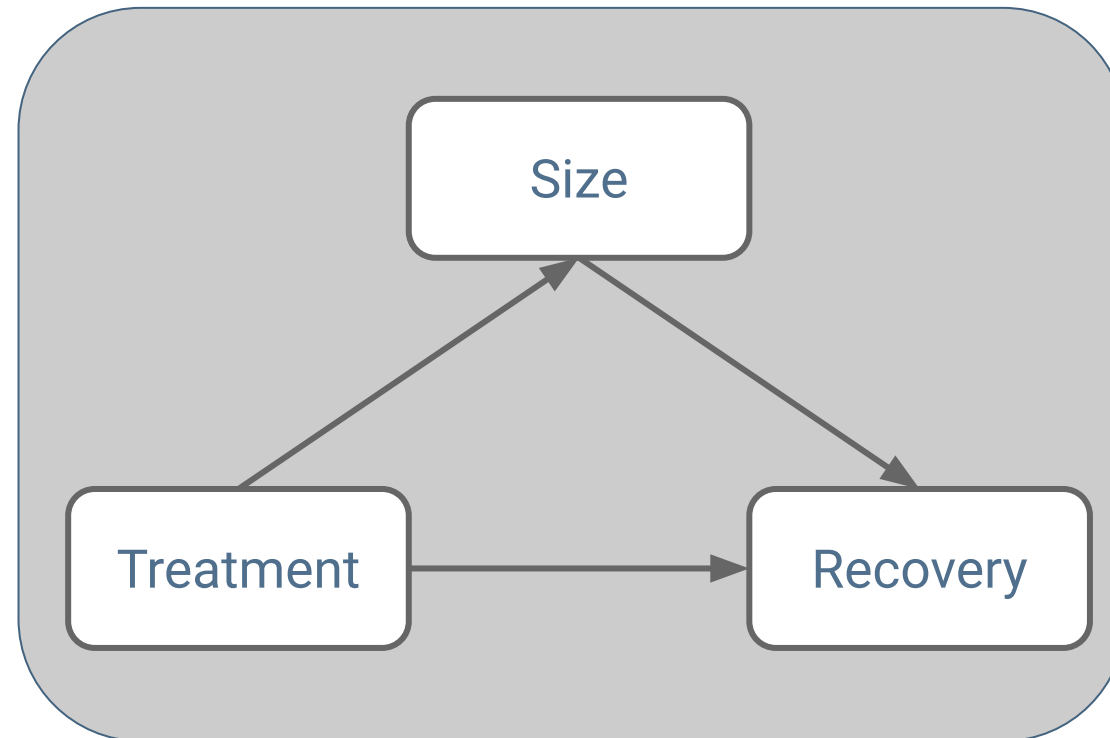
(b)



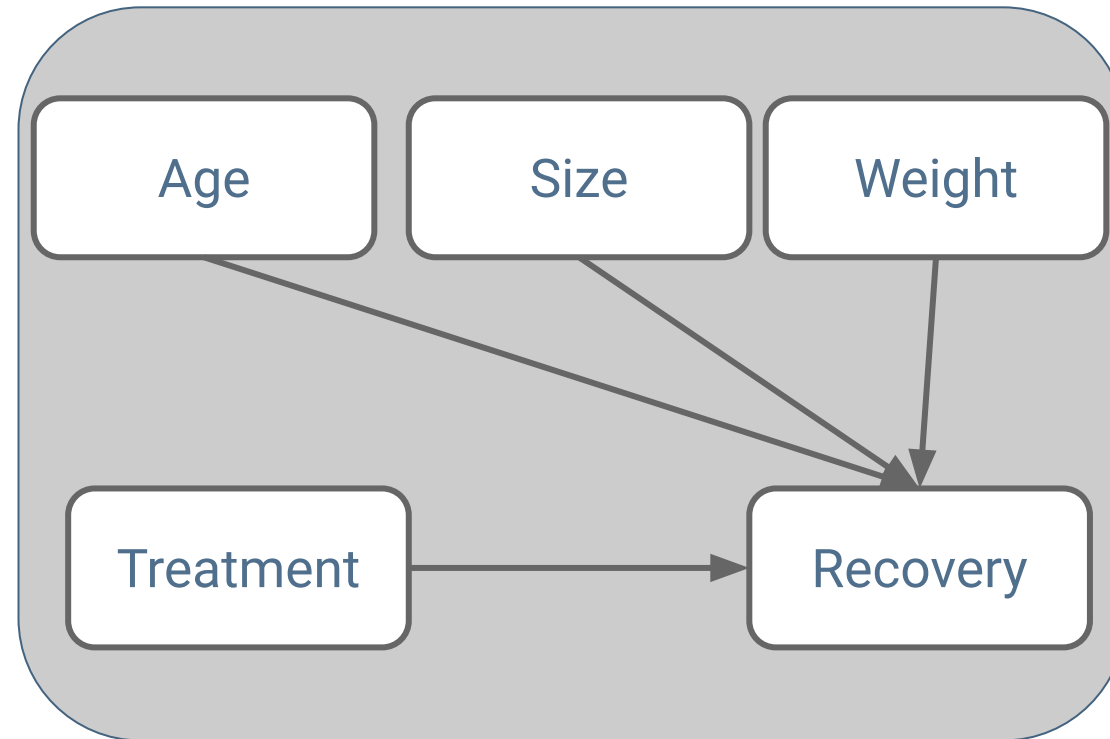
When to adjust?



