DAGS

Tools for reasoning about causes

Einar Holsbø, Department of Computer Science, UiT — October, 2023

The Marko fallacy



The fundamental problem of observational research:

seeing X ≠ doing X

Baby's first DAG: Marko's causal model

Less clothing

More thirsty



Less clothing

More thirsty

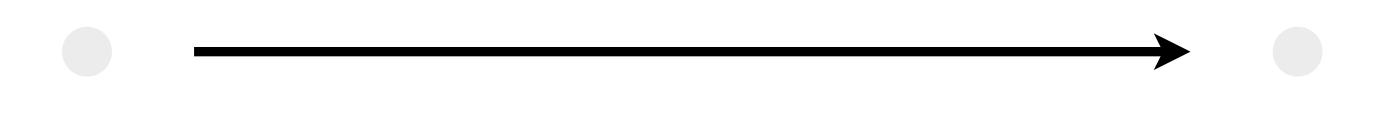
Technical term for some dots joined up with lines



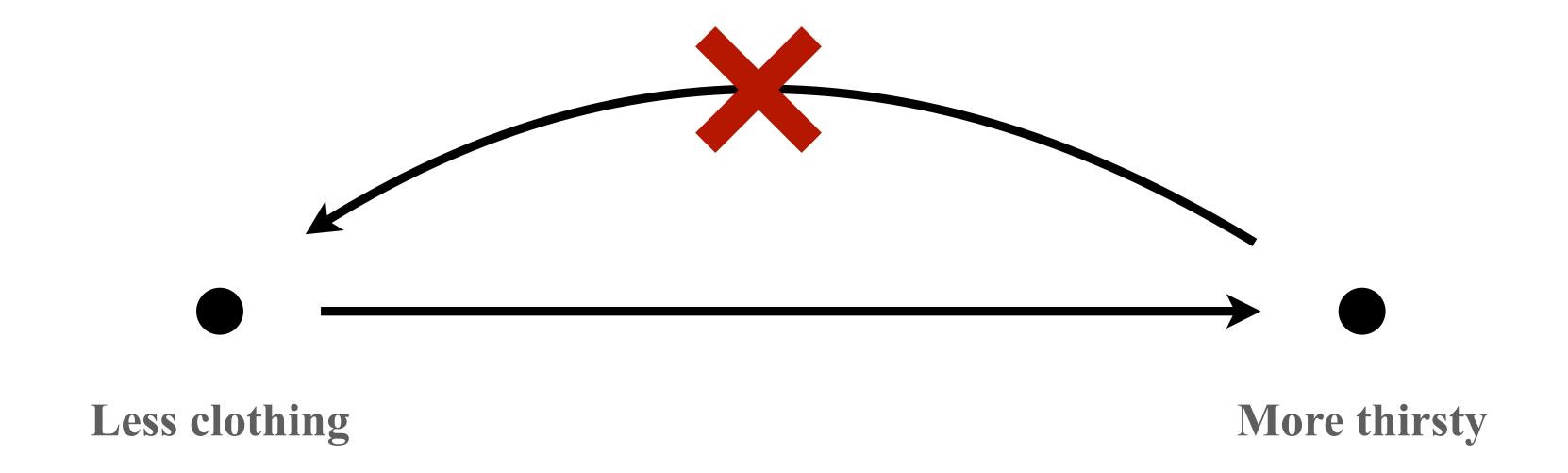
Less clothing

More thirsty

The lines have direction



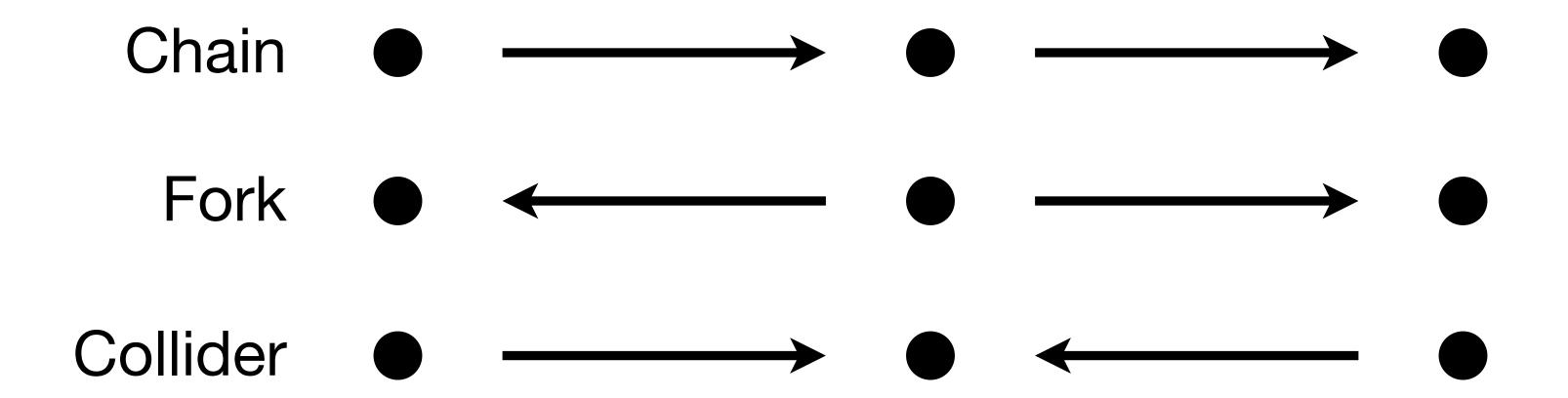
No cycles: you can't go backward, a thing cannot be its own cause



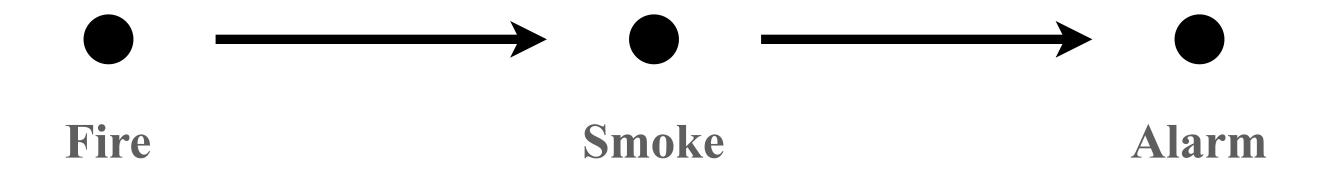
An arrow in the graph is a statement about a thought experiment

DAG anatomy

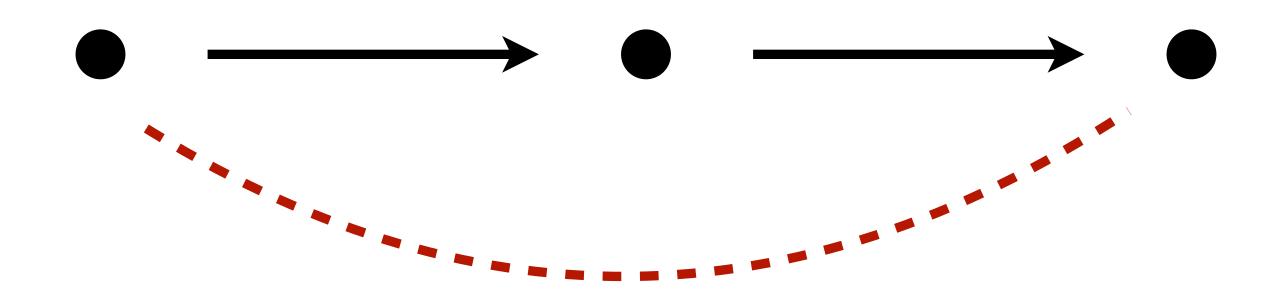
Three basic junctions



Chain: A causes B causes C

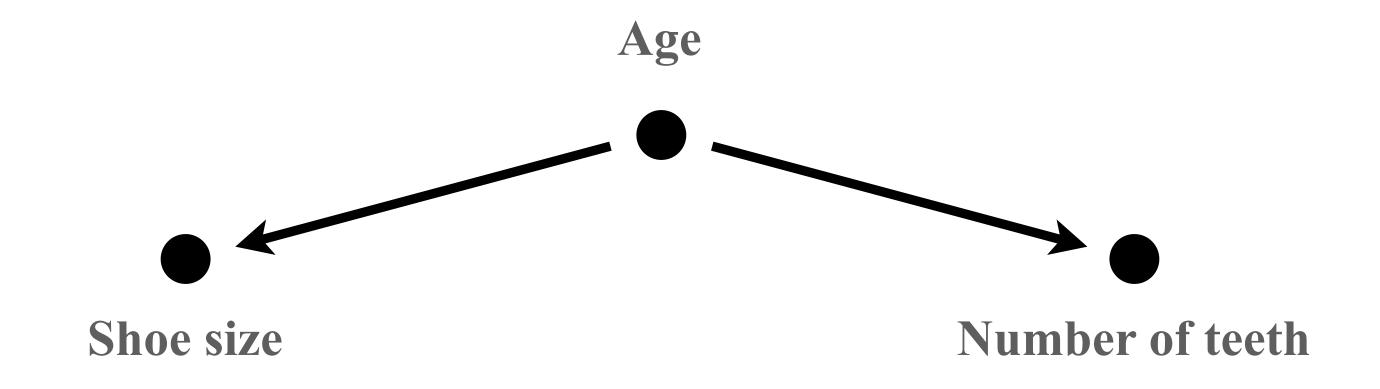


Chain: A causes B causes C

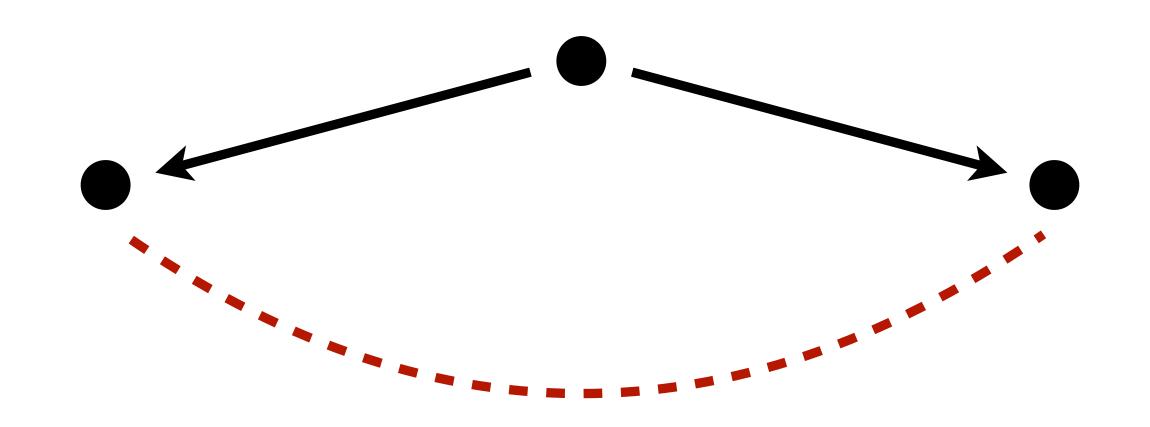


Transmits information: an alarm indicates fire, fire indicates alarm

Fork: your typical confounder

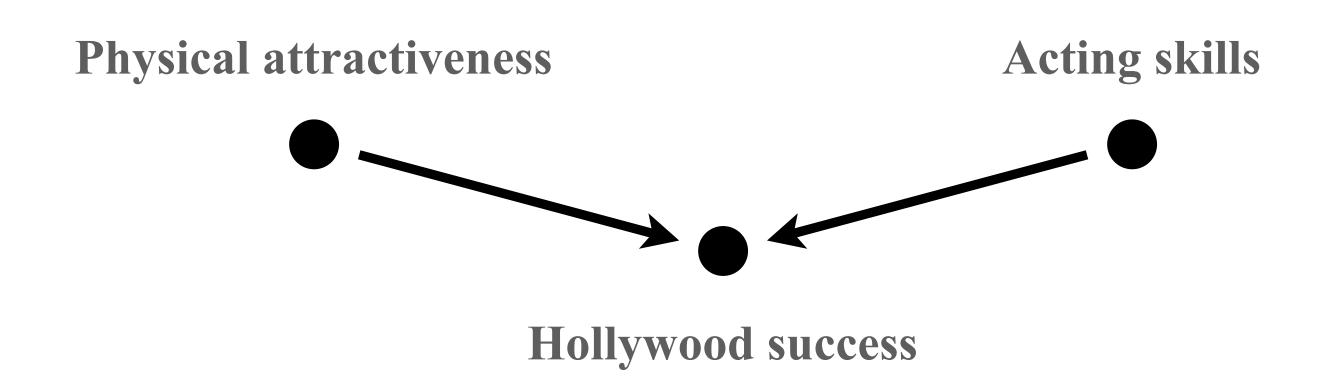


Fork: your typical confounder

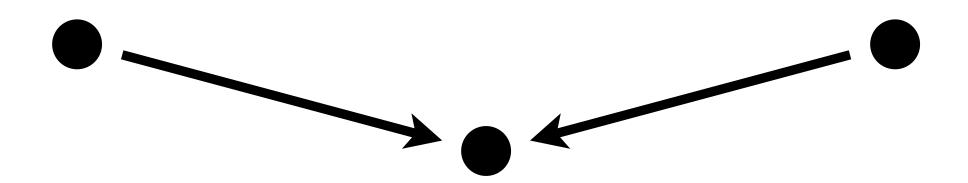


Transmits information: Few teeth indicates small shoes, small shoes indicates few teeth

Collider: a very interesting source of bias



Collider: a very interesting source of bias



Does <u>not</u> transmit information:

acting skills doesn't suggest attractiveness or vice versa

Puzzles: does information flow between A and B?

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$$A \longleftarrow \bullet \longleftarrow \bullet \longrightarrow \bullet \longrightarrow \bullet \longrightarrow B$$

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$$A \longleftarrow \bullet \longleftarrow \bullet \longleftarrow \bullet \longleftarrow \bullet \longrightarrow B$$

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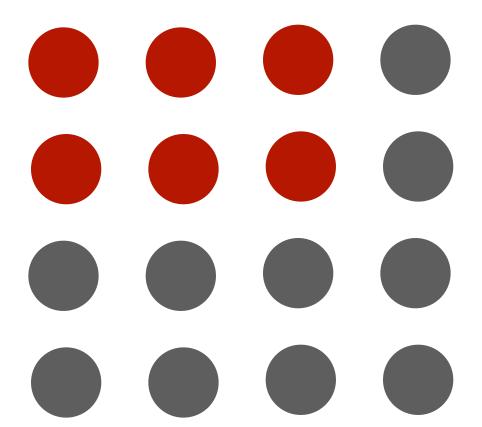
Probability intermission

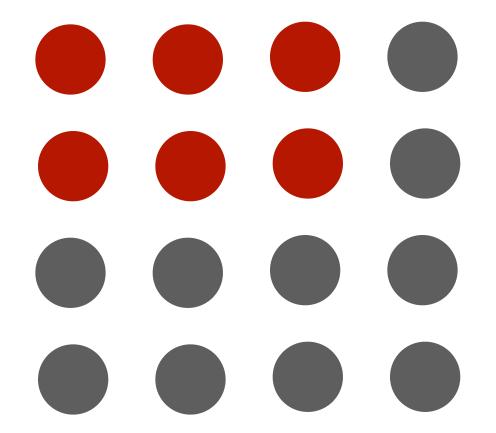
```
# interesting outcomes

# possible outcomes
```

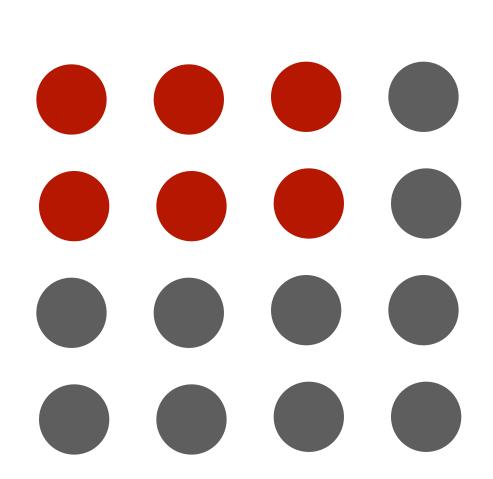
```
# interesting outcomes

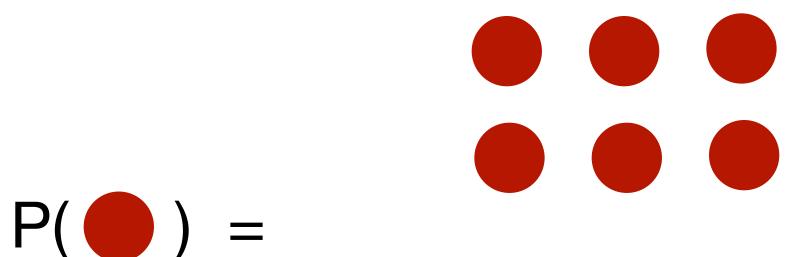
# possible outcomes
```