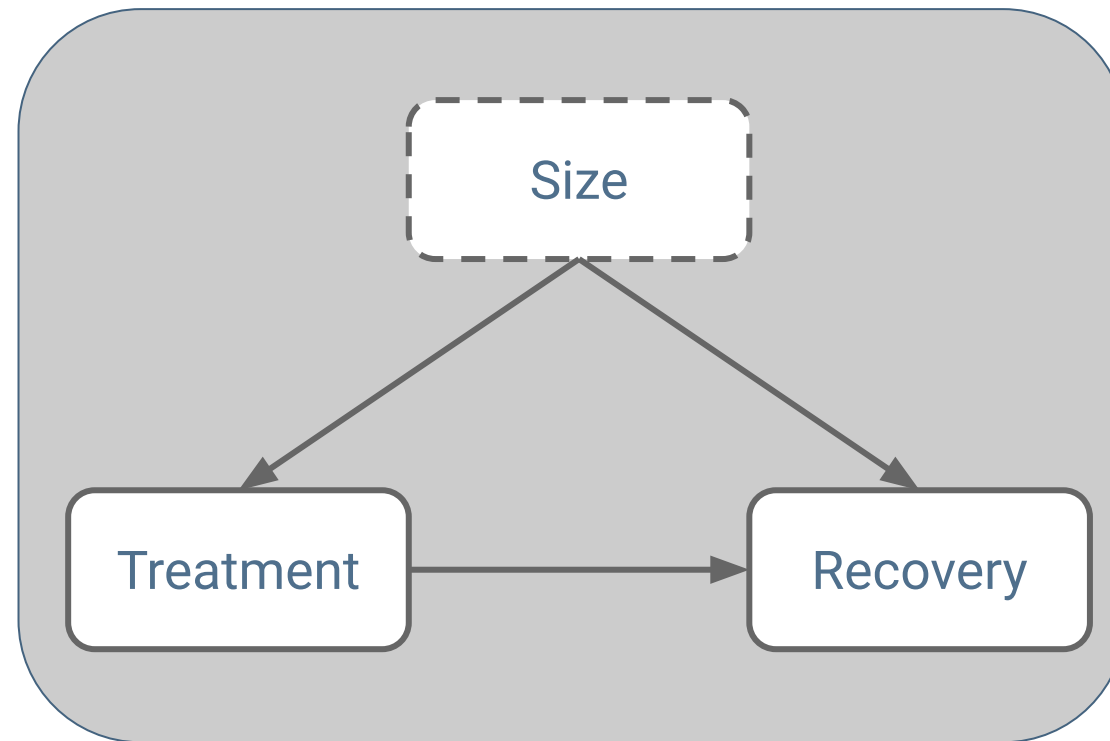
Do we need to adjust?



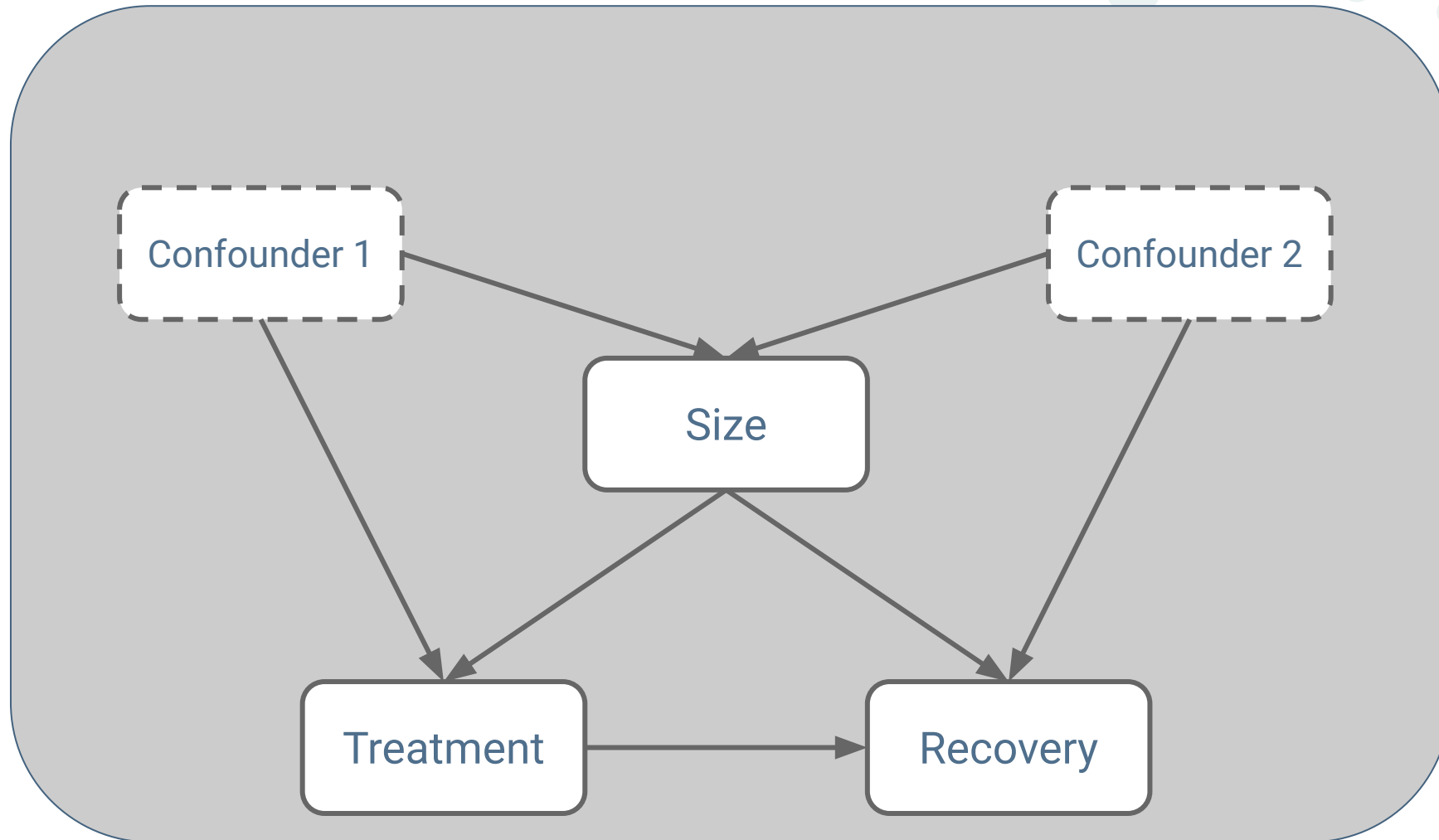
Do we need to adjust?