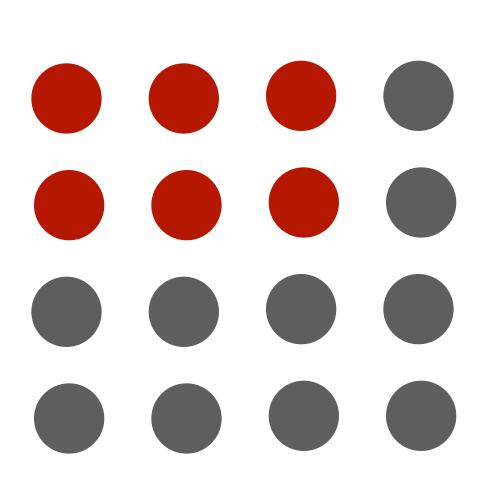


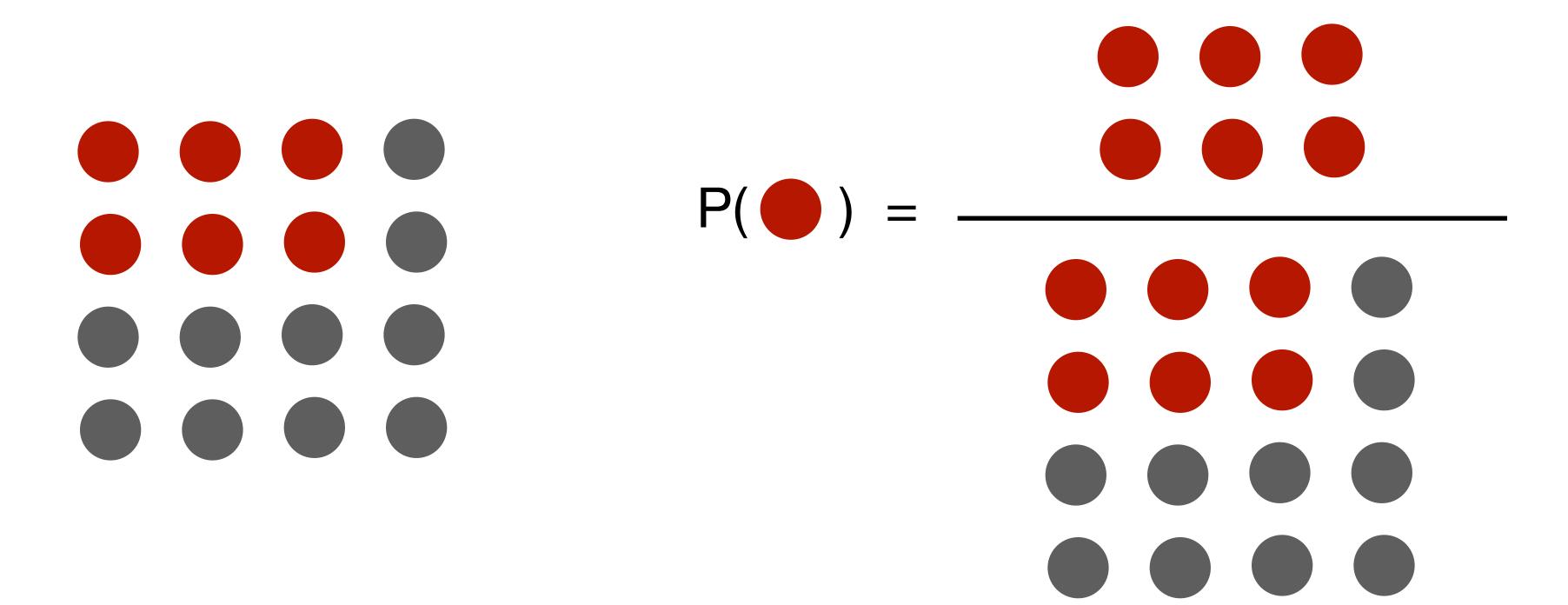


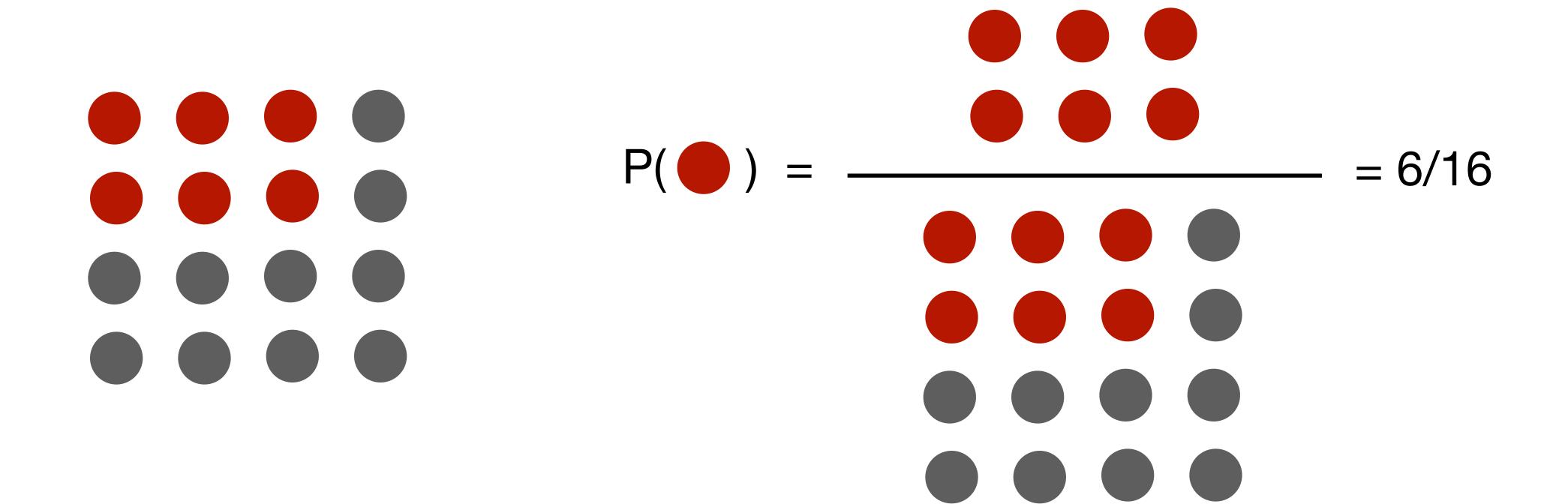
$$P(\bigcirc) =$$

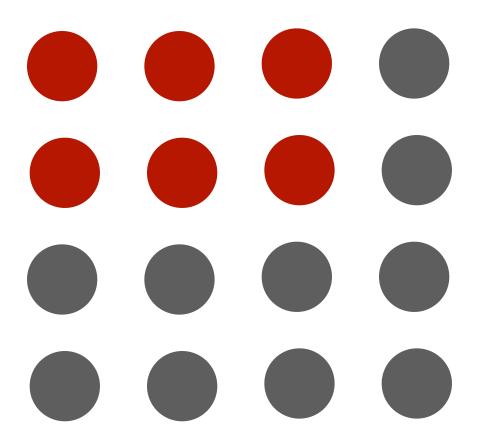


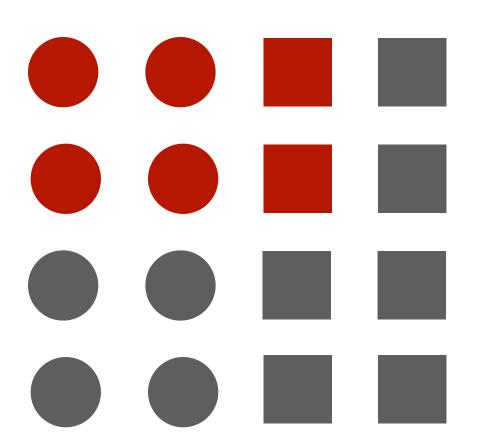


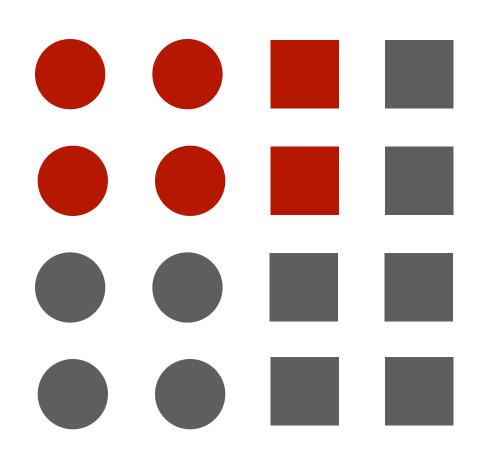




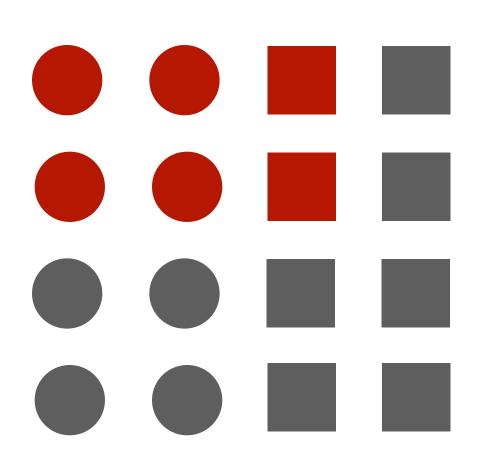


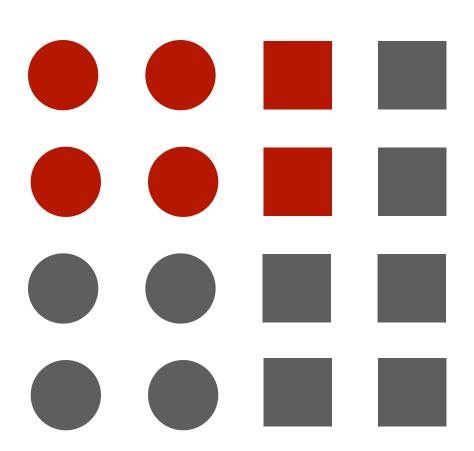


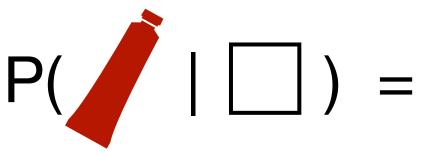


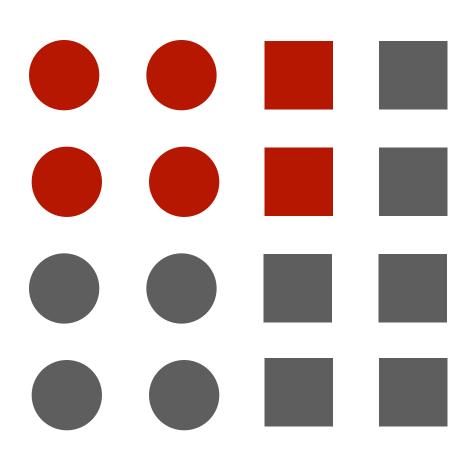


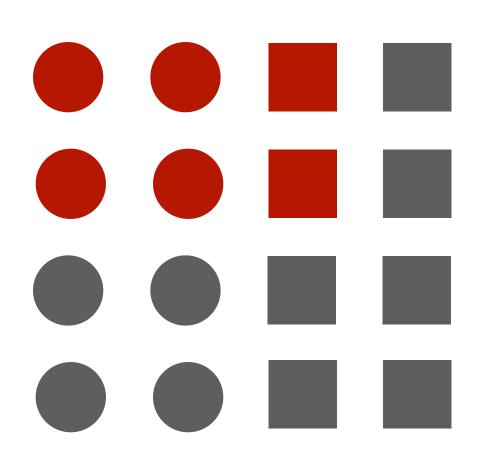


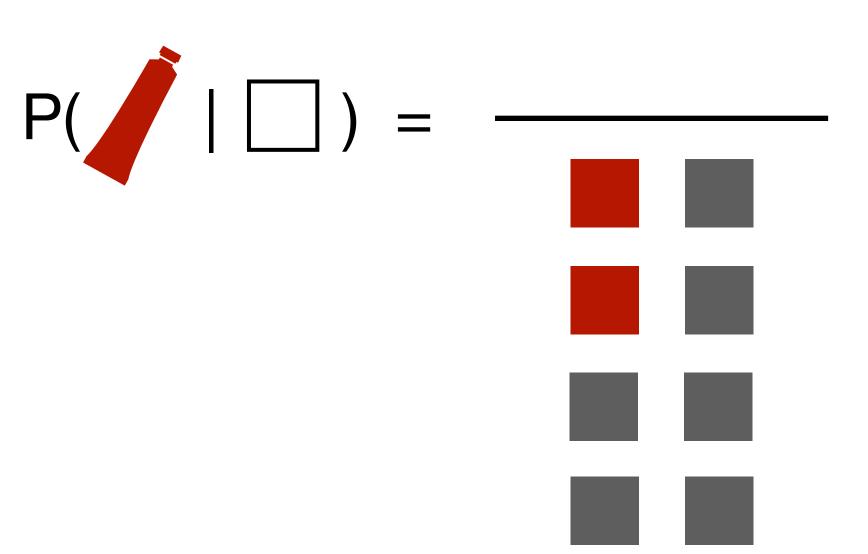


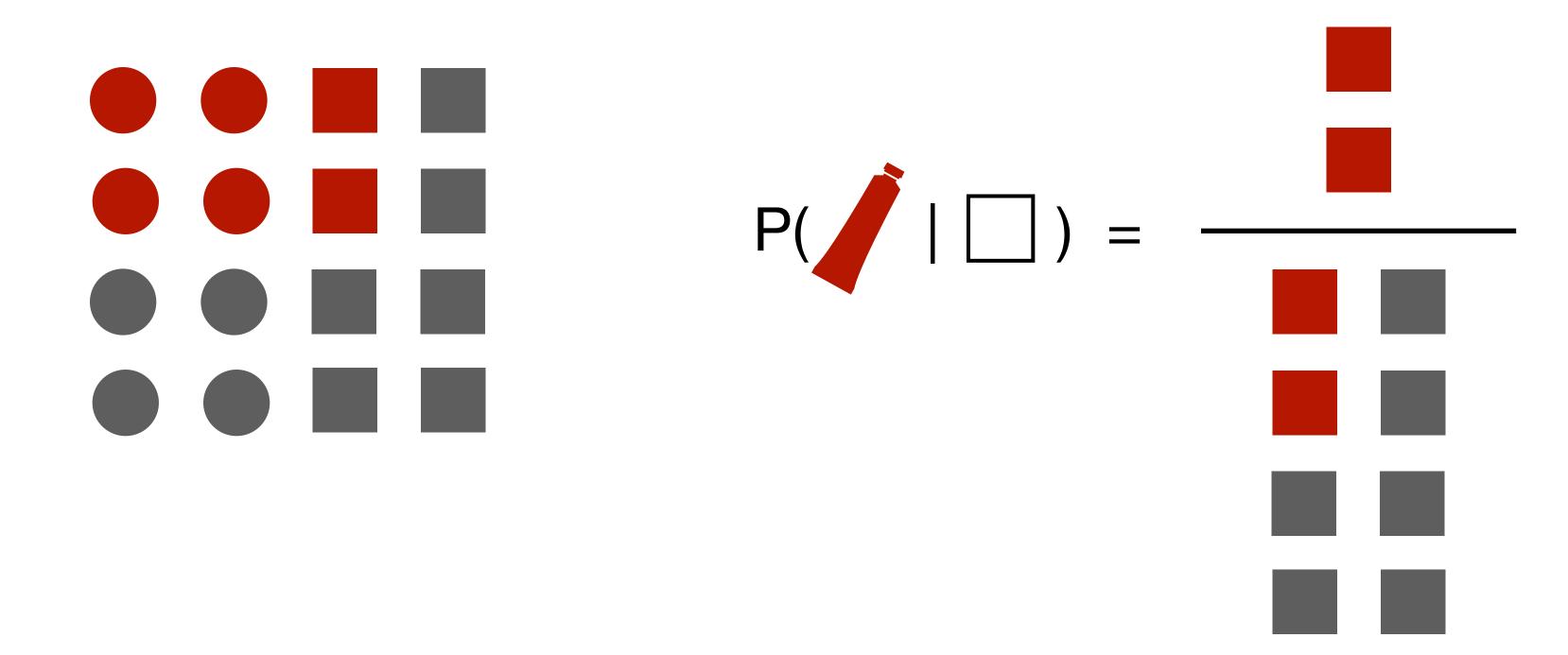


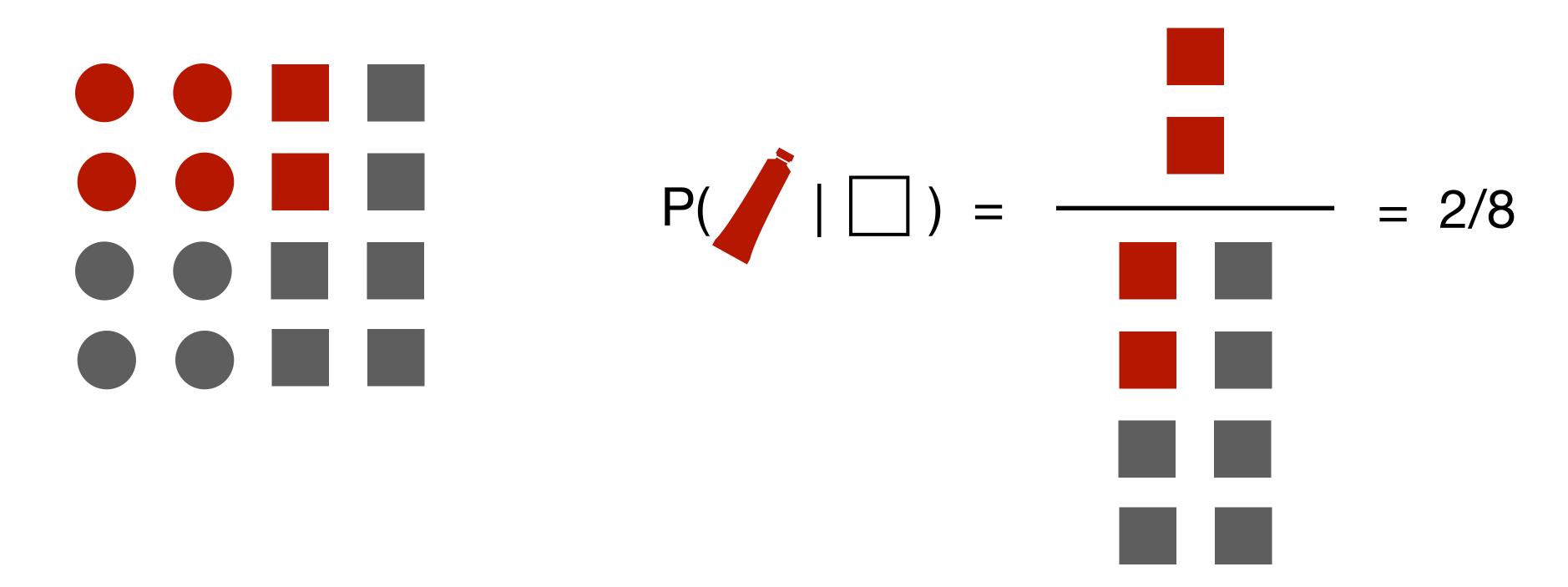


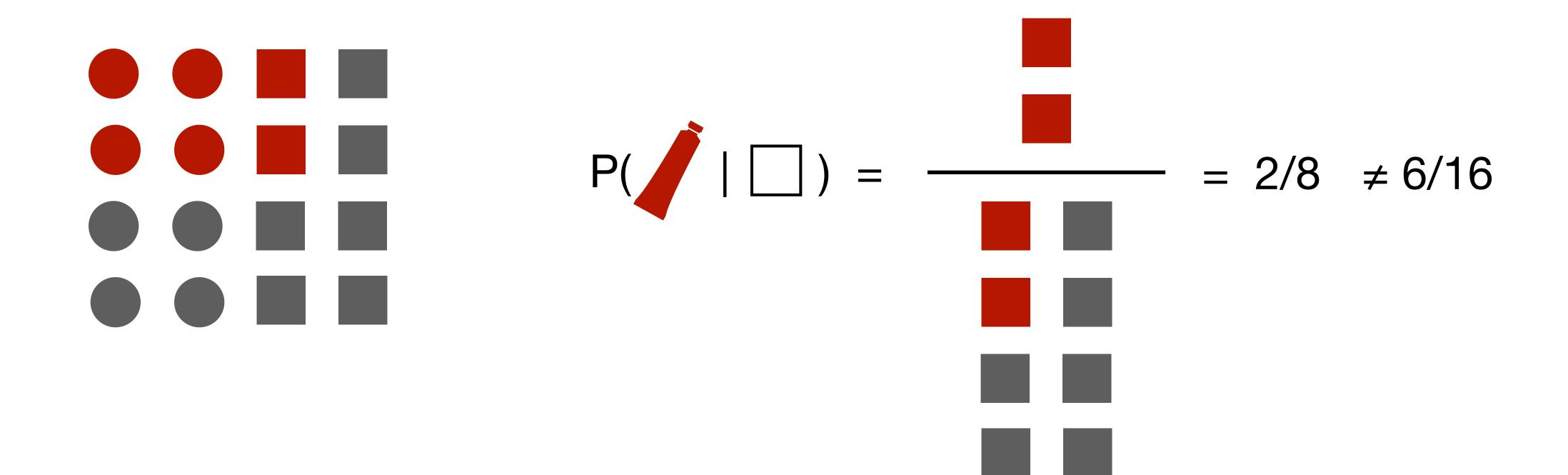






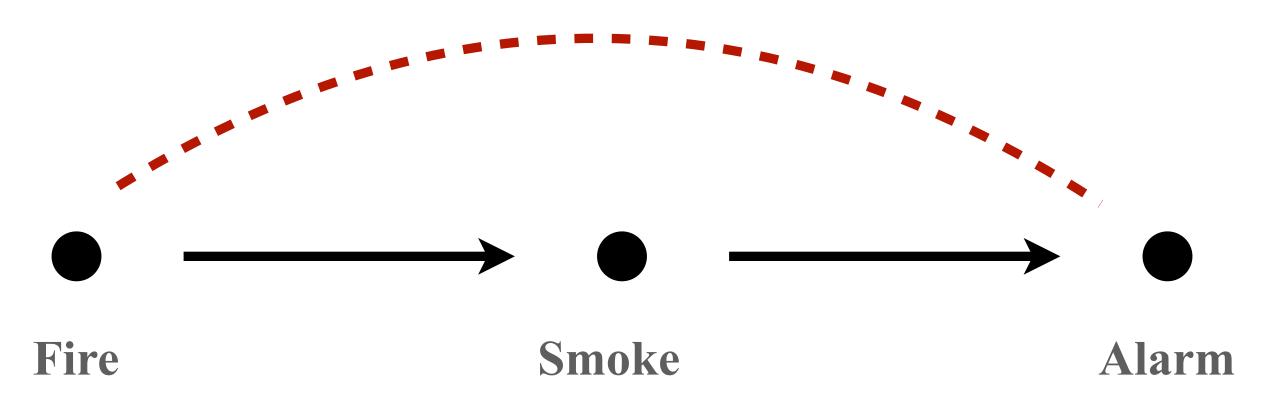






Conditioning alters the flow of information in the 3 basic junctions

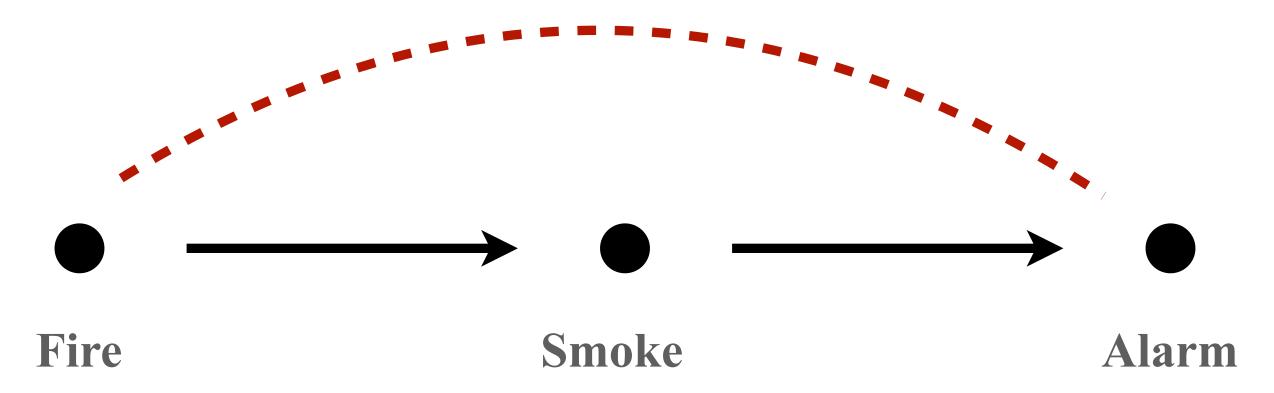
NB: the three data sets I use in this section are fake — I generated them with a computer to illustrate my points



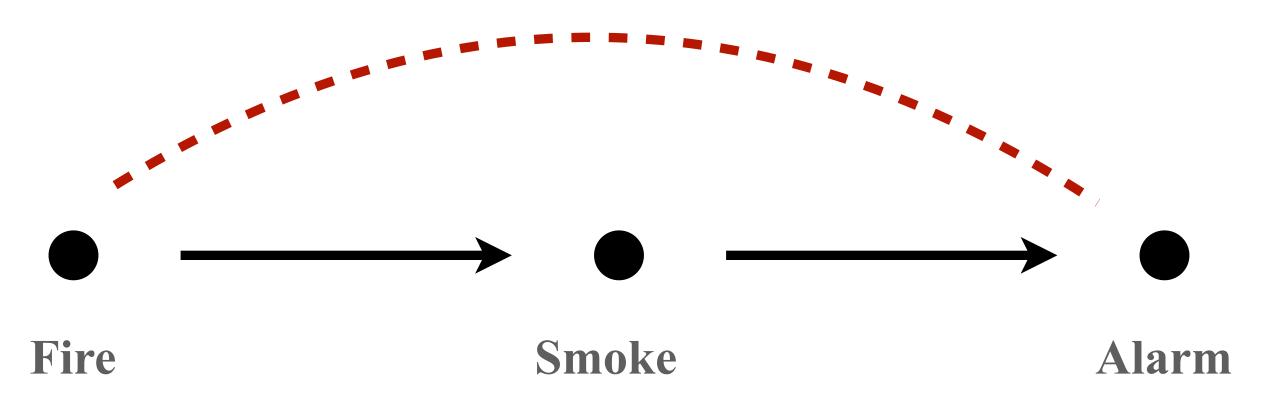
Smoke

Alarm

| | No alarm | Alarm |
|---------|----------|-------|
| No fire | 9338 | 153 |
| Fire | 9 | 500 |



| | No alarm | Alarm | P(fire) = |
|---------|----------|-------|-------------|
| No fire | 9338 | 153 | |
| Fire | 9 | 500 | |

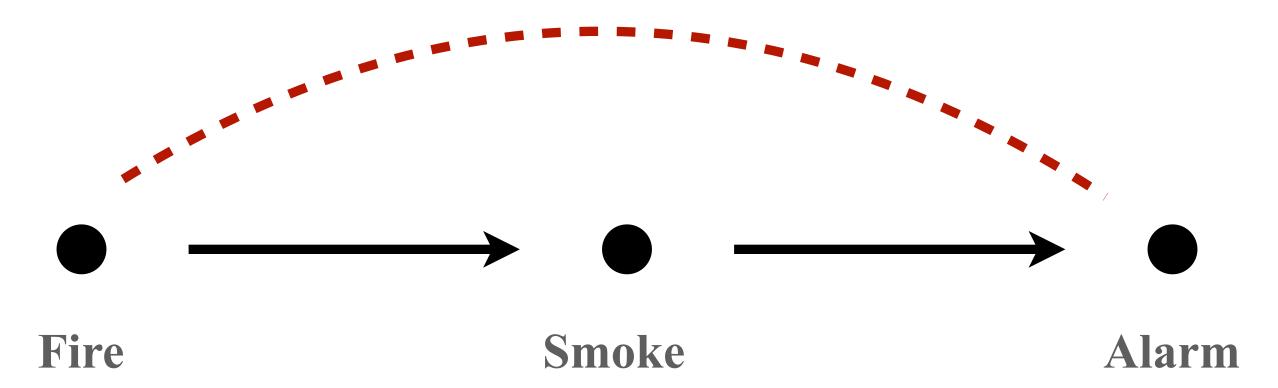


9 + 500

No alarm Alarm
P(fire) = ————
9338 153

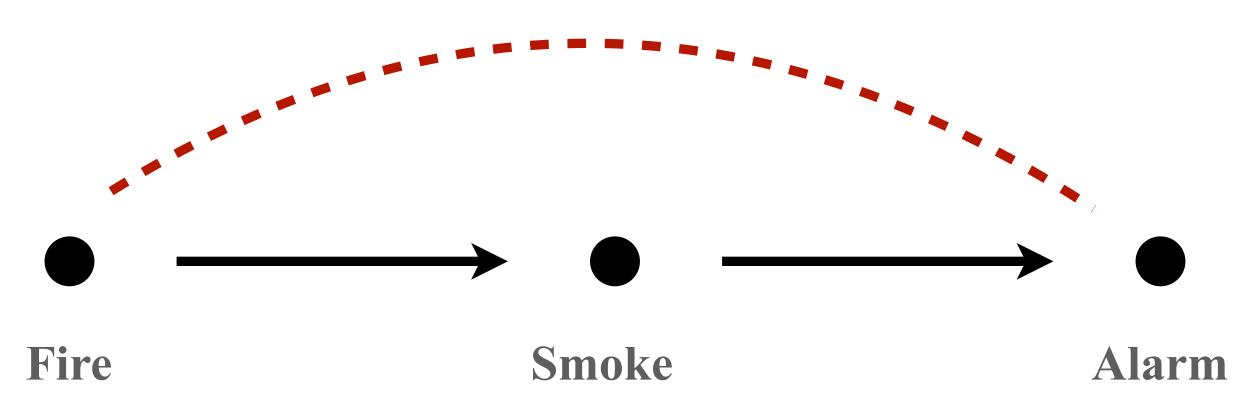
500

No fire



+ 153

| | No alarm | Alarm | P(fire) = | 9 + 500 |
|---------|----------|-------|-------------|----------------|
| No fire | 9338 | 153 | | 9 + 500 + 9338 |
| Fire | 9 | 500 | | |

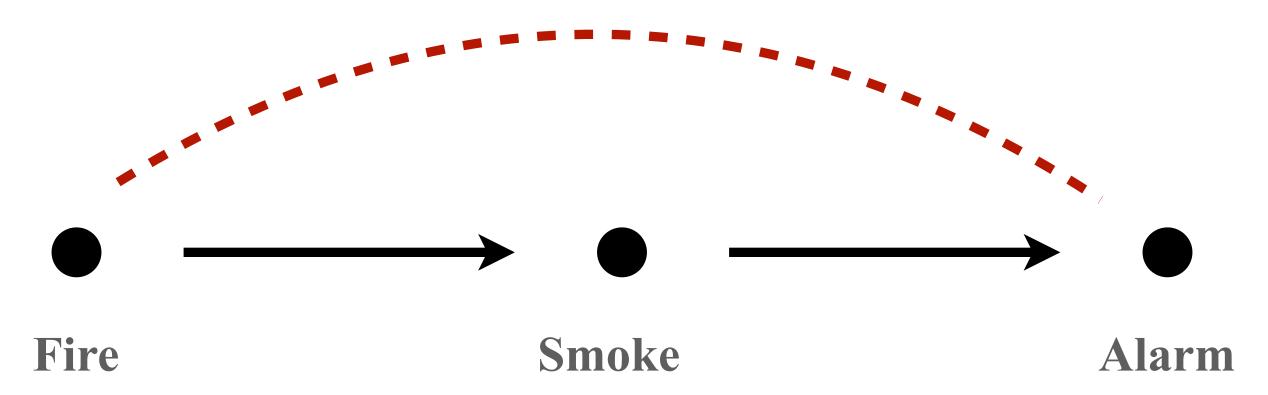


509

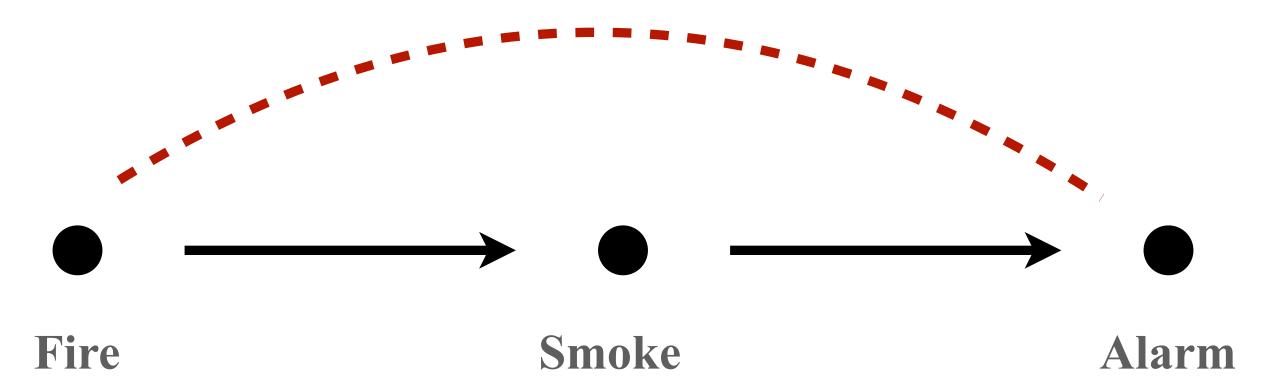
10 000

≈ **0.05**

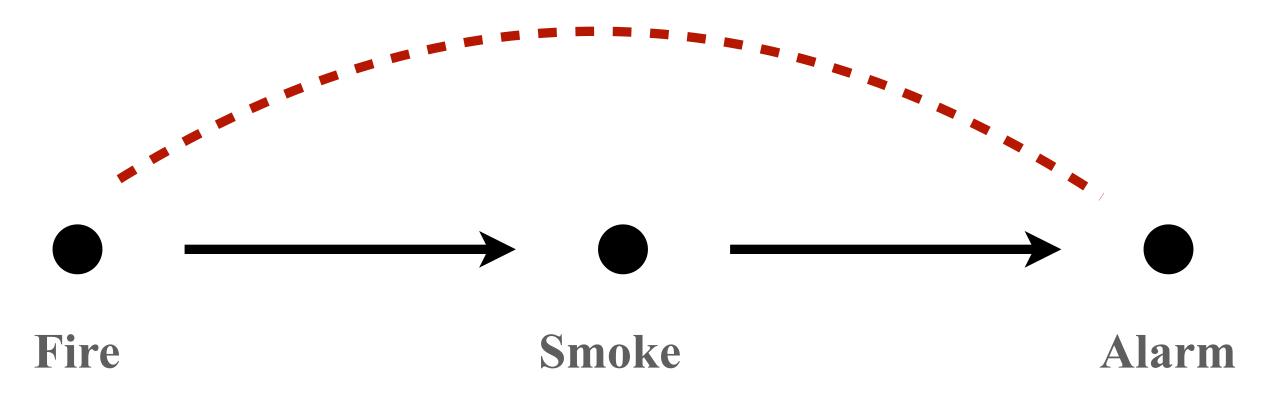
| | No alarm | Alarm | P(fire) = $\frac{9 + 500}{}$ | |
|---------|----------|-------|--------------------------------|---|
| No fire | 9338 | 153 | 9 + 500 + 9338 + 153 | 3 |
| Fire | 9 | 500 | | |



| | No alarm | Alarm | P(fire) = $\frac{9 + 500}{}$ = $\frac{509}{}$ ≈ 0.05 |
|----------|----------|-------|--|
| NIa fira | 0220 | 150 | 9 + 500 + 9338 + 153 10 000 |
| No fire | 9338 | 153 | P(fire alarm) = |
| Fire | 9 | 500 | i (iii o aiaiiii) — |

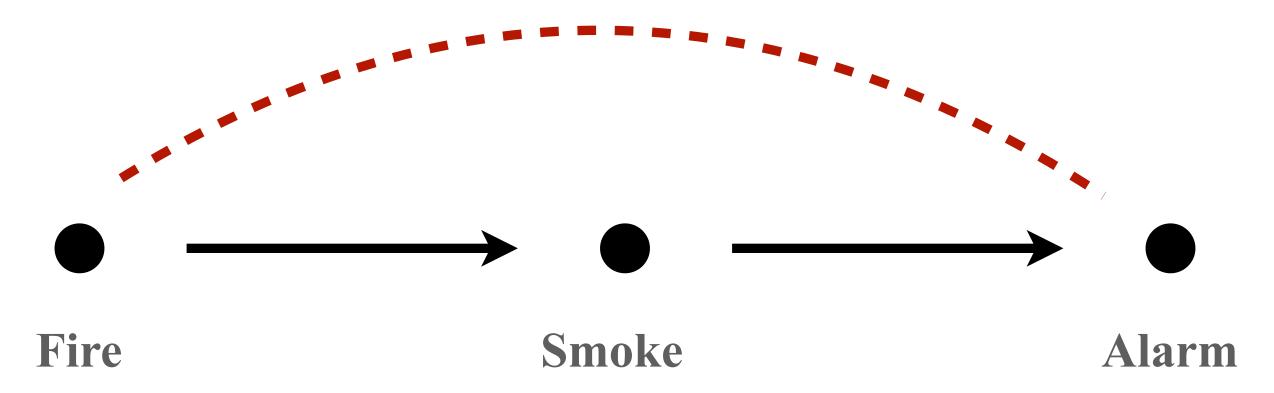


P(fire) =
$$\frac{9 + 500}{9 + 500 + 9338 + 153} = \frac{509}{10000} \approx 0.05$$



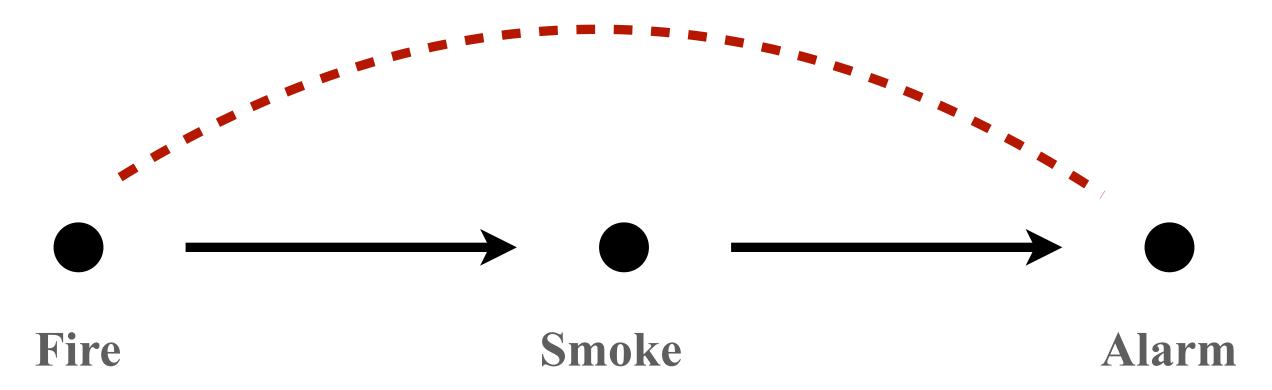
P(fire) =
$$\frac{9 + 500}{9 + 500 + 9338 + 153} = \frac{509}{10000} \approx 0.05$$

No fire



P(fire) =
$$\frac{9 + 500}{9 + 500 + 9338 + 153} = \frac{509}{10000} \approx 0.05$$

P(fire | alarm) =
$$\frac{500}{500 + 153}$$



P(fire) =
$$\frac{9 + 500}{9 + 500 + 9338 + 153} = \frac{509}{10000} \approx 0.05$$

No fire

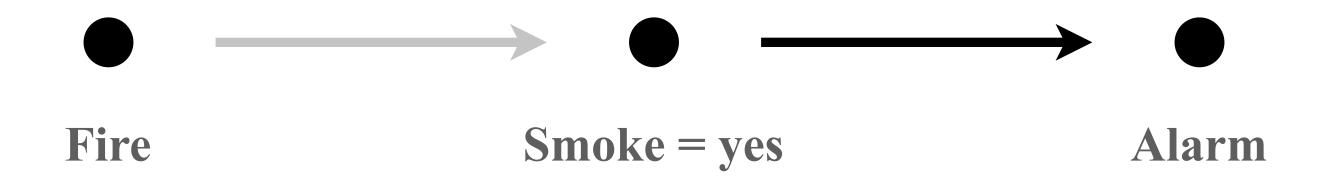
153

500

P(fire | alarm) =
$$\frac{500}{500 + 153} \approx 0.77$$

Transmits info Smoke Transmits info Alarm

POINT: hearing the alarm gives us information about the likelihood of any ongoing fires



| | No alarm | Alarm |
|---------|----------|-------|
| No fire | 9338 | 153 |
| Fire | 9 | 500 |

| | No alarm | Alarm | No alarm | Alarm |
|---------|----------|---------|----------|---------|
| No fire | 1 | 92 | 9337 | 61 |
| Fire | 6 | 500 | 3 | 0 |
| | Smoke | e = yes | Smo | ke = no |

| | No alarm | Alarm | P(fire | smoke)= |
|---------|----------|---------|--------|---------|
| No fire | 1 | 92 | | |
| Fire | 6 | 500 | | |
| | Smoke | e = yes | | |

| | No alarm | Alarm | P(fire smoke) = 506/599 |
|-------------|----------|-------|---------------------------|
| No fire | 1 | 92 | |
| Fire | 6 | 500 | |
| Smoke = yes | | | |

| | No alarm | Alarm | P(fire smoke) = $506/599 \approx 0.84$ |
|---------|----------|---------|--|
| No fire | 1 | 92 | |
| Fire | 6 | 500 | |
| | Smok | e = yes | |

| | No alarm | Alarm | P(fire smoke) = $506/599 \approx 0.84$ |
|---------|----------|---------|--|
| No fire | 1 | 92 | P(fire alarm, smoke) = |
| Fire | 6 | 500 | |
| | Smoke | e = yes | |

No alarm Alarm P(fire | smoke) =
$$506/599 \approx 0.84$$

No fire 92 P(fire | alarm, smoke) =

Smoke = yes

500

No alarm Alarm P(fire | smoke) = $506/599 \approx 0.84$ No fire 92 P(fire | alarm, smoke) = 500/592

Smoke = yes

500

No alarm Alarm $P(\text{ fire } | \text{ smoke }) = 506/599 \approx 0.84$ $P(\text{ fire } | \text{ alarm, smoke }) = 500/592 \approx 0.84$ $P(\text{ fire } | \text{ alarm, smoke }) = 500/592 \approx 0.84$

Smoke = yes

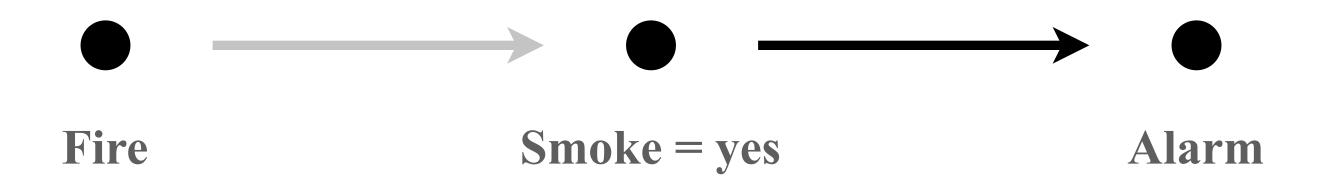
500

No alarm Alarm $P(\text{ fire } | \text{smoke}) = 506/599 \approx 0.84$

Conditional on smoke, the alarm gives no extra information!

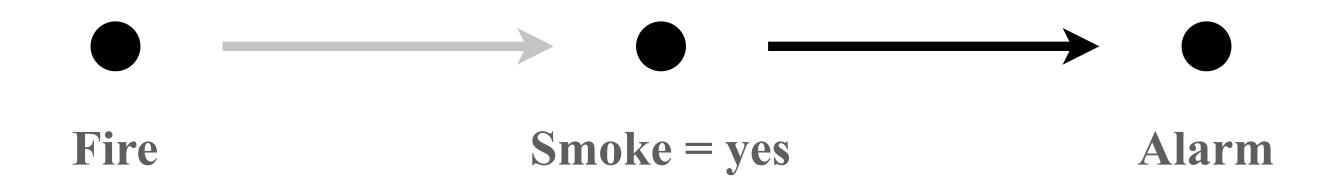
Fire 6 50

Smoke = yes



Conditional on smoke, the alarm gives no extra information!

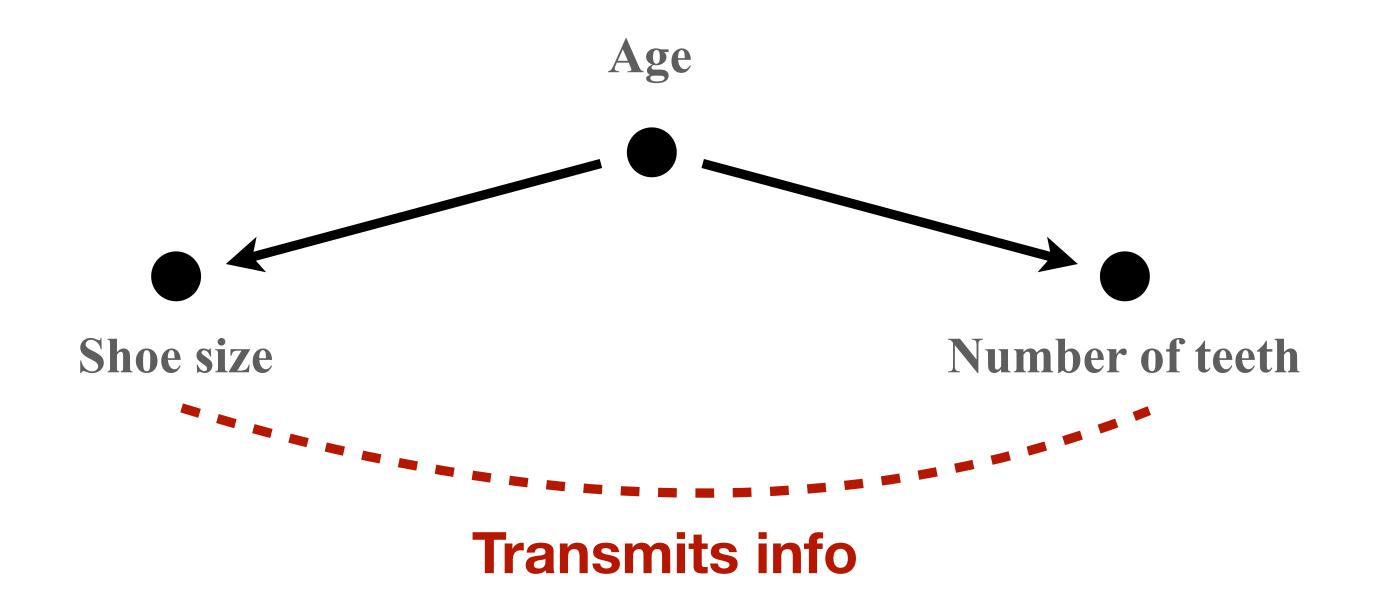
If I knew nothing about fire, smoke, or alarms, and analyzed these data haphazardly, I would conclude that smoke alarms are useless

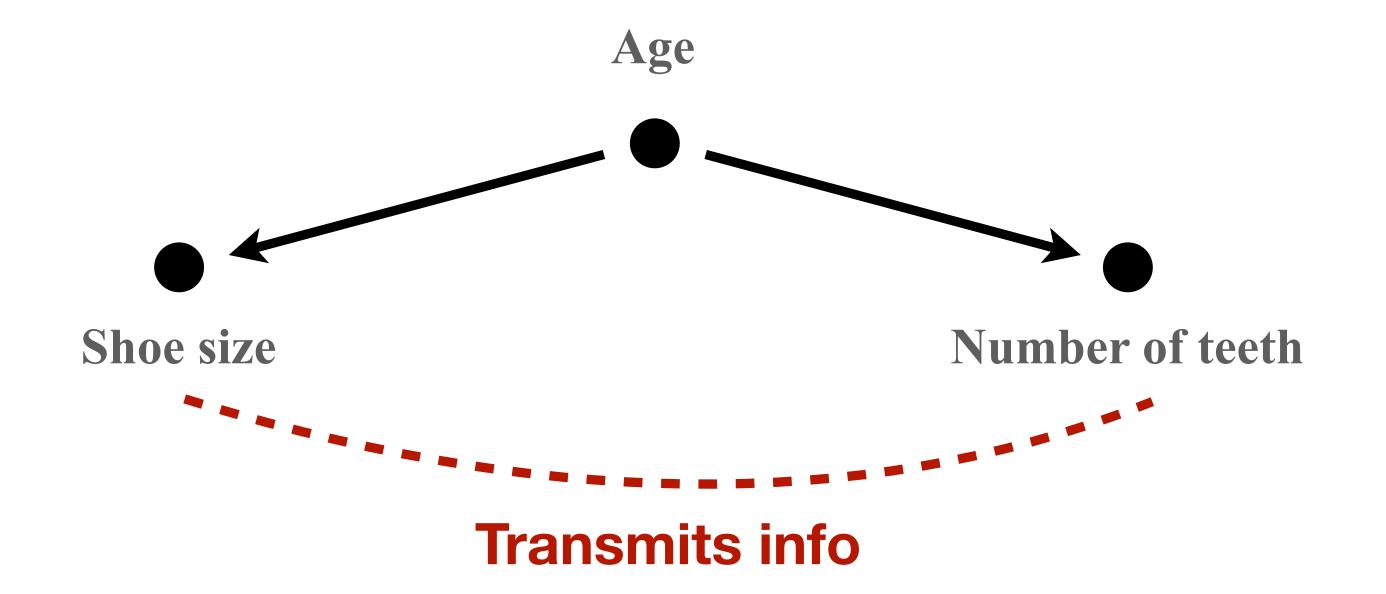


Conditional on smoke, the alarm gives no extra information!