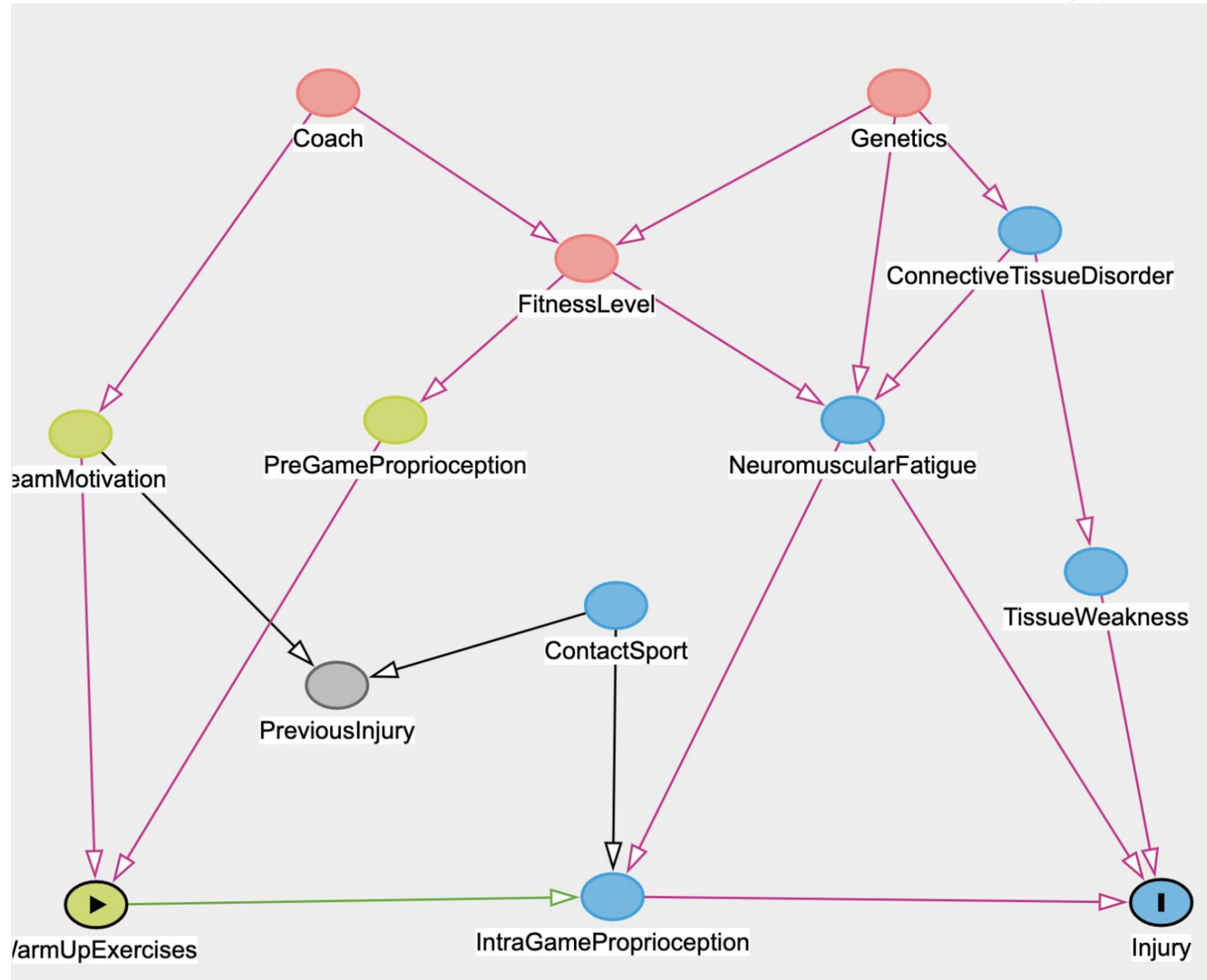
Do we need to adjust?



Do we need to adjust?



# Example



Adjustment (total effect) ▾

Minimal sufficient adjustment sets for estimating the total effect of WarmUpExercises on Injury:

- Coach, FitnessLevel
- Coach, PreGameProprioception
- ConnectiveTissueDisorder, NeuromuscularFatigue
- FitnessLevel, Genetics
- FitnessLevel, TeamMotivation
- NeuromuscularFatigue, TissueWeakness
- PreGameProprioception, TeamMotivation

▾ Testable implications

The model implies the following conditional independences:

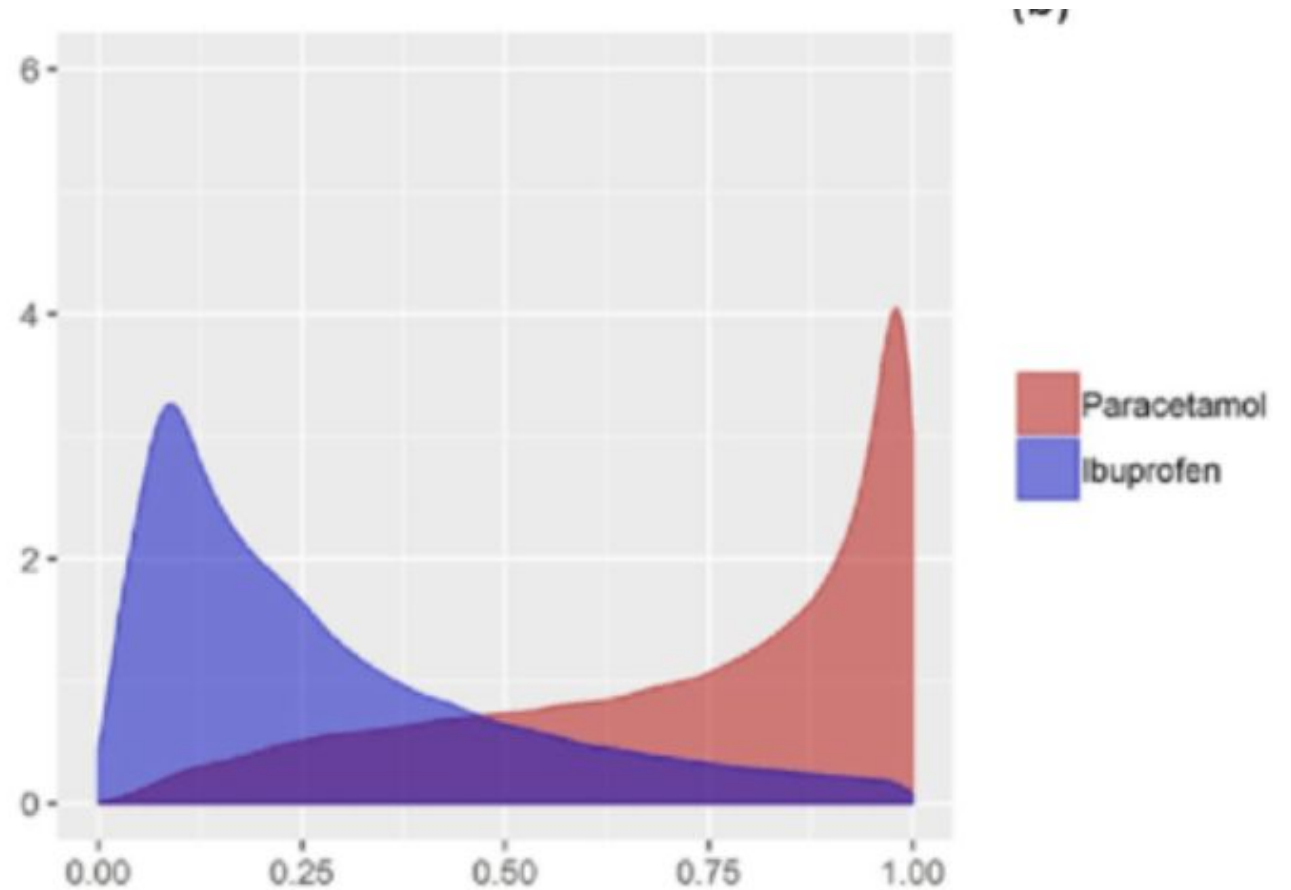
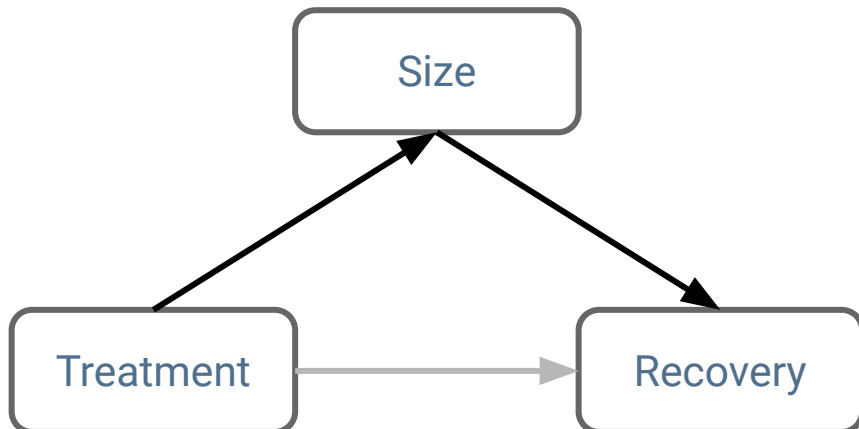
- WarmUpExercises  $\perp$  Injury | IntraGameProprioception, NeuromuscularFatigue, TissueWeakness
- WarmUpExercises  $\perp$  Injury | ConnectiveTissueDisorder, IntraGameProprioception, NeuromuscularFatigue
- WarmUpExercises  $\perp$  Injury | FitnessLevel, Genetics, IntraGameProprioception, NeuromuscularFatigue