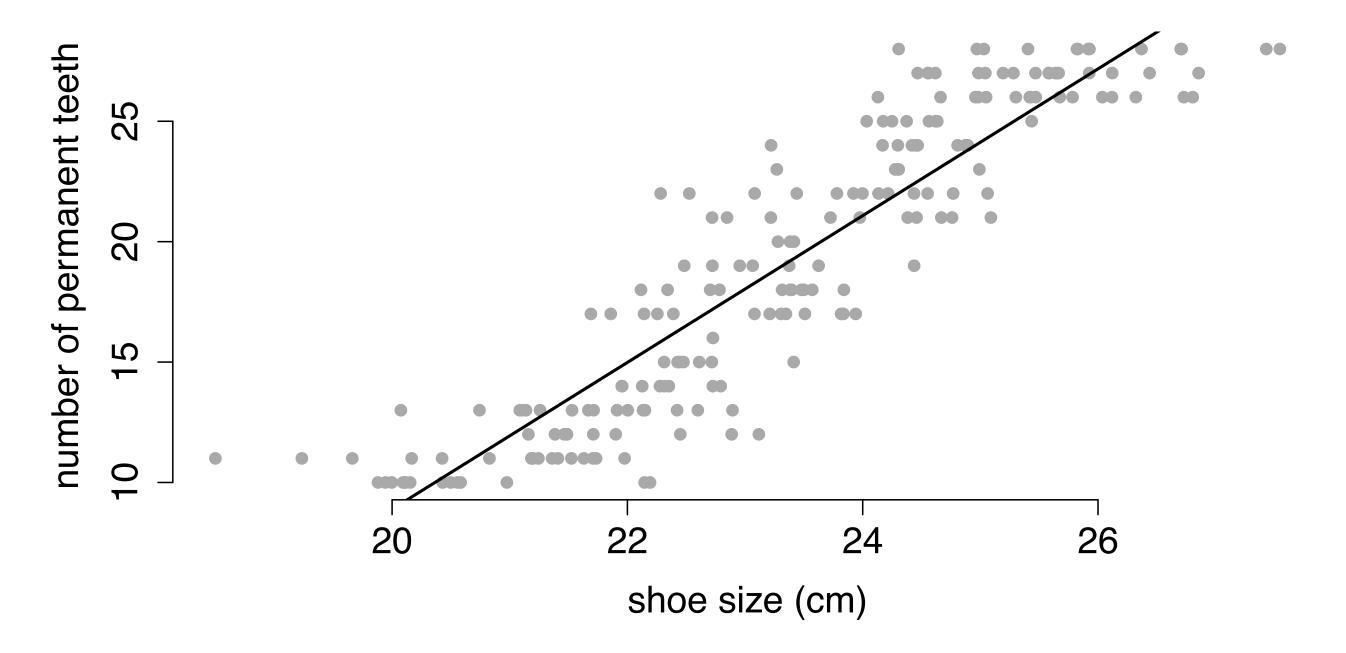
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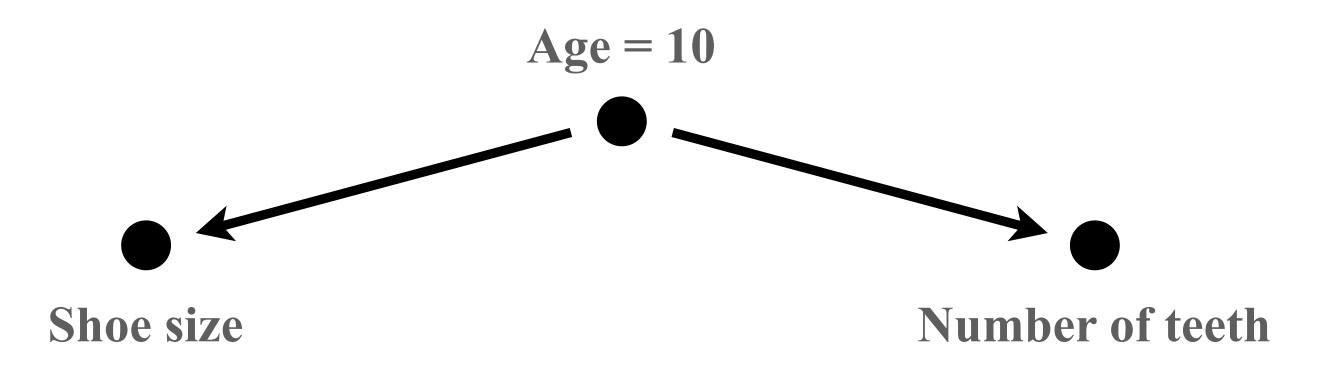
"Conditioning on a mediator" — adjusting away the effect of interest



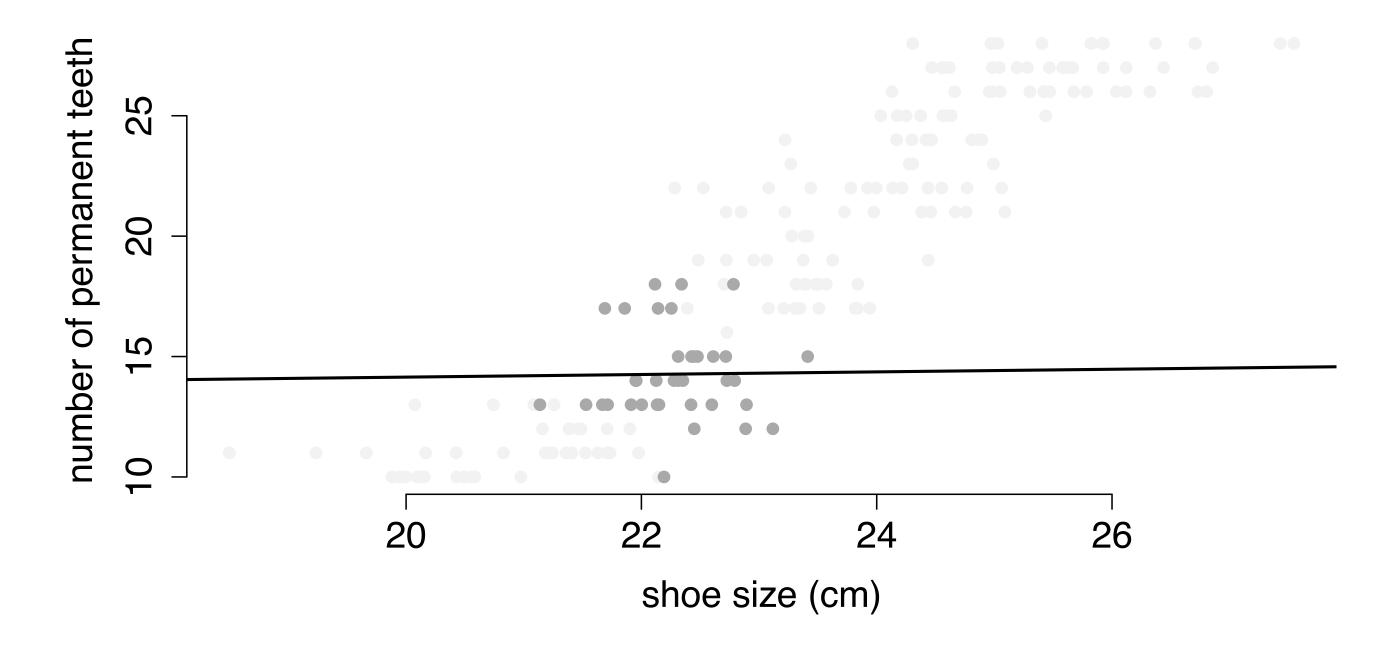


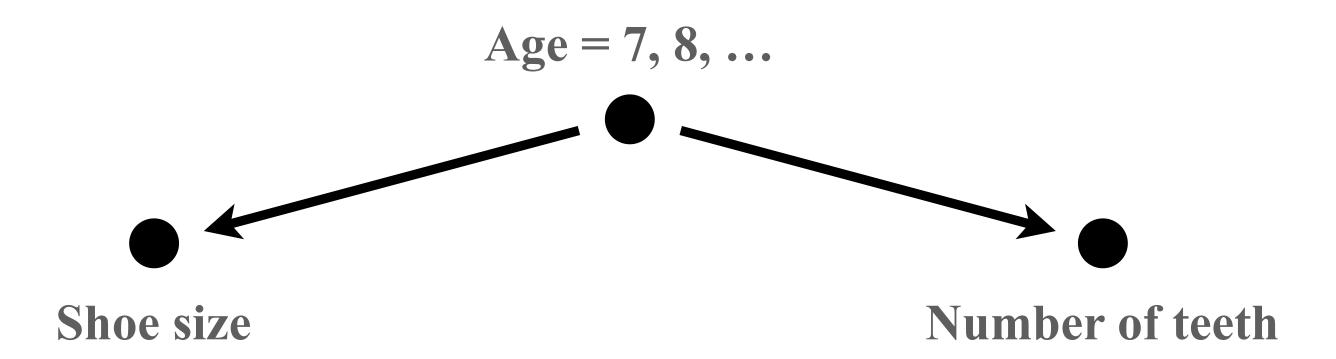
teeth vs shoe size, kids aged 7-15



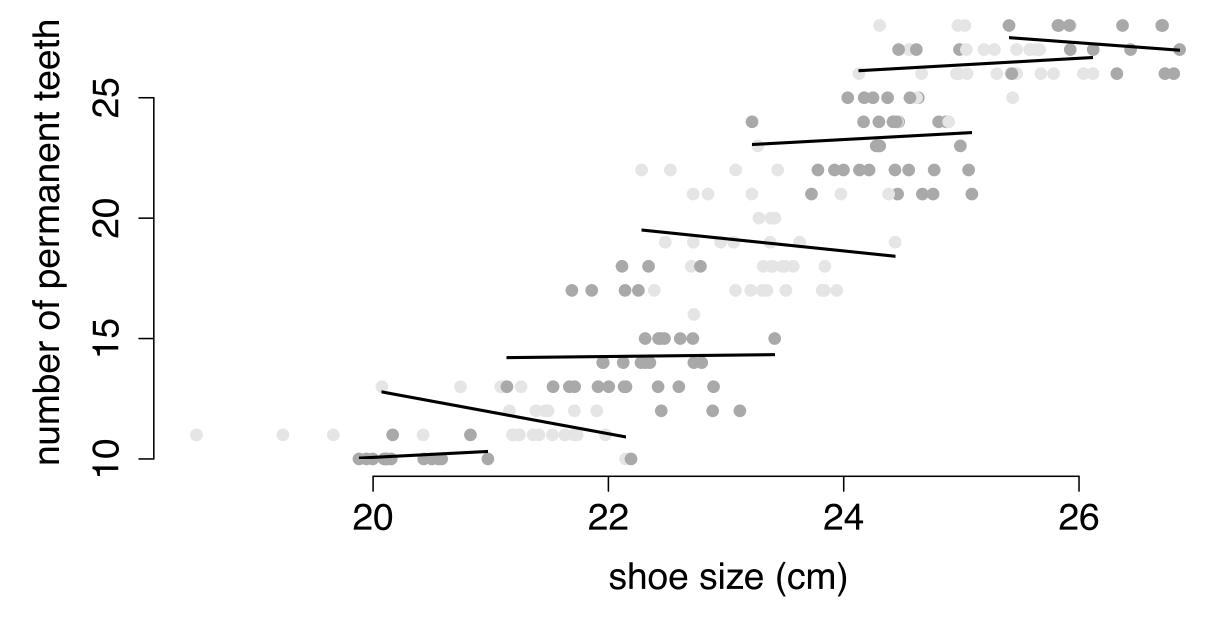


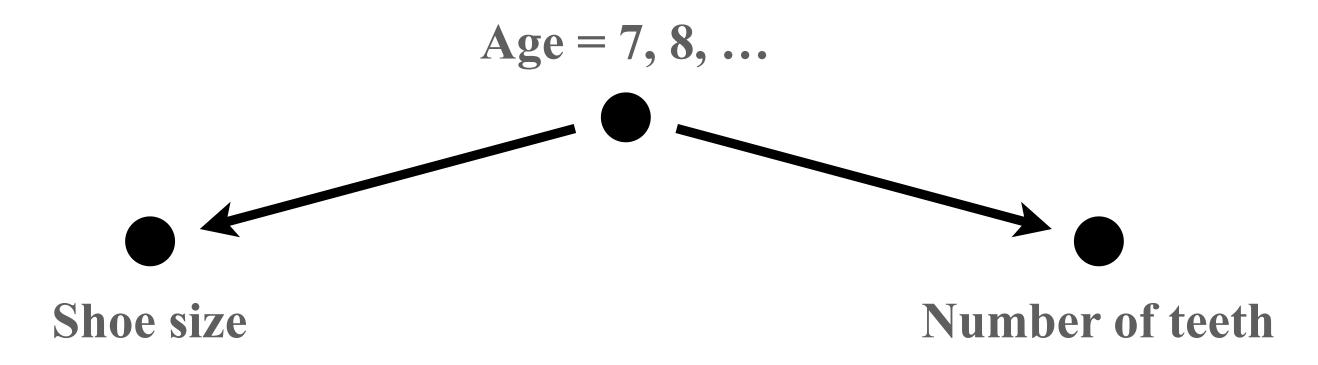
teeth vs shoe size, kids aged 10





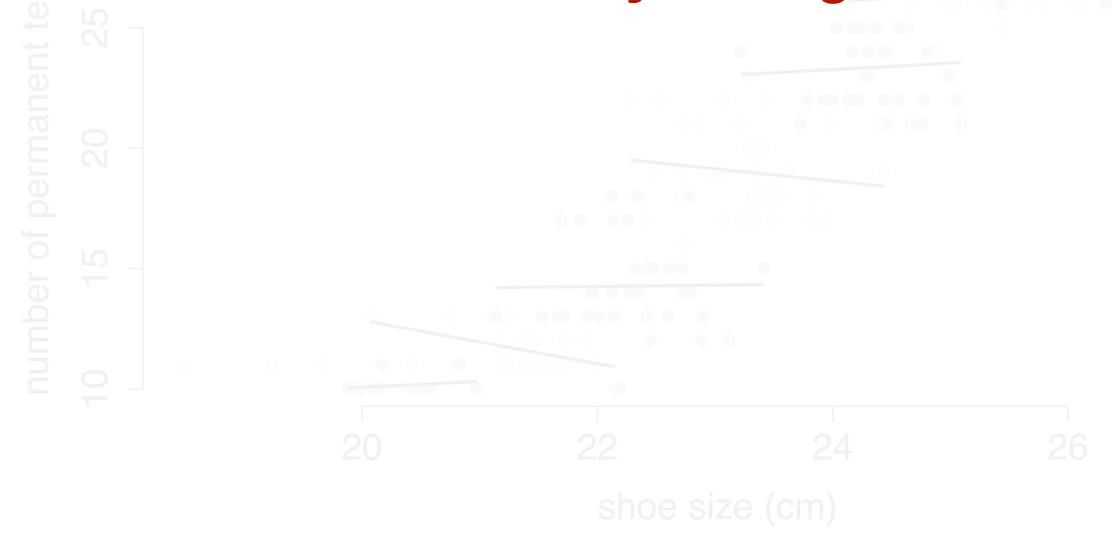


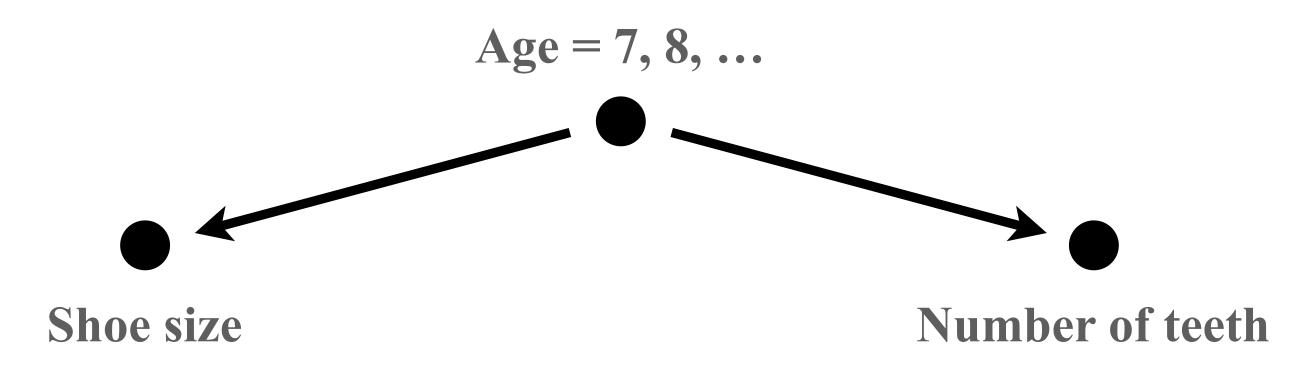




teeth vs shoe size, regressions conditional on age

Number of teeth informs on shoe size only through their mutual association with age

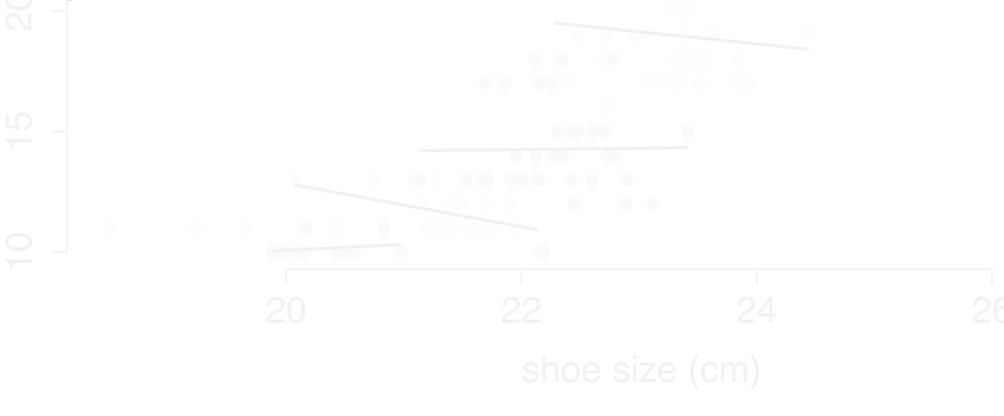


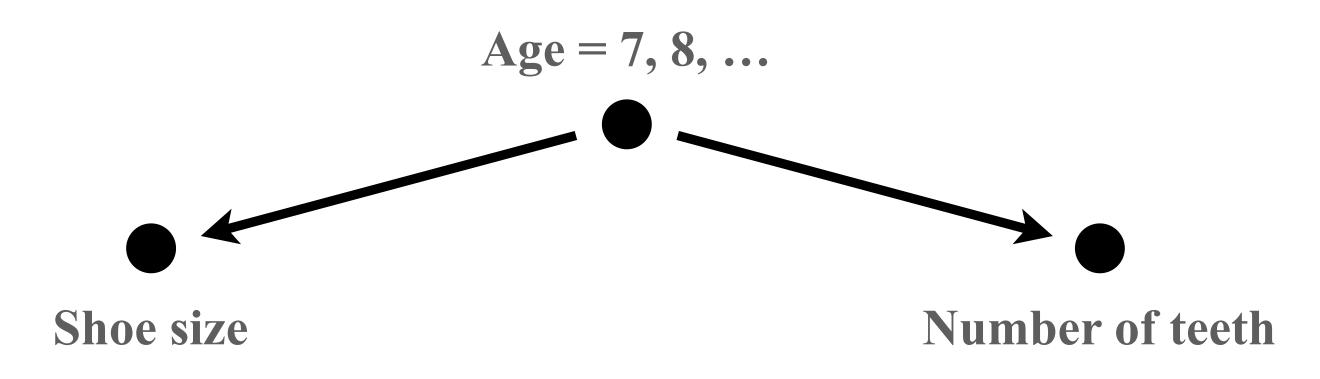


teeth vs shoe size, regressions conditional on age

Number of teeth informs on shoe size only through their mutual association with age

Controlling age by fixing it to some value blocks the "information flow"





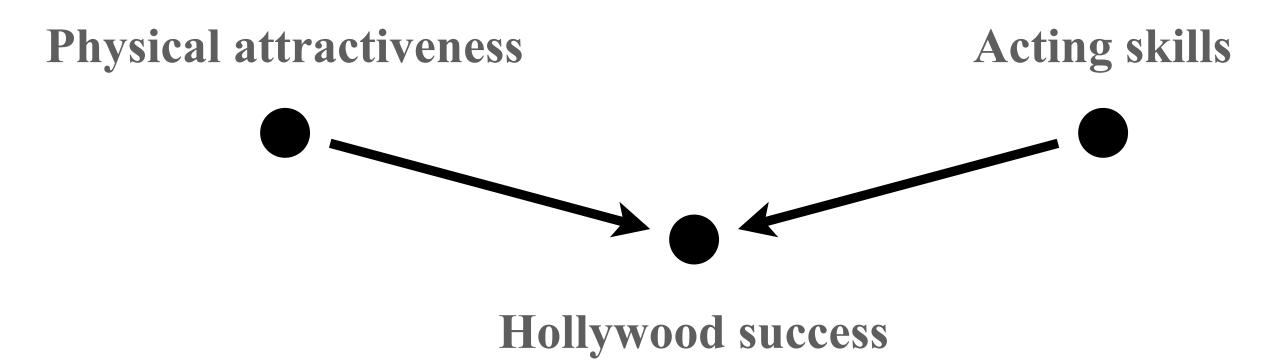
Does not transmit info

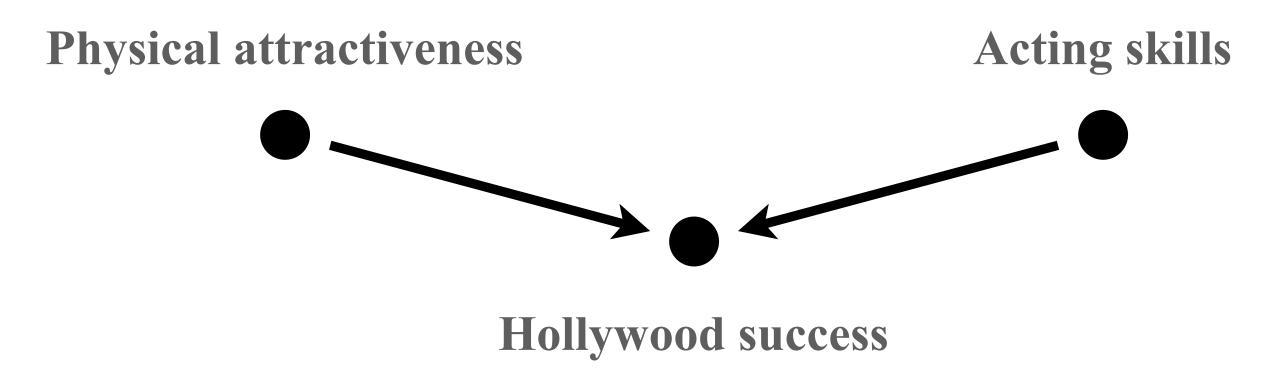
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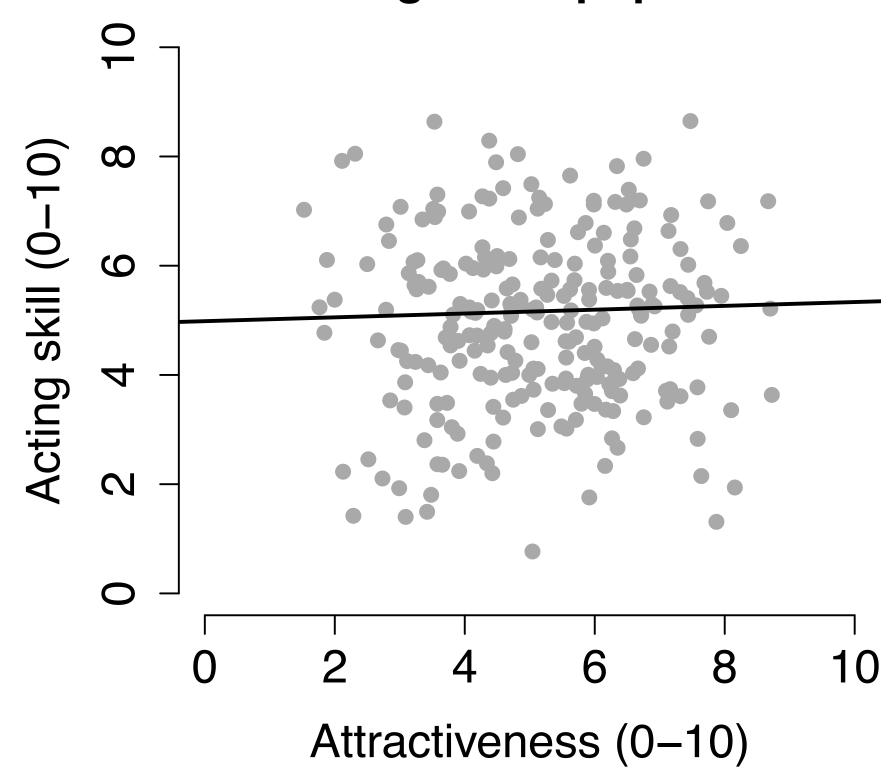
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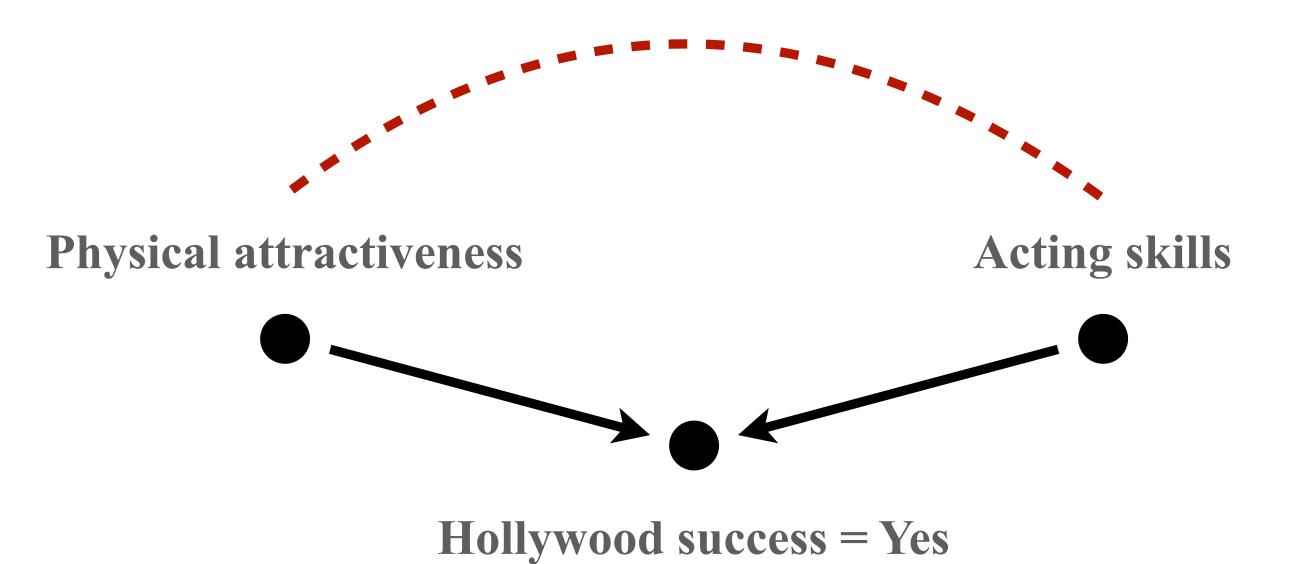
Ignoring age leaves "uncontrolled confounding"



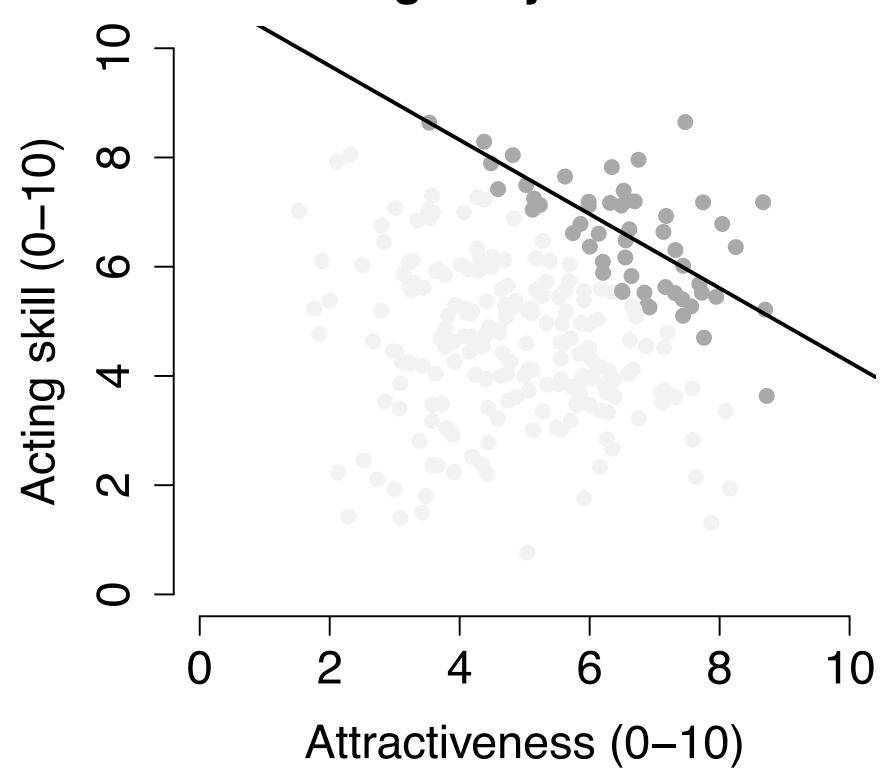


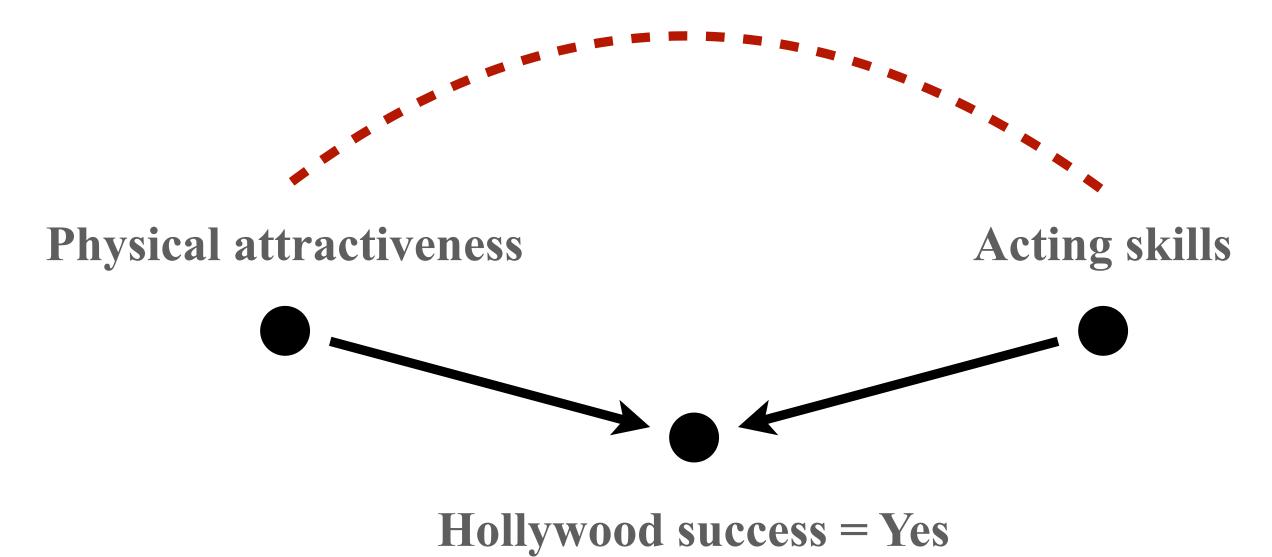
Acting skill vs. attractiveness in the general population





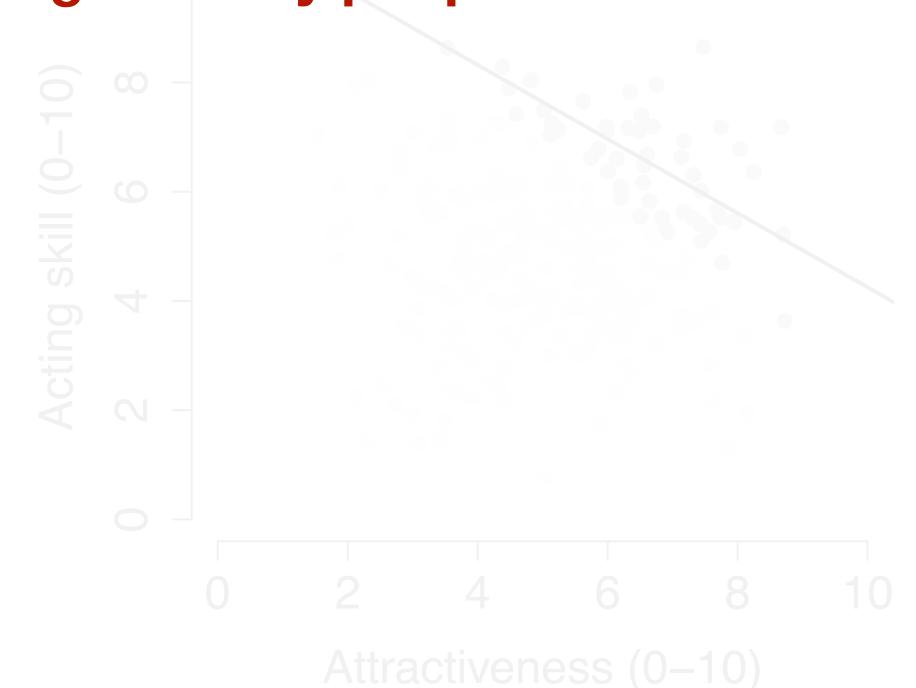
Acting skill vs. attractiveness among Hollywood stars

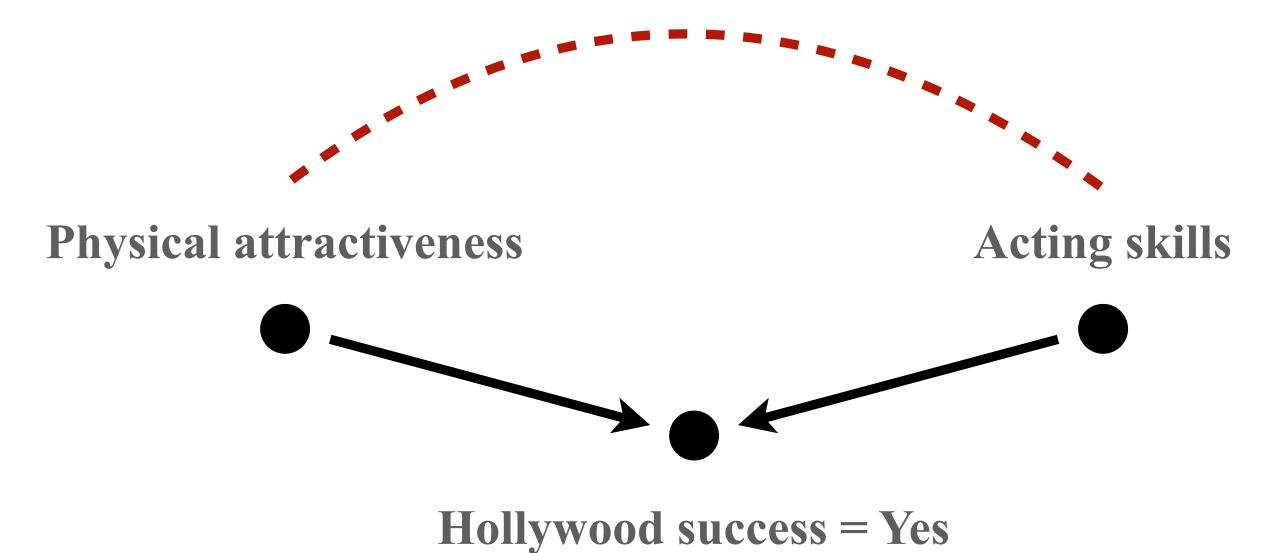




Acting skill vs. attractiveness

Acting skill and attractiveness not related among ordinary people

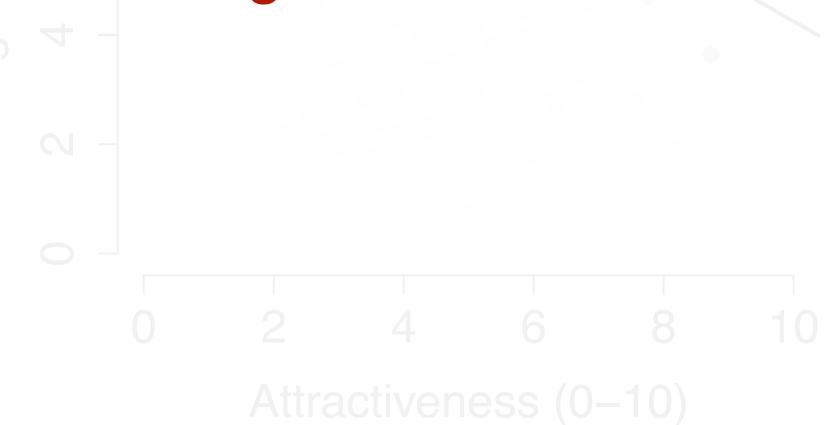


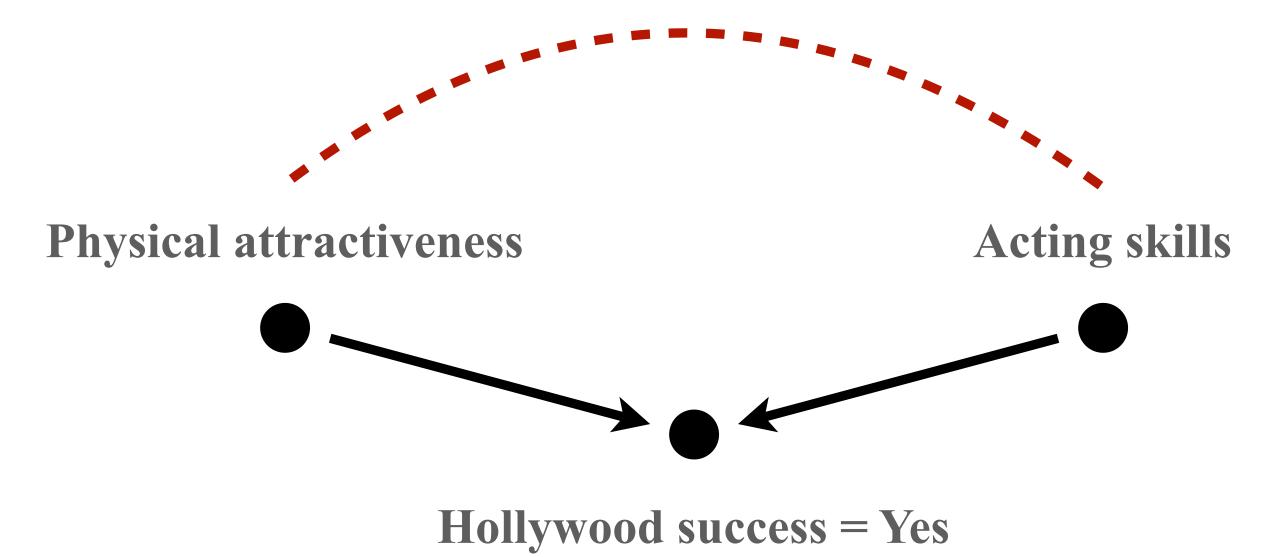


Acting skill vs. attractiveness

Acting skill and attractiveness not related among ordinary people

Very high attractiveness or very high acting skill alone enough to ensure success





Acting skill vs. attractiveness

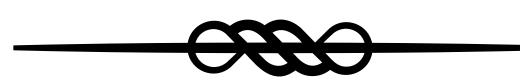
Acting skill and attractiveness not related among ordinary people

Very high attractiveness or very high acting skill alone enough to ensure success

Conditioning on collider called "collider bias:" introduces an effect where there should be none

Attractiveness (0–10)

break probably



The Marko fallacy: confounding

Lessons so far

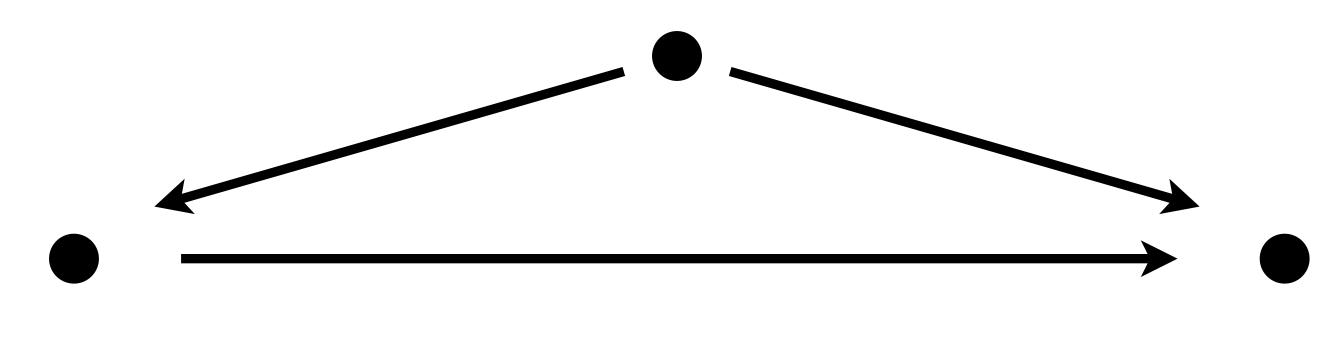
- The three basic junctures are like "information pipes"
- The chain and the fork are open, information flows through them
- The collider is closed, information does not flow through
- Conditioning on the middle node of a chain or fork closes the pipe
- Conditioning on the middle node of a collider opens the pipe

Please suggest a more sane model by applying one of the basic junctions to this mess