# Applications

## Variations of the adjustment formula

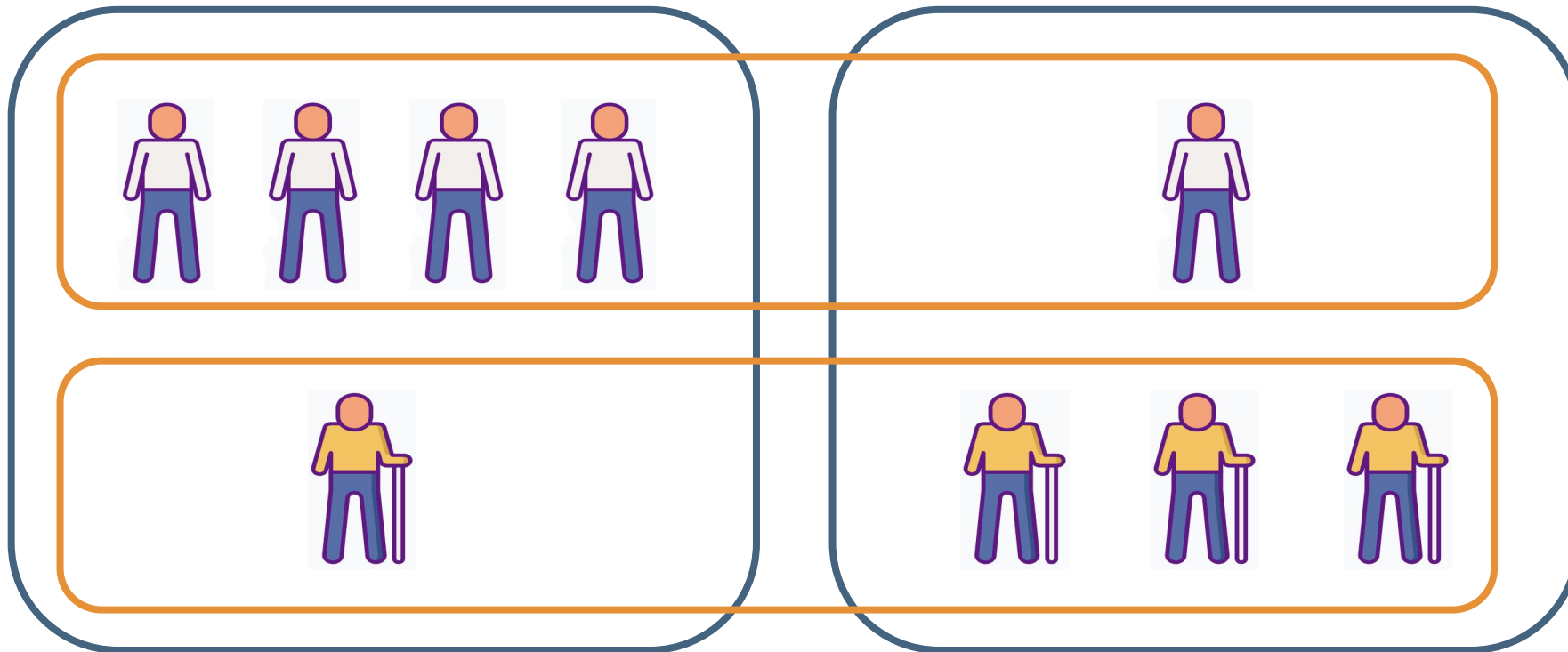
1. Propensity Scores
2. In linear models: controlling for some variables
3. Mediation Analysis