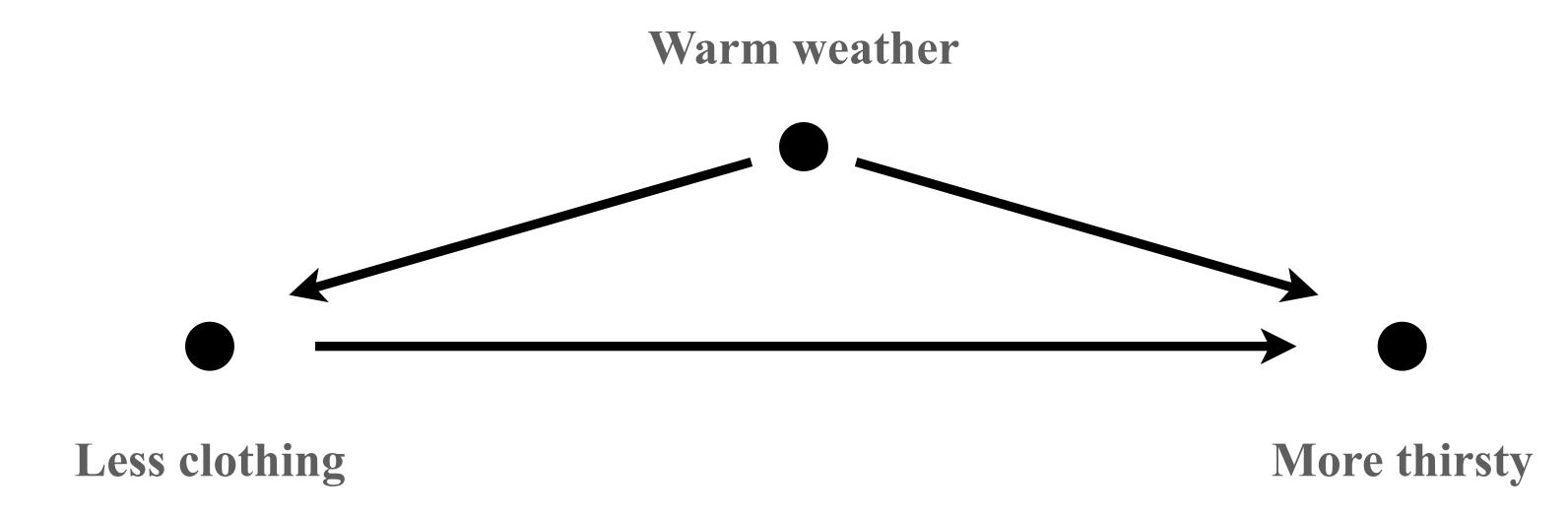
Less clothing

More thirsty

Warm weather

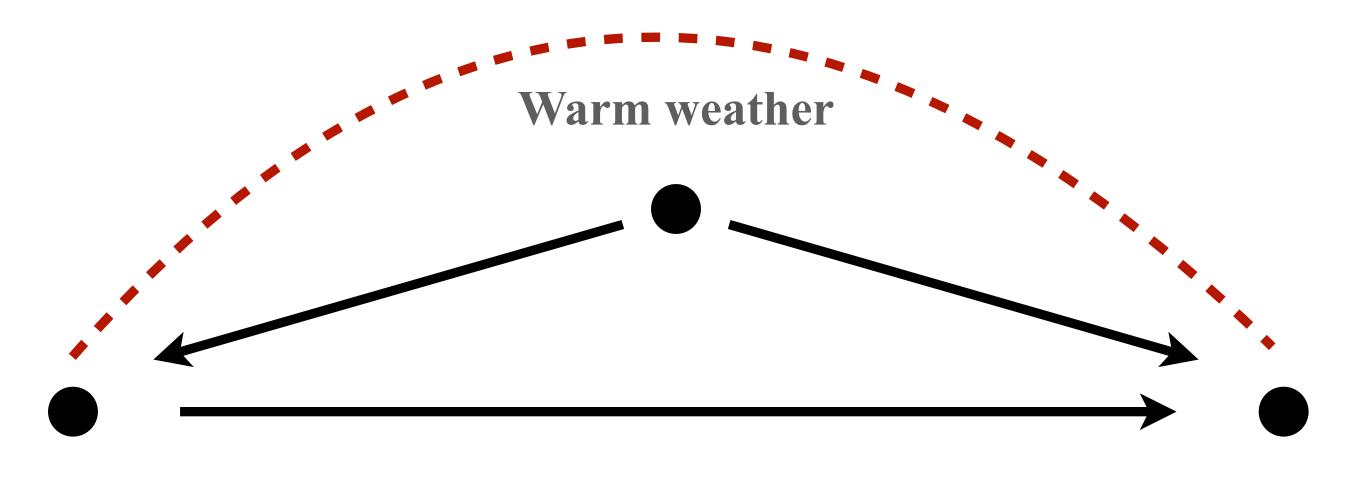


Less clothing More thirsty



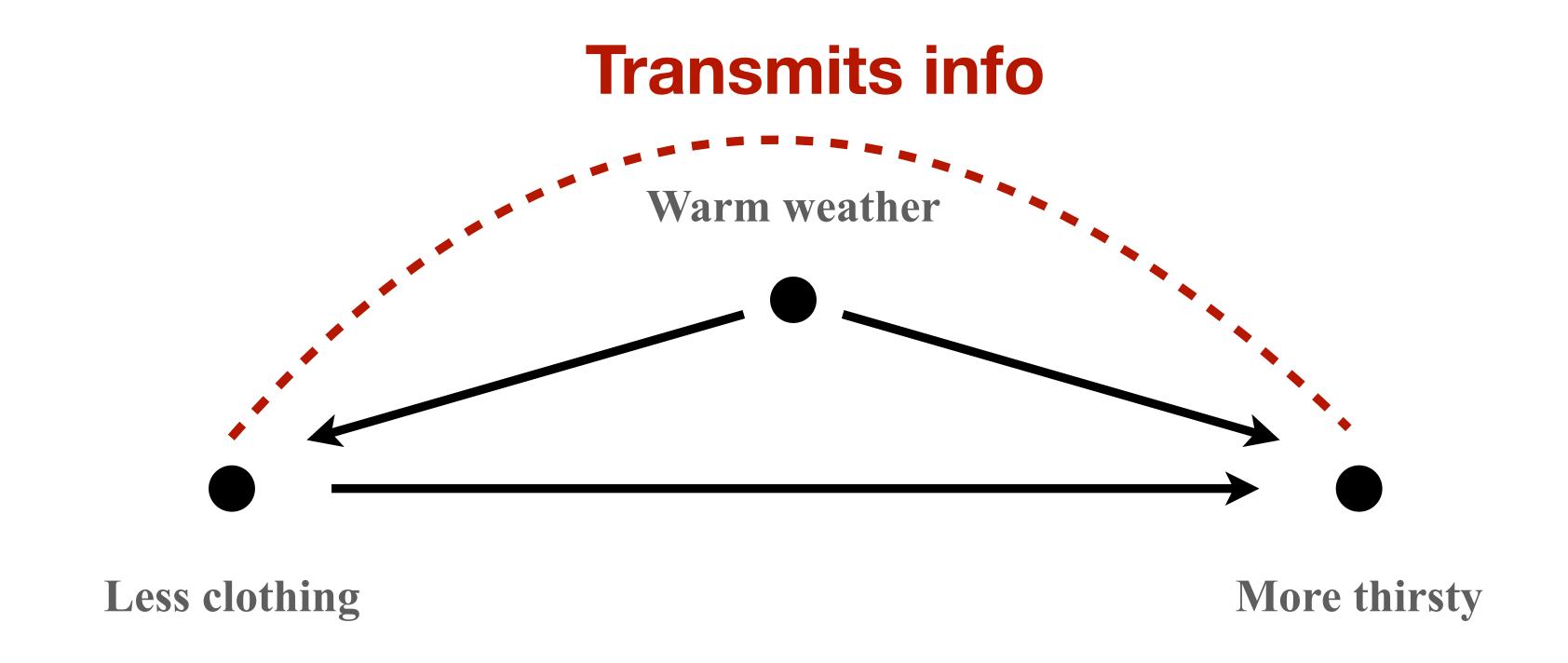
Note: leaving an arrow in is a weaker assumption than removing it.

assuming an effect might be anything (including 0) VS assuming that it is 0 exactly

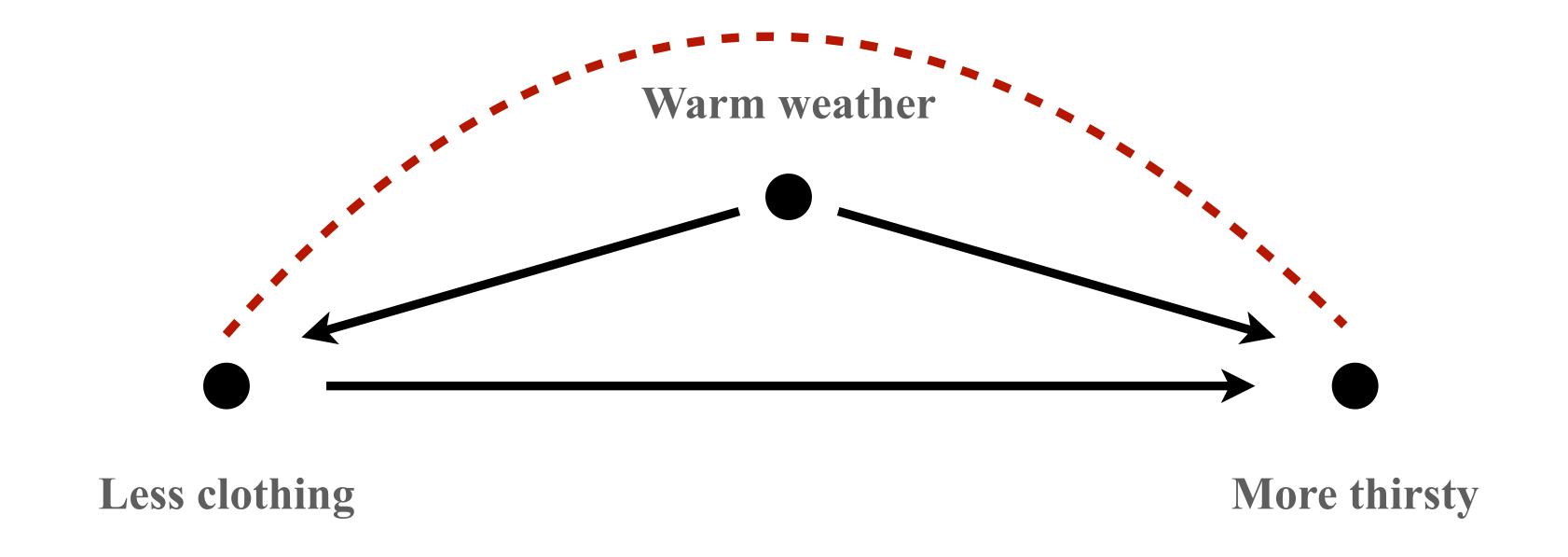


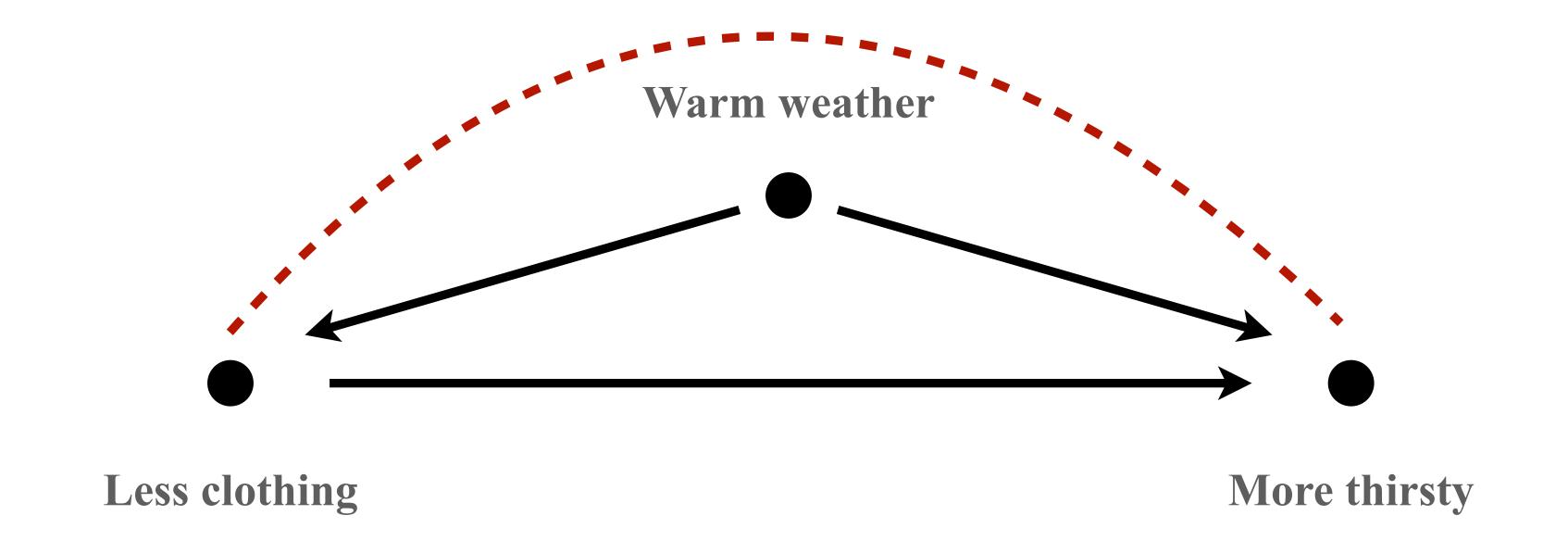
Less clothing

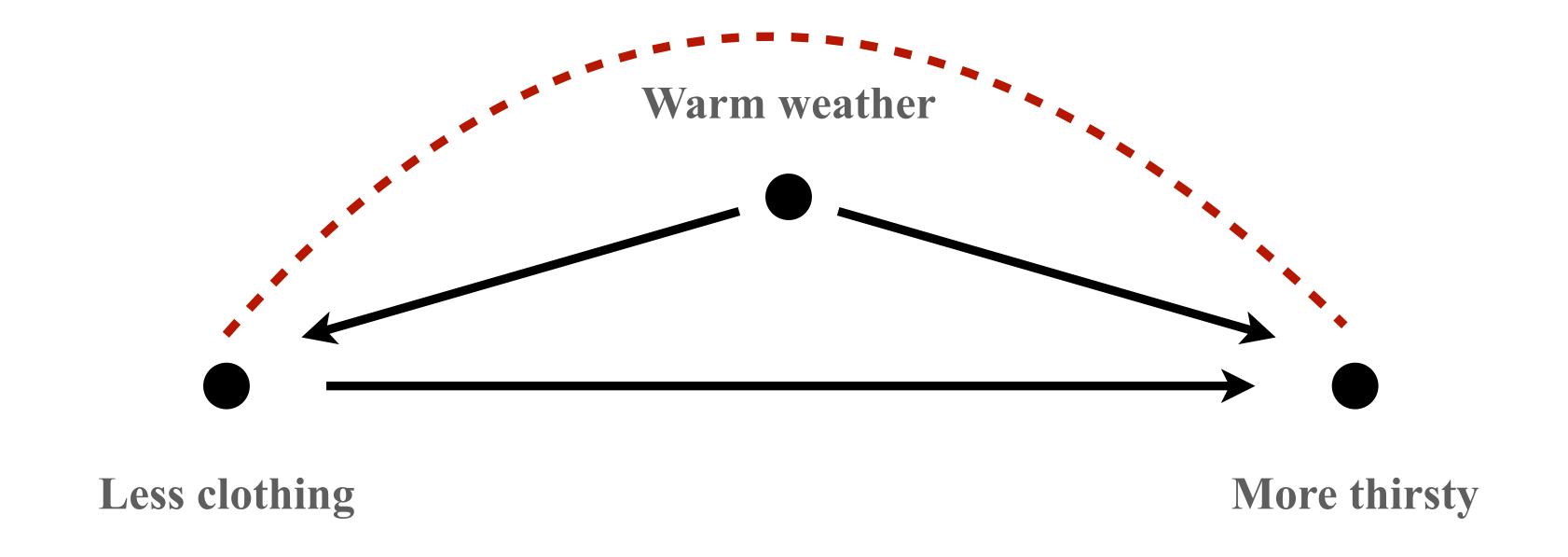
More thirsty



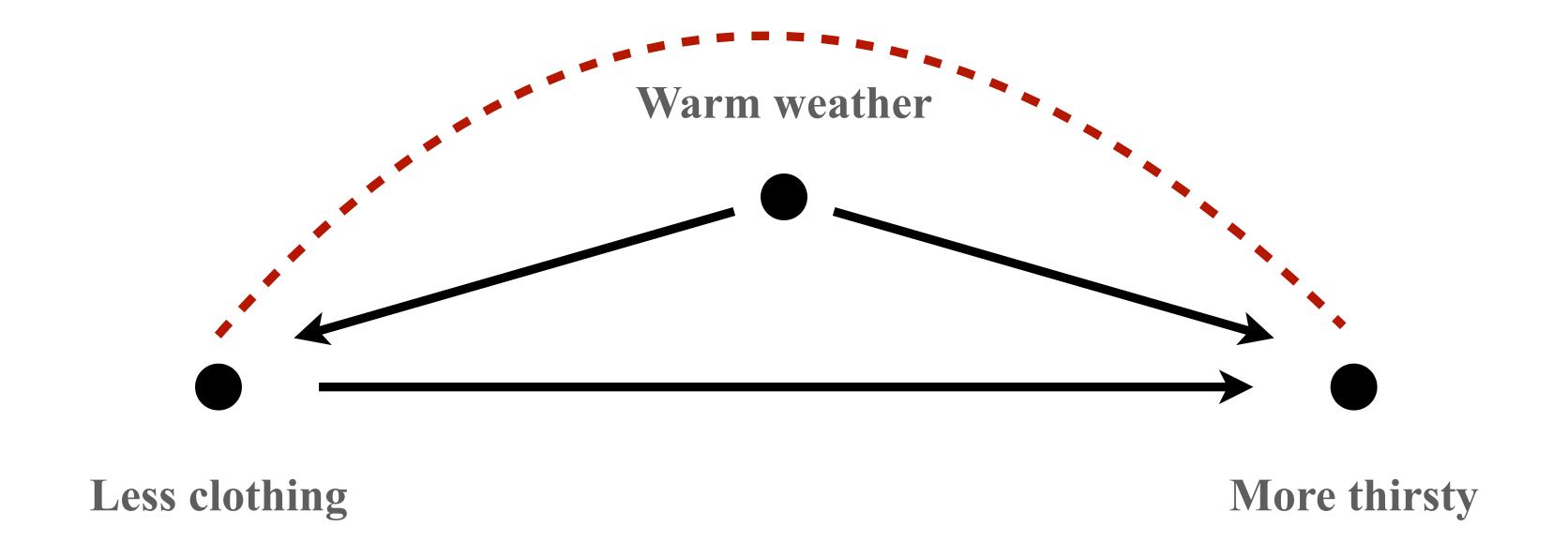
to confound: to mix or confuse



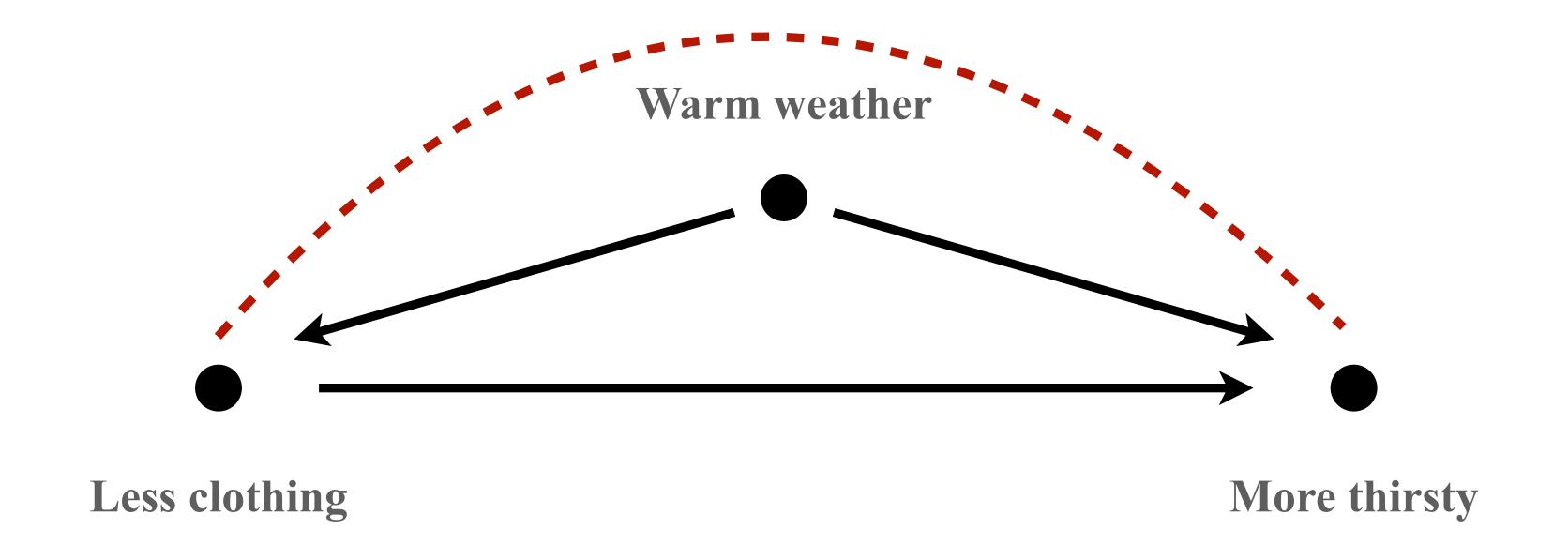




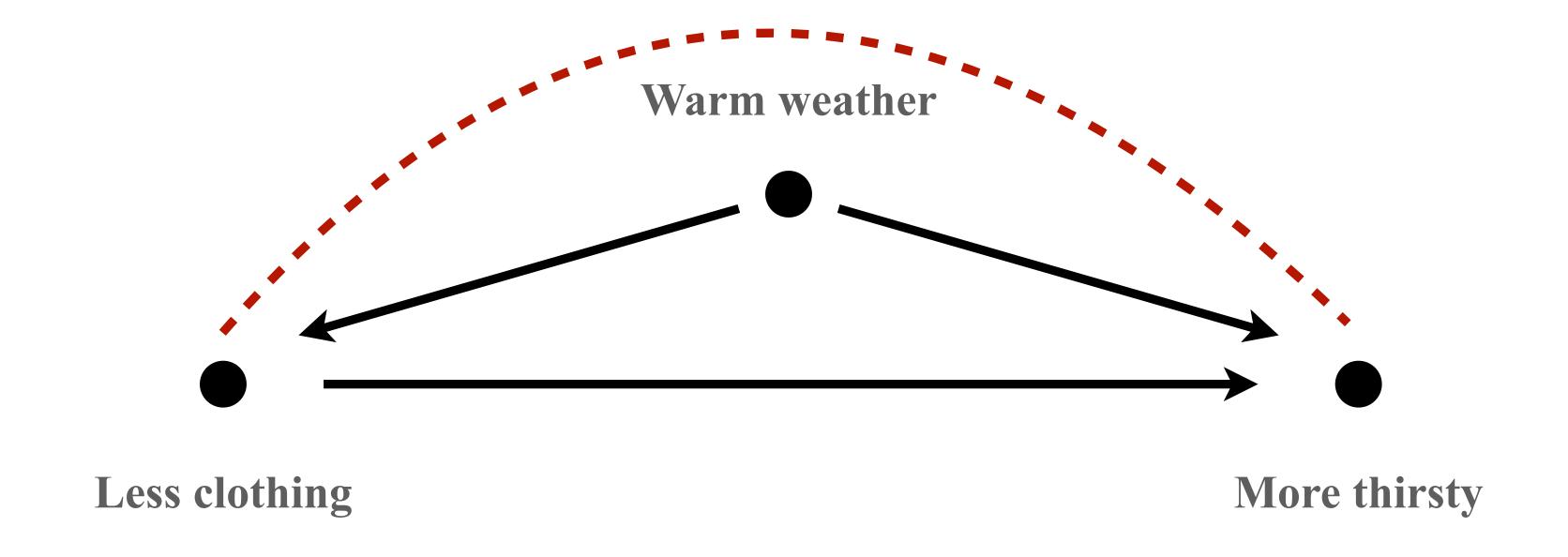
• open: no collider along the path



- open: no collider along the path
- back-door: an arrow goes into the exposure

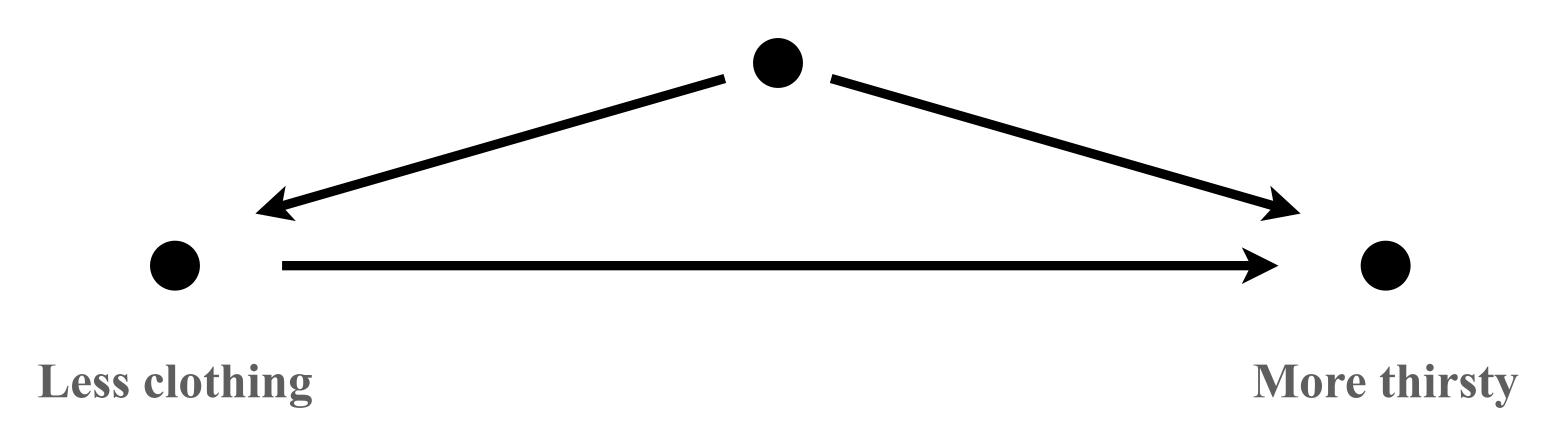


- open: no collider along the path
- back-door: an arrow goes into the exposure
- The problem: there is a mixing of the presumed causal relation along the direct path and the purely associational relation through the back-door path

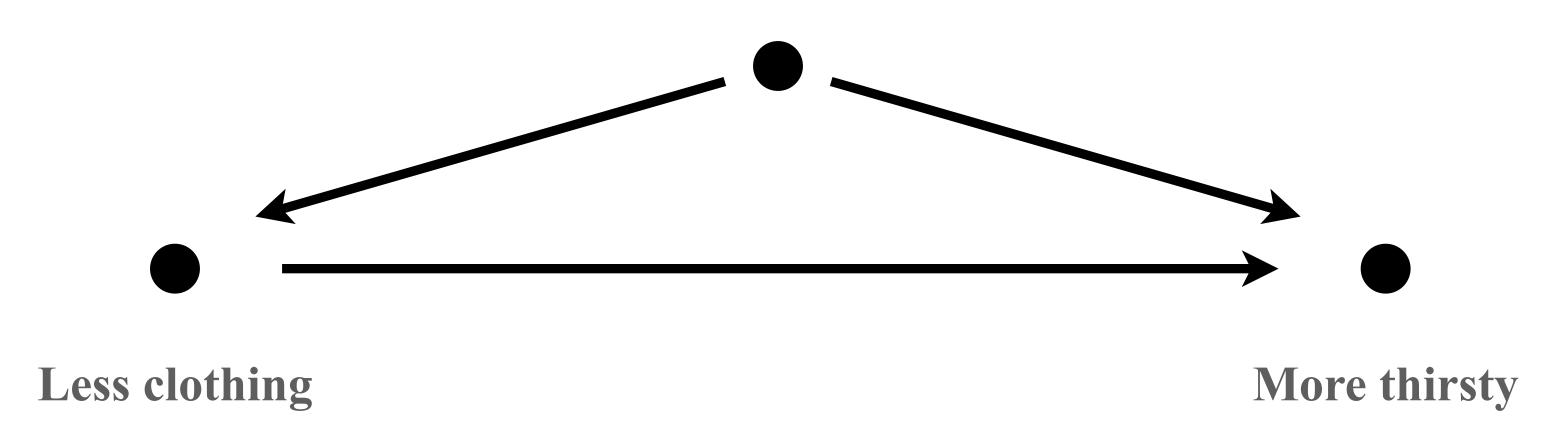


Please to tell me what I can do about this situation.

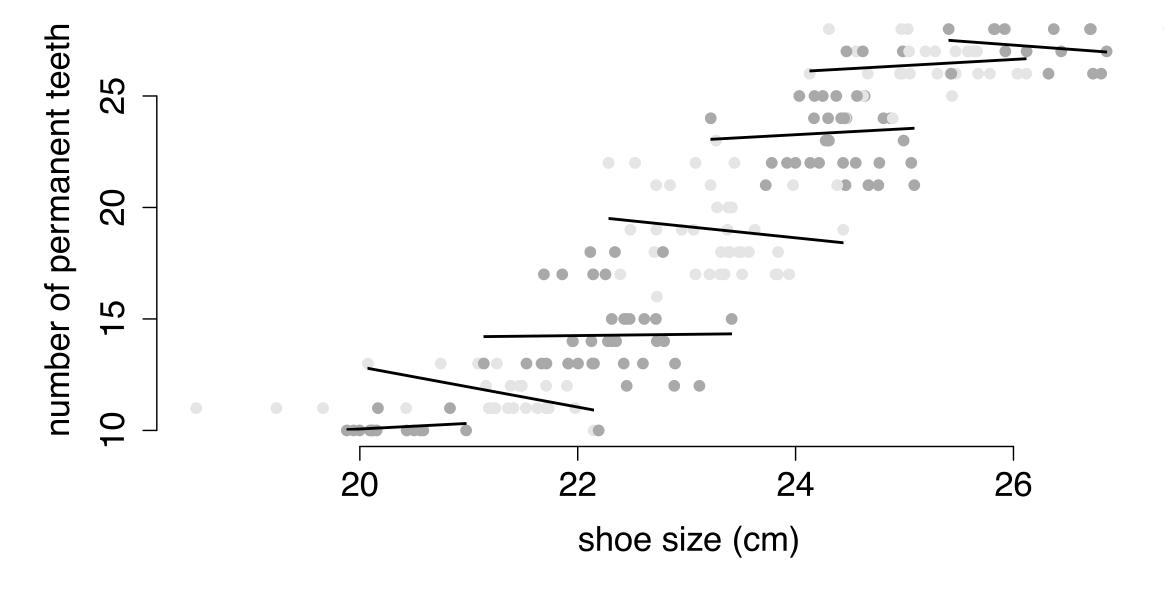
Temperature = x, y, z, ...



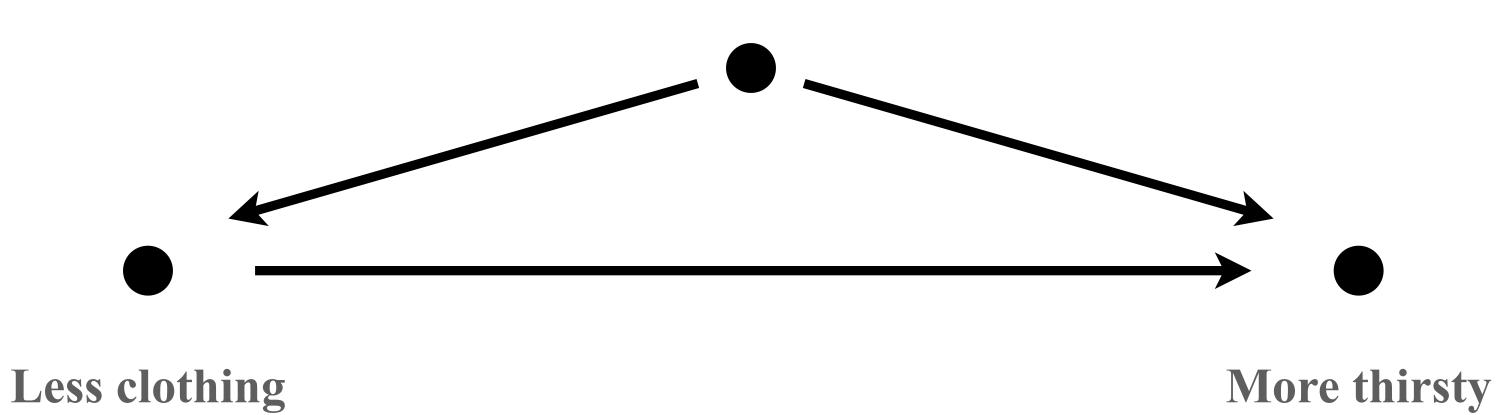




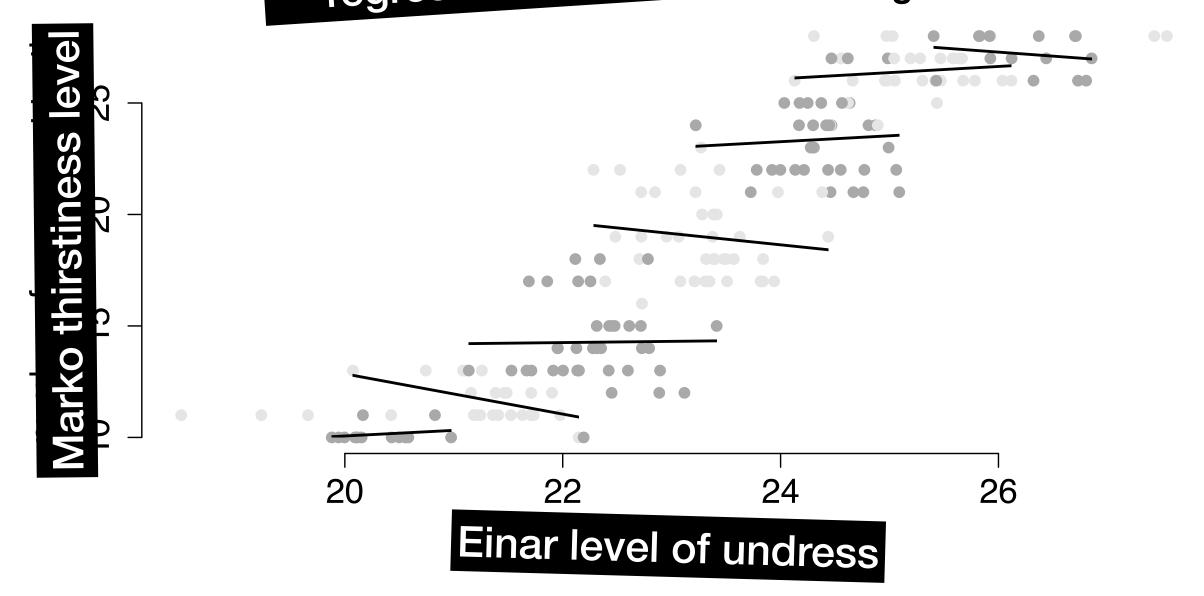
teeth vs shoe size, regressions conditional on age



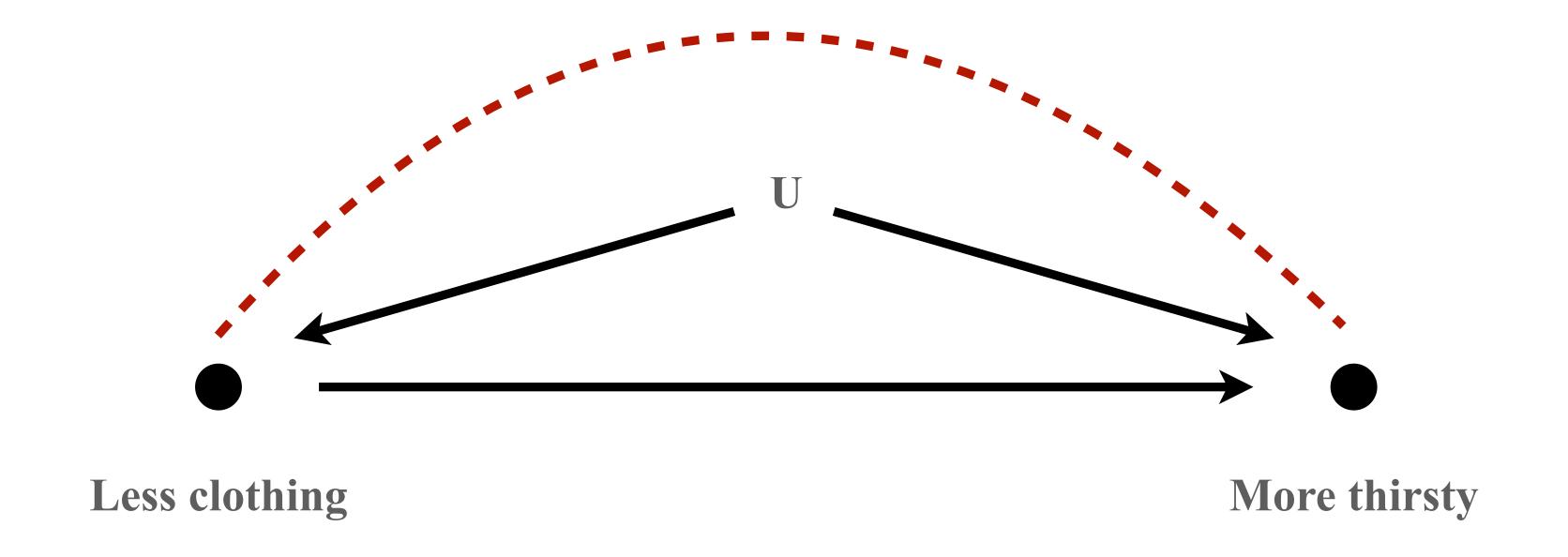
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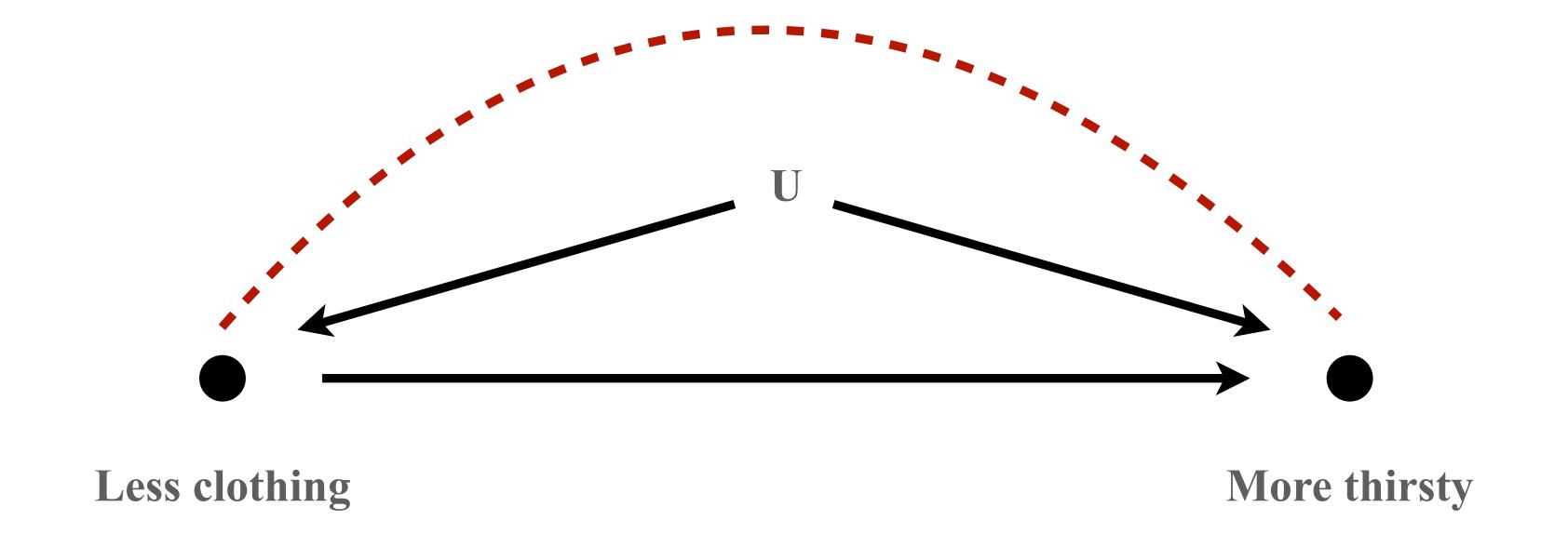
Thirst level as a function of state of undress, regressions conditional on temperature



Comparing 25 degree days with one another: comparing like with like

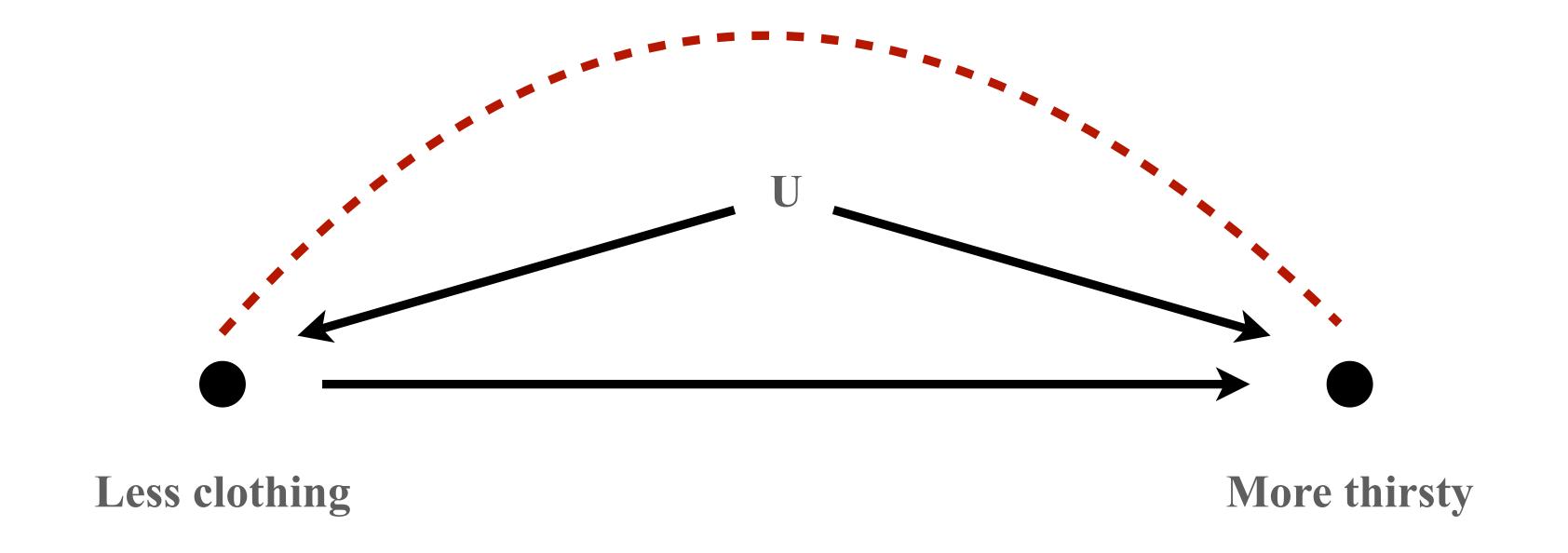


Temperature UNOBSERVED: what do we do???



Temperature UNOBSERVED: what do we do???

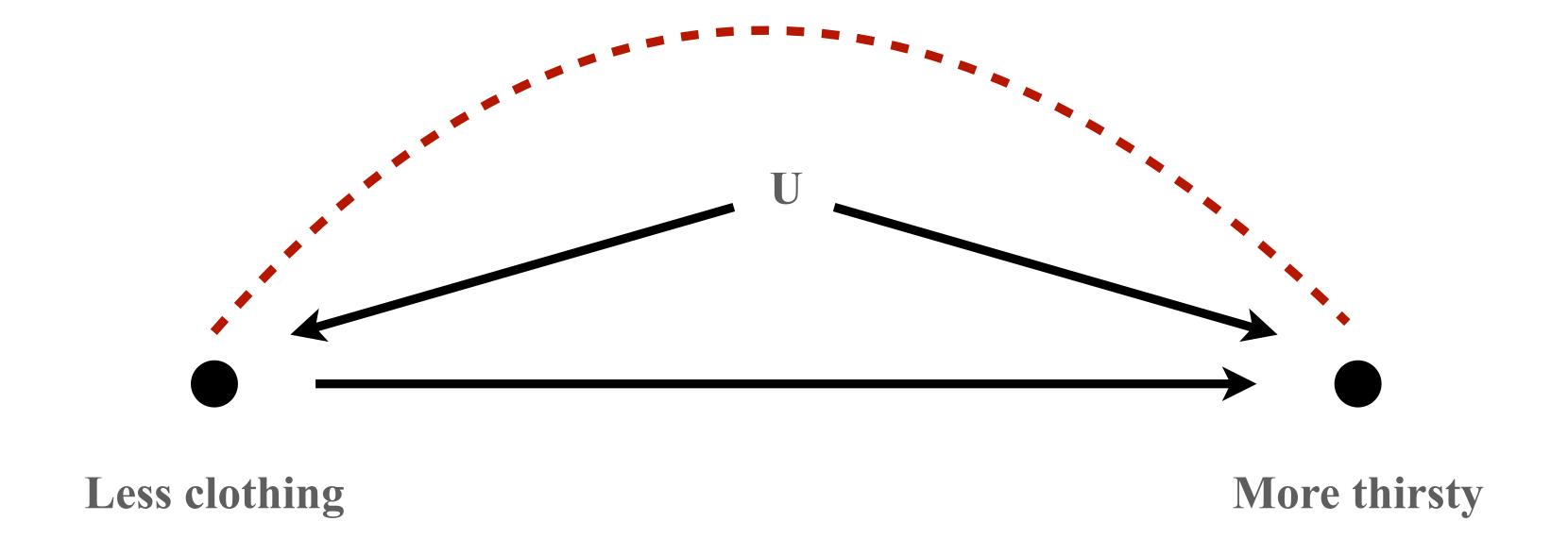
Not much to do: the effect is non-identifiable (get more data or do an actual experiment)



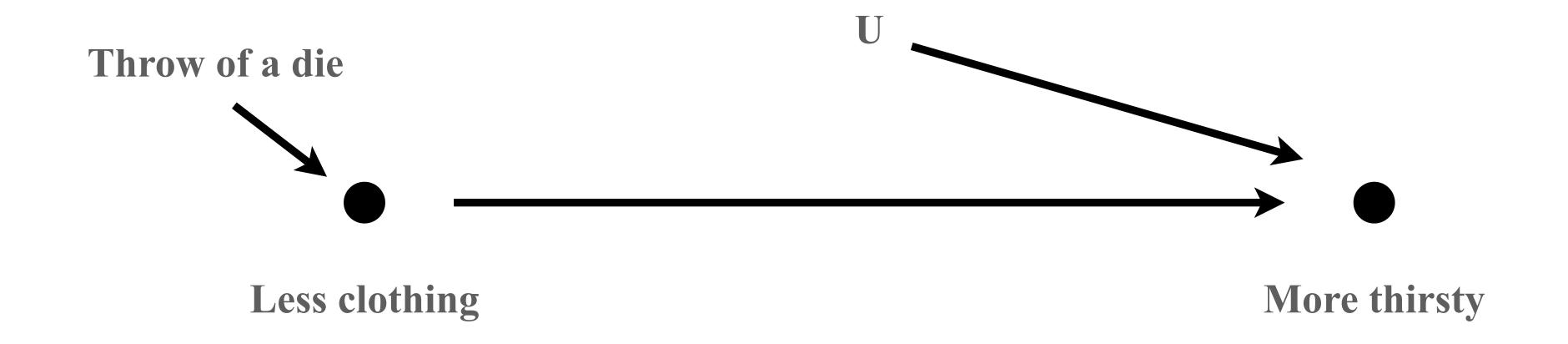
Temperature UNOBSERVED: what do we do???