## Propensity Score

TREATMENT

CONTROL



**We can compare  
if they have  
same attributes**

**Propensity  
Score:**

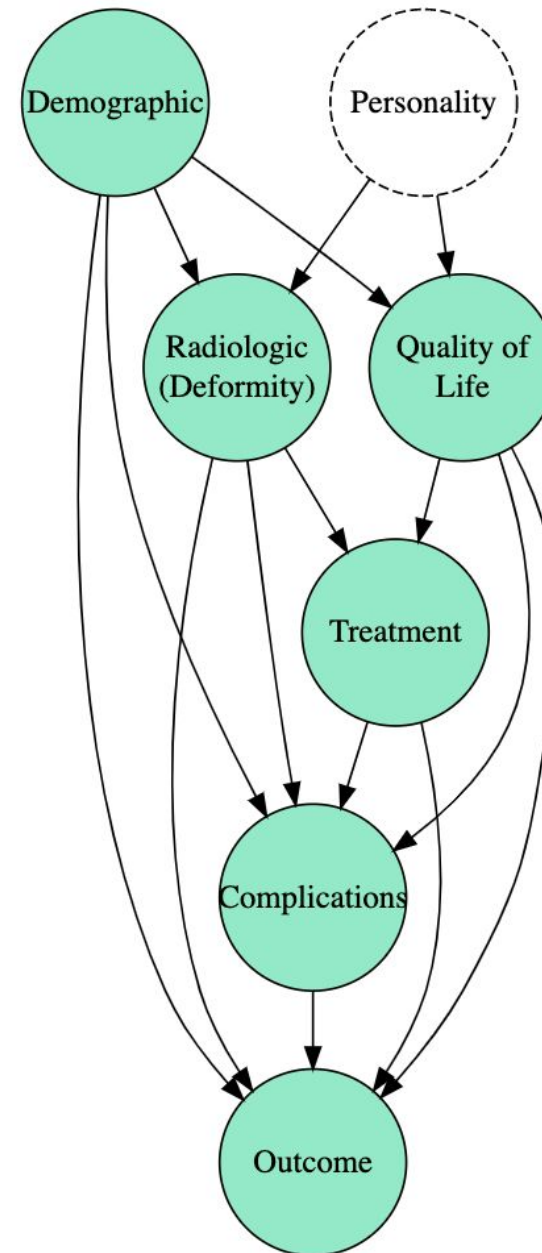
**We can compare  
if they have the  
same chances to  
be treated**



# Example

Differences between

- What is the impact of a particular type of treatment
- What is the difference between treating the patient or not





# Why RCTs are so important



## Why RCTs are so important

- Which are the confounders of a RCT?
- In general, how are we sure that are considering all possible confounders?

### **RCT**

- Measuring an outcome.
- Mean risk: uncertainty

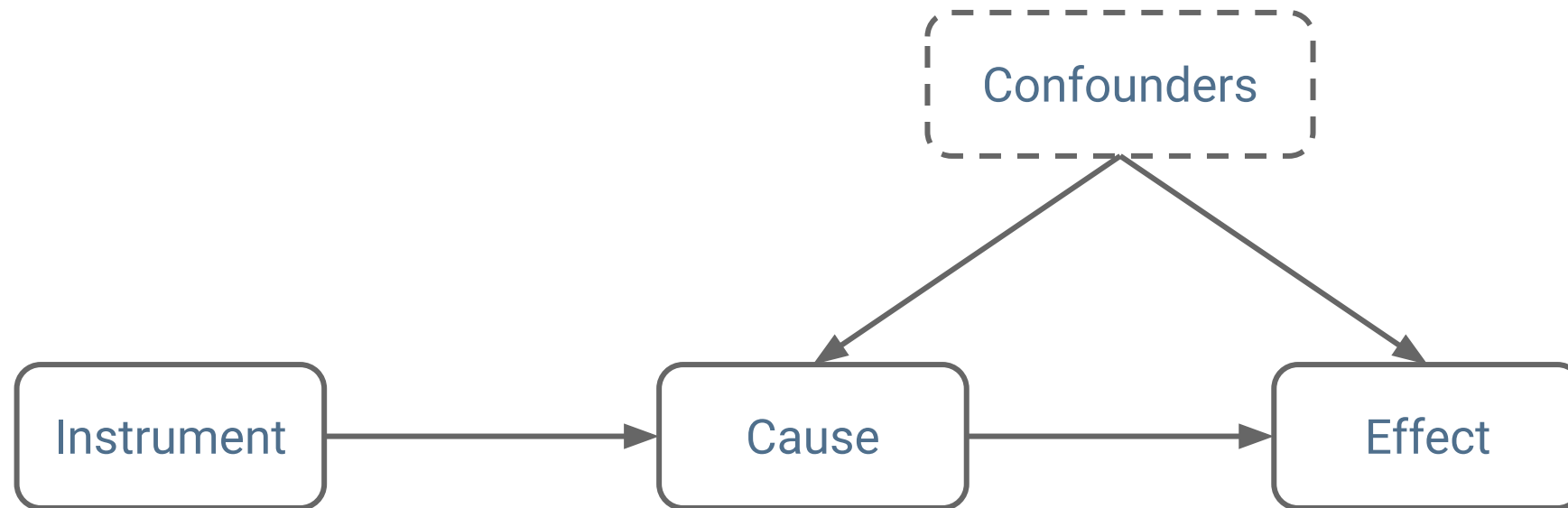
### **Causal Modeling**

- Modeling Causes + Measuring outcome
- Main risk: errors in modeling + uncertainty

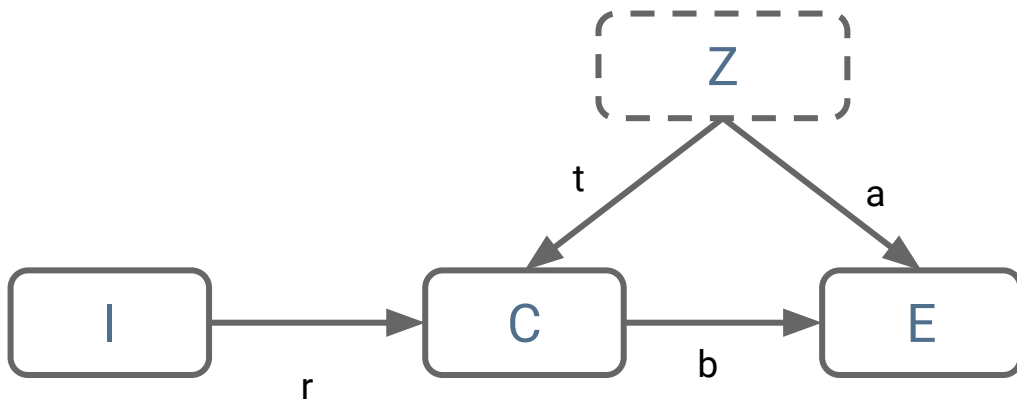


# Instrumental Variables

# Graph



# Estimation



## Model

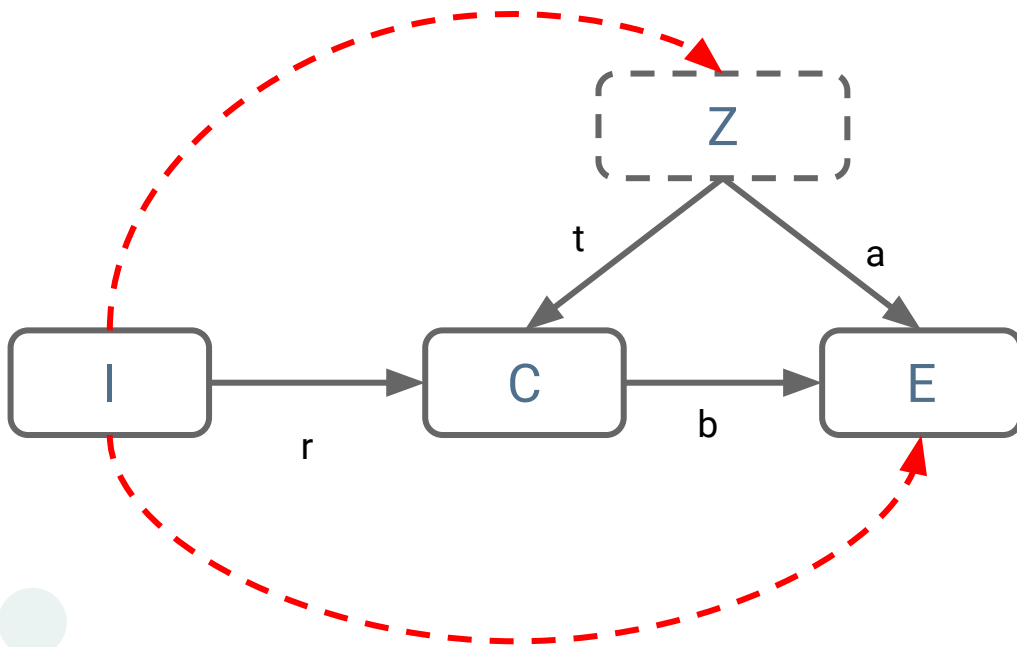
$$E = a Z + b C$$

$$C = r I + t Z$$

## Formulation

$$\begin{aligned} E &= a Z + b (r I + t Z) = \\ &= (a + bt) Z + br I \end{aligned}$$

# Assumptions



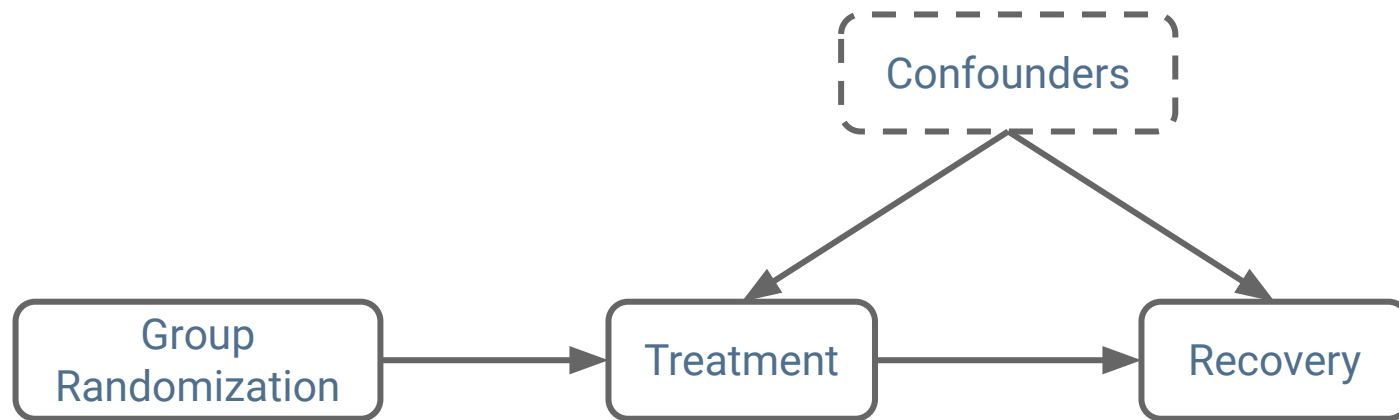
Exclusion restriction:

There is no other path from I to E

Non compliant RCTs



# Graph



Per Protocol

Intention to Treat

## Intention to Treat

- Unbiased
- Diluted

## Per Protocol

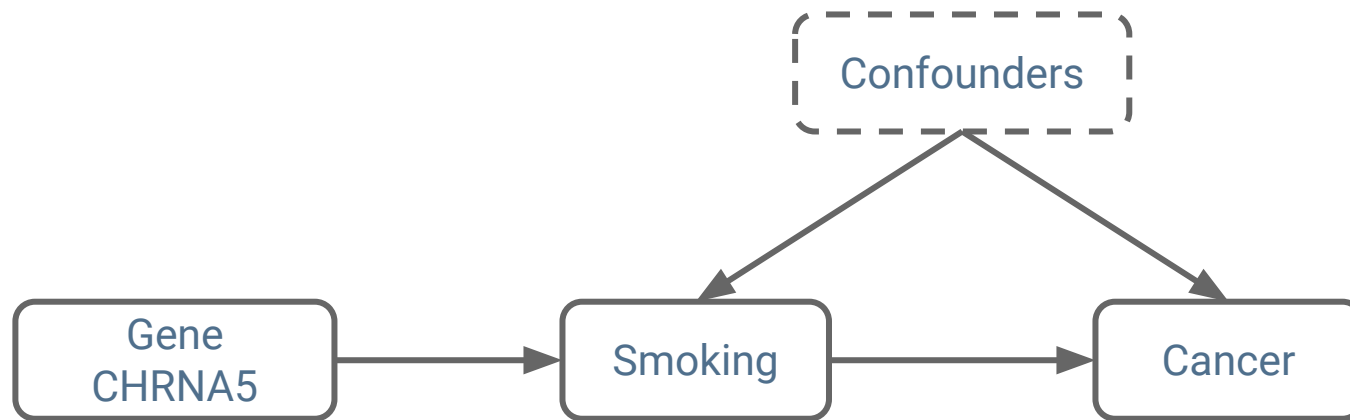
- Potentially biased

## Instrumental Variables:

- Unbiased
- Not diluted

# Mendelian Randomization

# Mendelian Randomization




- CHRNA 4 highly correlated with Smoking (well estimated)
- Exclusion restriction: Out of the smoking group there seems not be any correlation between CHRNA5 and Cancer



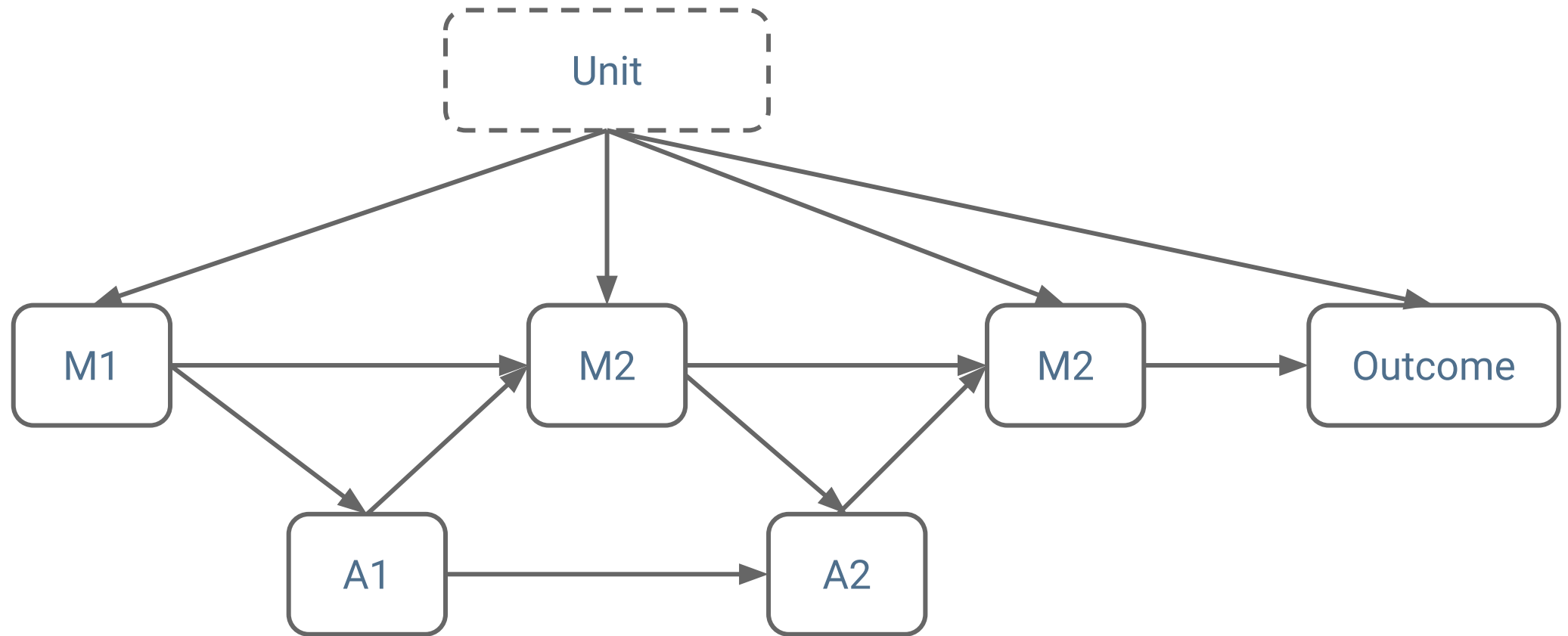
# Advanced Stuff

# Statistical Estimation of Average Treatment Effects

A decorative network graph in the top right corner, consisting of several light blue circular nodes of varying sizes connected by thin grey lines, forming a sparse, interconnected web.

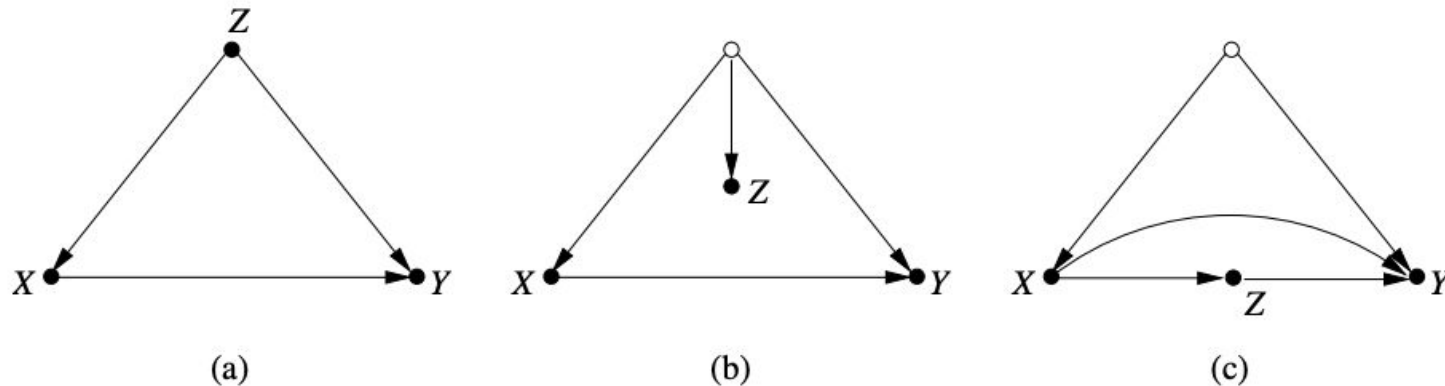
- Using Machine Learning to calculate Propensity Scores and Adjustment Formula
  - Using Double Machine Learning to Estimate Adjustment Formula and Time-Varying treatment effects
  - Confidence intervals and p-values
- 
- A decorative network graph in the bottom left corner, consisting of several light blue circular nodes of varying sizes connected by thin grey lines, forming a sparse, interconnected web.

# Time-Varying Treatment Effects



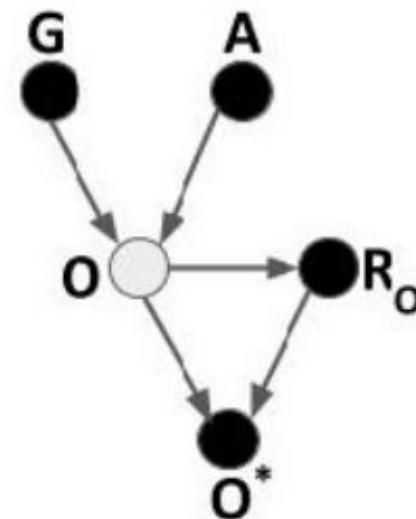
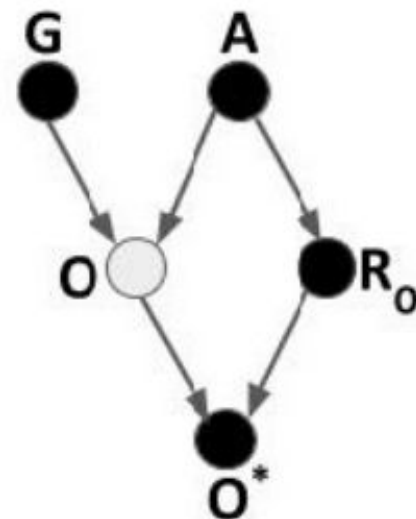
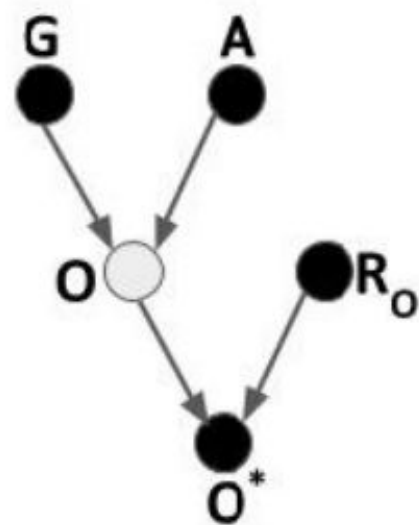
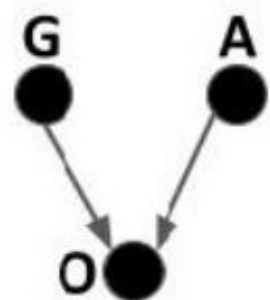
# External Validity and Transportability

**Example 1** We conduct a randomized trial in Los Angeles (LA) and estimate the causal effect of treatment  $X$  on outcome  $Y$  for every age group  $Z = z$  as depicted in Fig. 1(a). We now wish to generalize the results to the population of New York City (NYC), but we find that the distribution  $P(x, y, z)$  in LA is different from the one in NYC (call the latter  $P^*(x, y, z)$ ). In particular, the average age in NYC is significantly higher than that in LA. How are we to estimate the causal effect of  $X$  on  $Y$  in NYC, denoted  $P^*(y|do(x))$ .<sup>1</sup>



**Figure 1:** Causal diagrams depicting Examples 1–3. In (a)  $Z$  represents “age.” In (b)  $Z$  represents “linguistic skills” while age (hollow circle) is unmeasured. In (c)  $Z$  represents a biological marker situated between the treatment ( $X$ ) and a disease ( $Y$ ).

# Recoverability of missing values



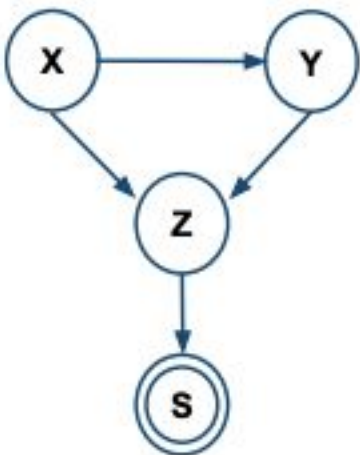


# Recovery from selection Bias

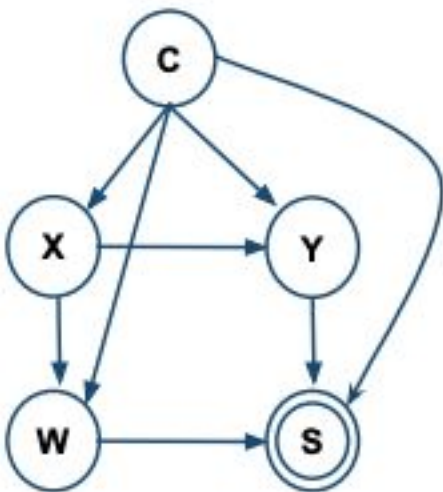
(a)



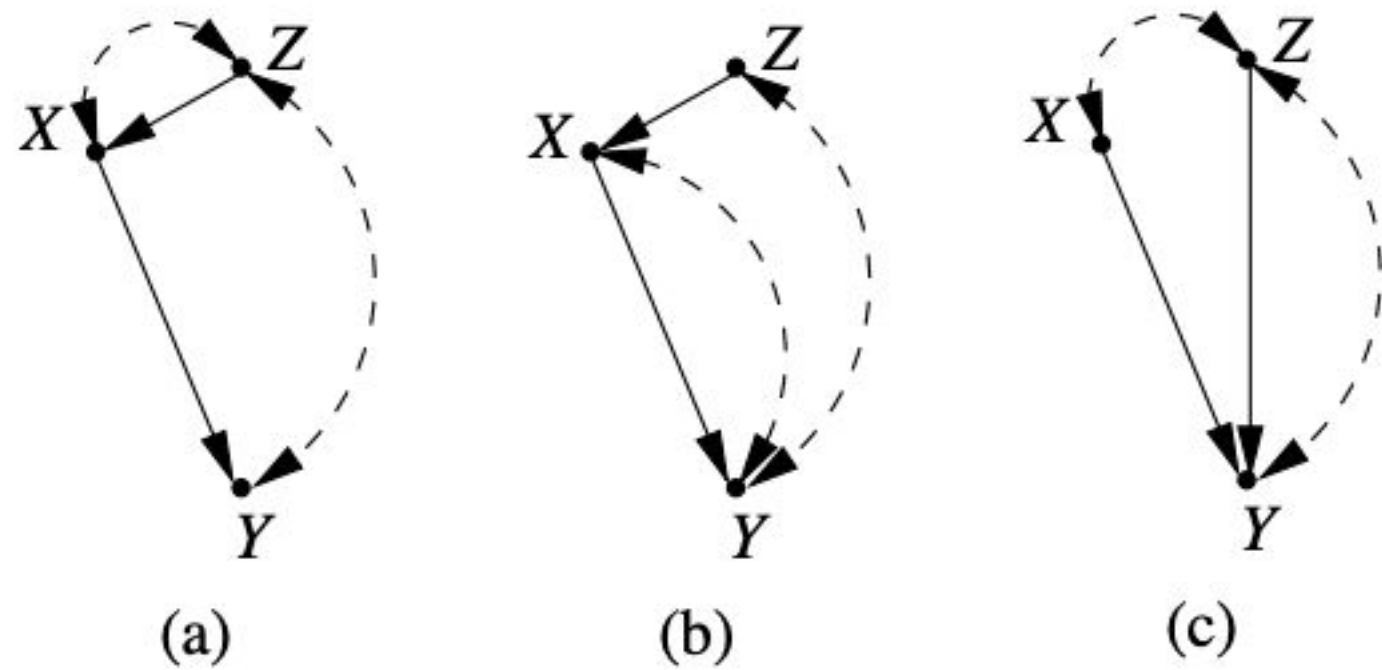
(b)



(c)



Z-identifiability (difficult): identification through auxiliary experiments





# Conclusions



## Conclusions

- CI are more flexible than RCTs, but come at a price: making assumptions with its associated risk
- Causal Inference builds on top of classical statistics, where causality plays a central role
- Causal **modeling** is about (formal) modeling

## Applications

- CI when there is no alternative, prioritizing RCTs, noncompliant RCT, mendelian randomization, ...