Not much to do: the effect is non-identifiable (get more data or do an actual experiment)

An effect is identifiable if it is possible to close all back-door paths without opening new ones



Why randomization is so good

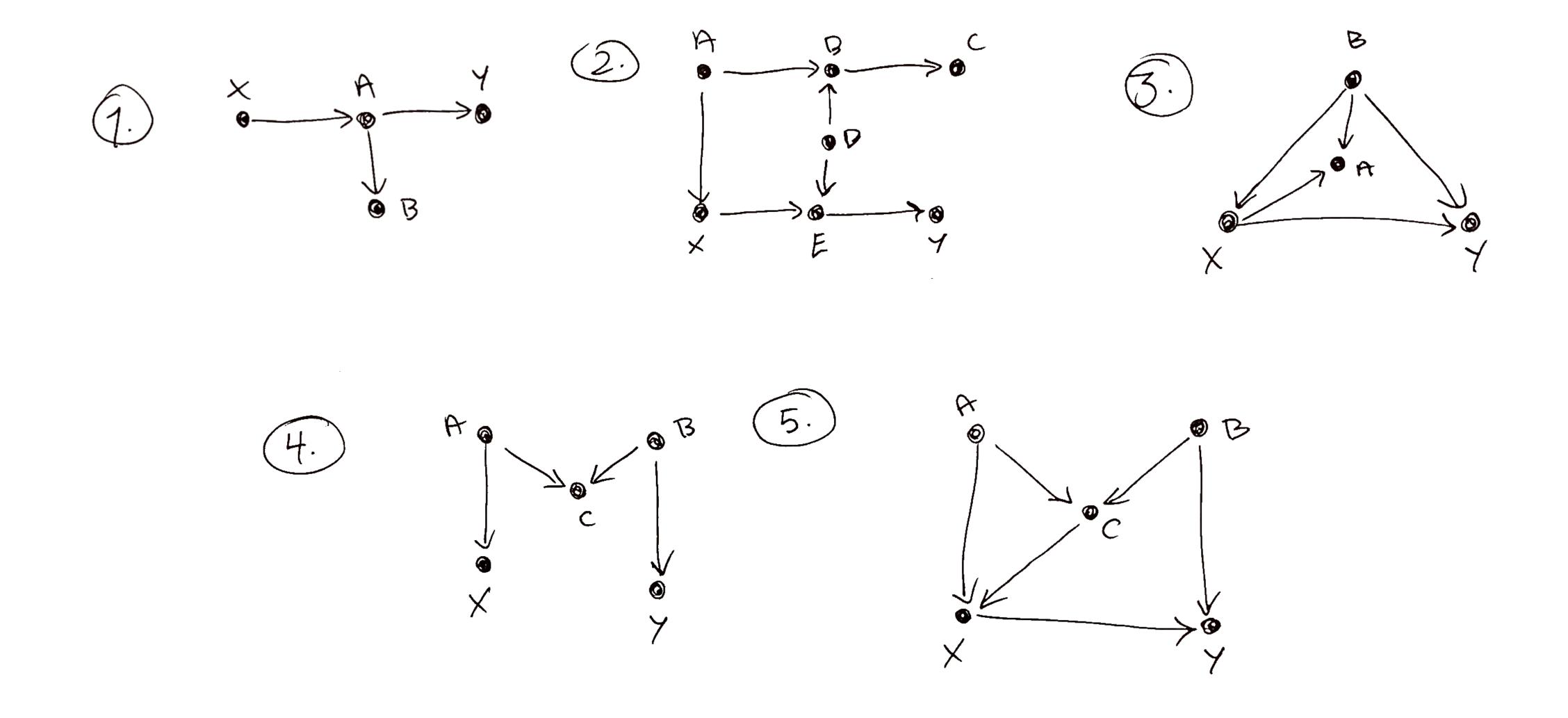


Why randomization is so good

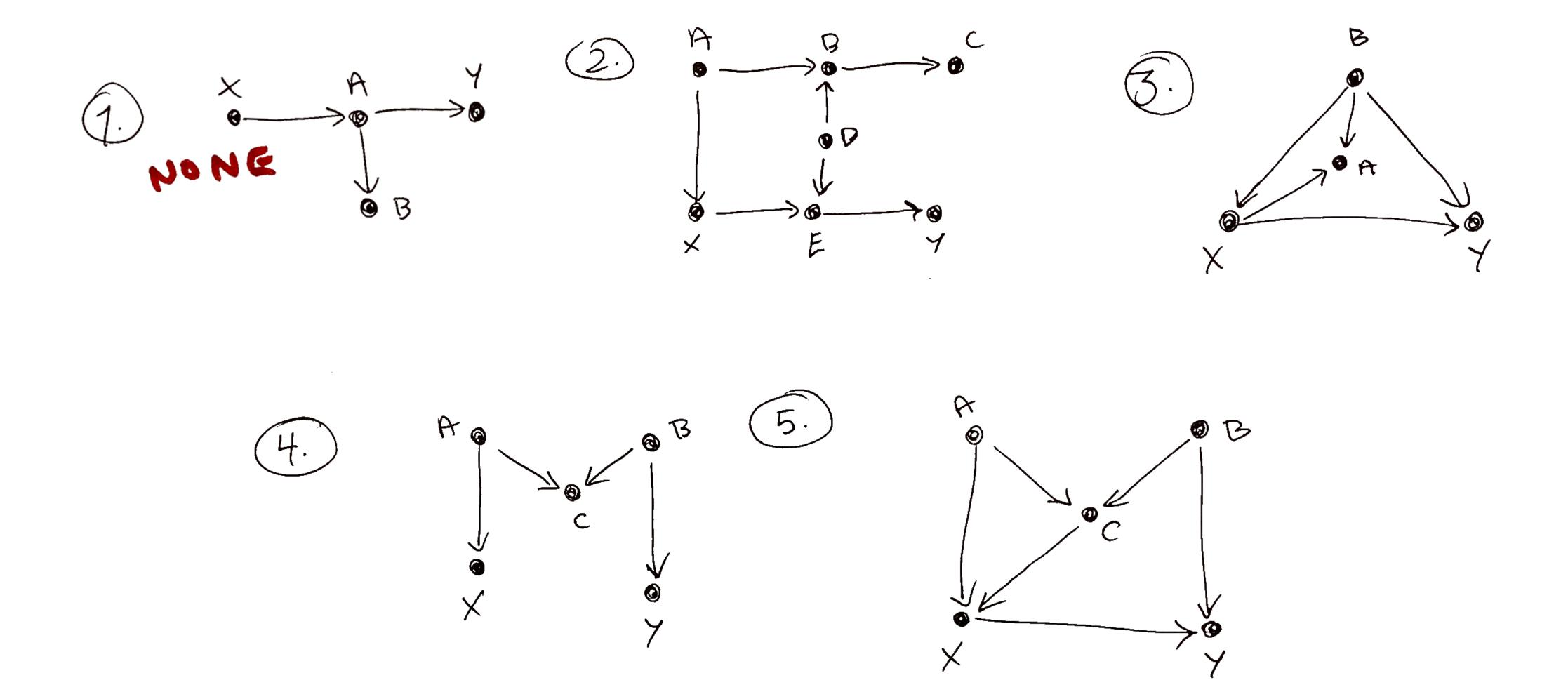
Point: For the price of making some causal assumptions, the rule about closing backdoor paths tells us exactly what to adjust for

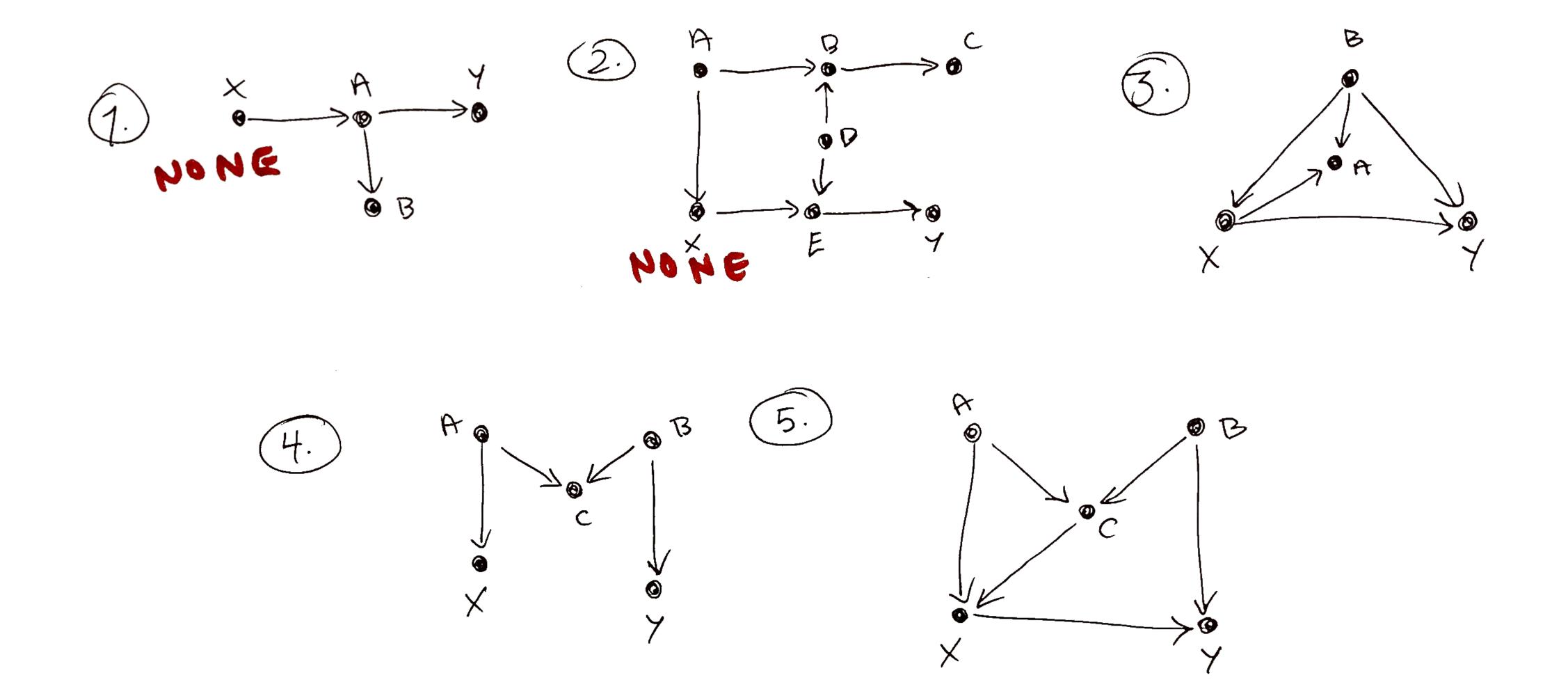
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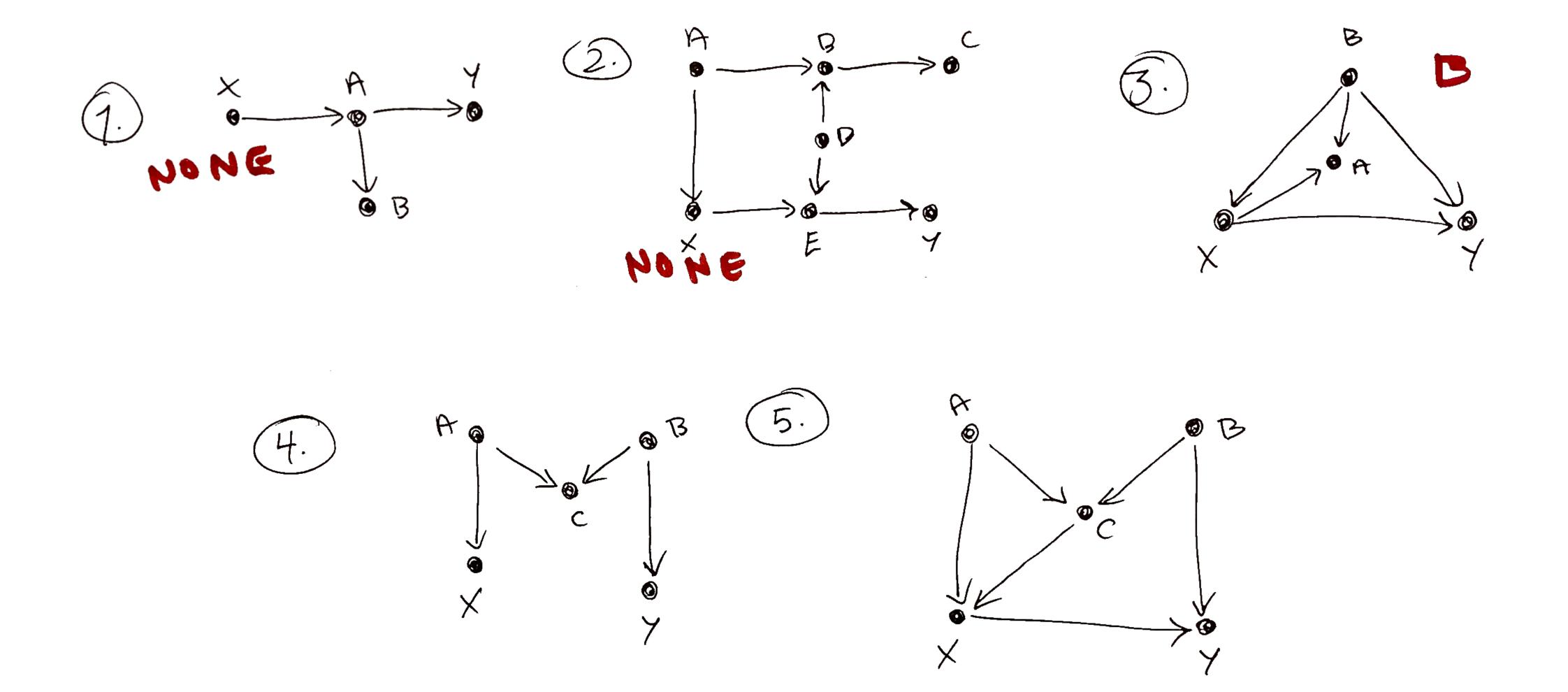
(Also you need to have made the right measurements)

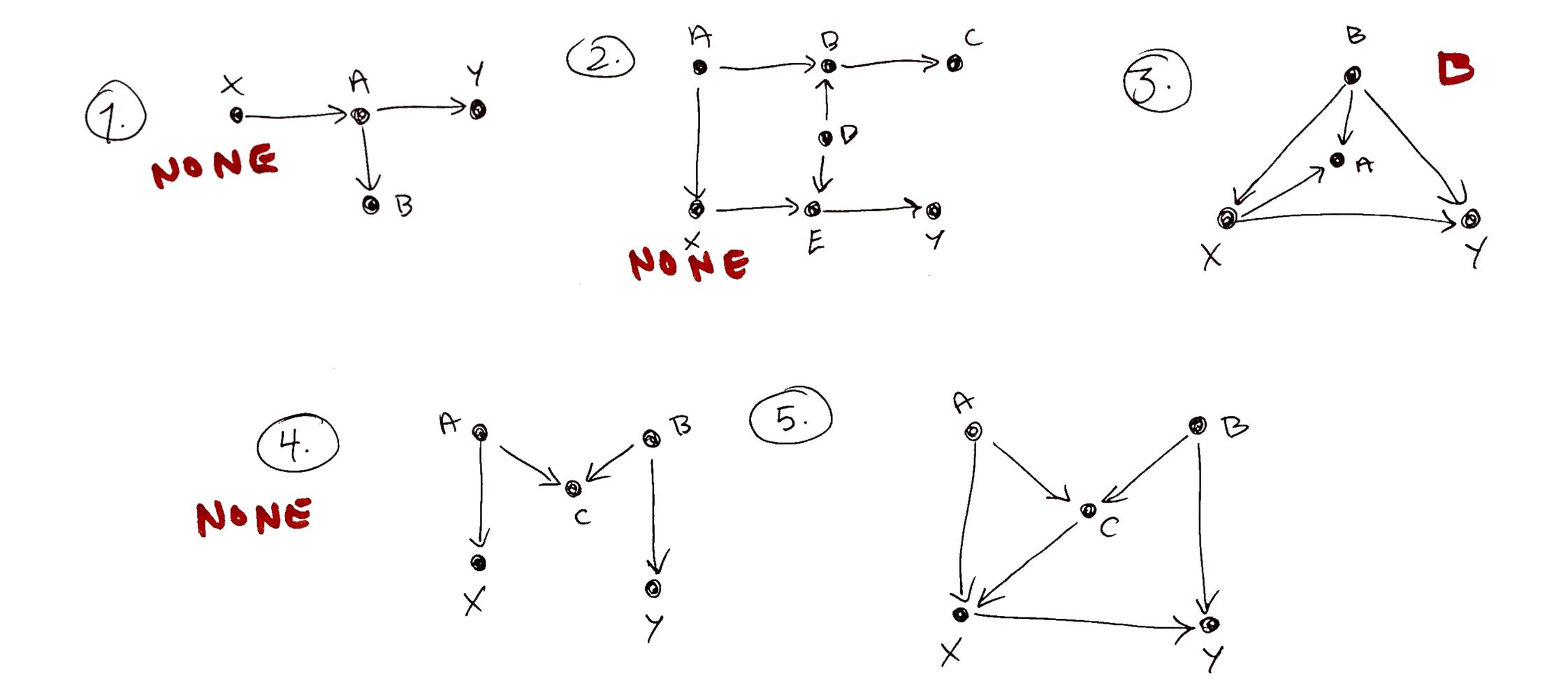


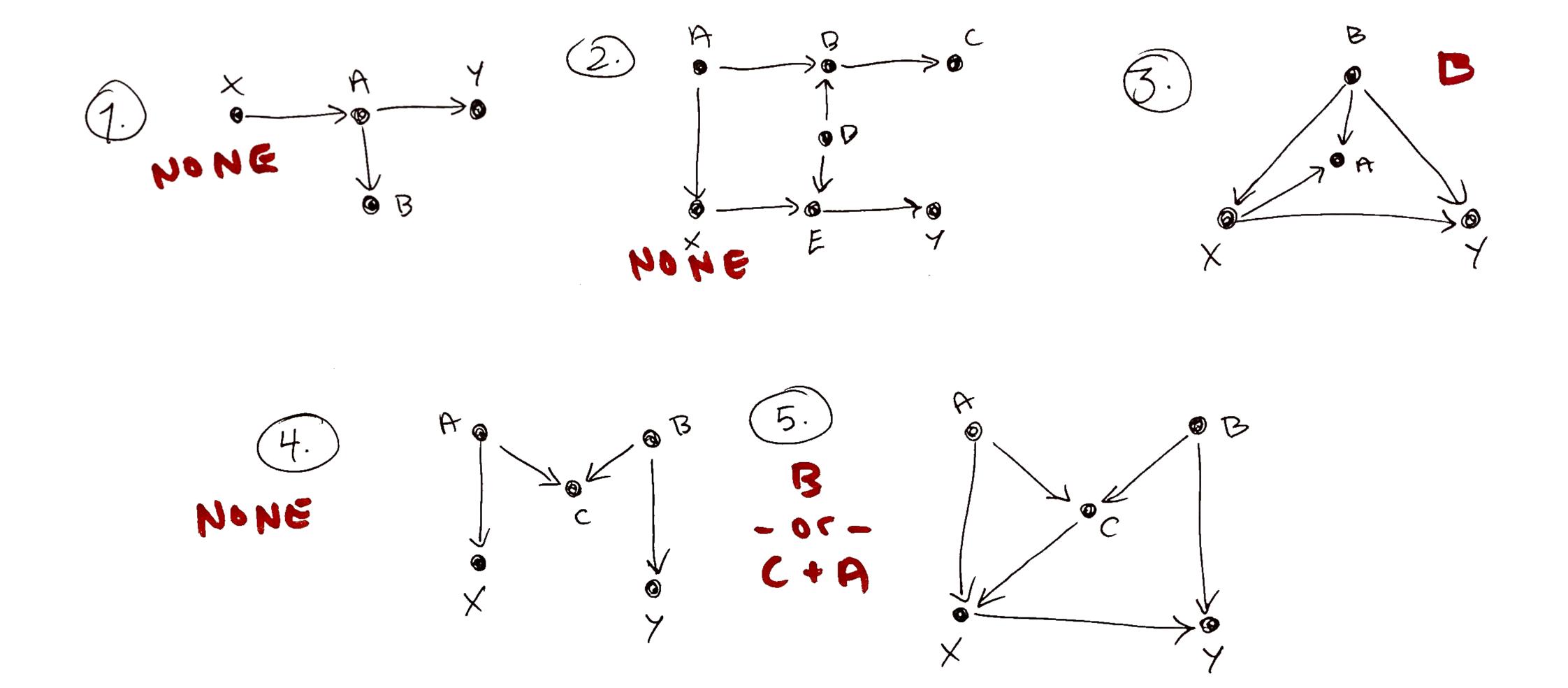
Puzzles: what should we adjust for to close the back-door paths between x and y?

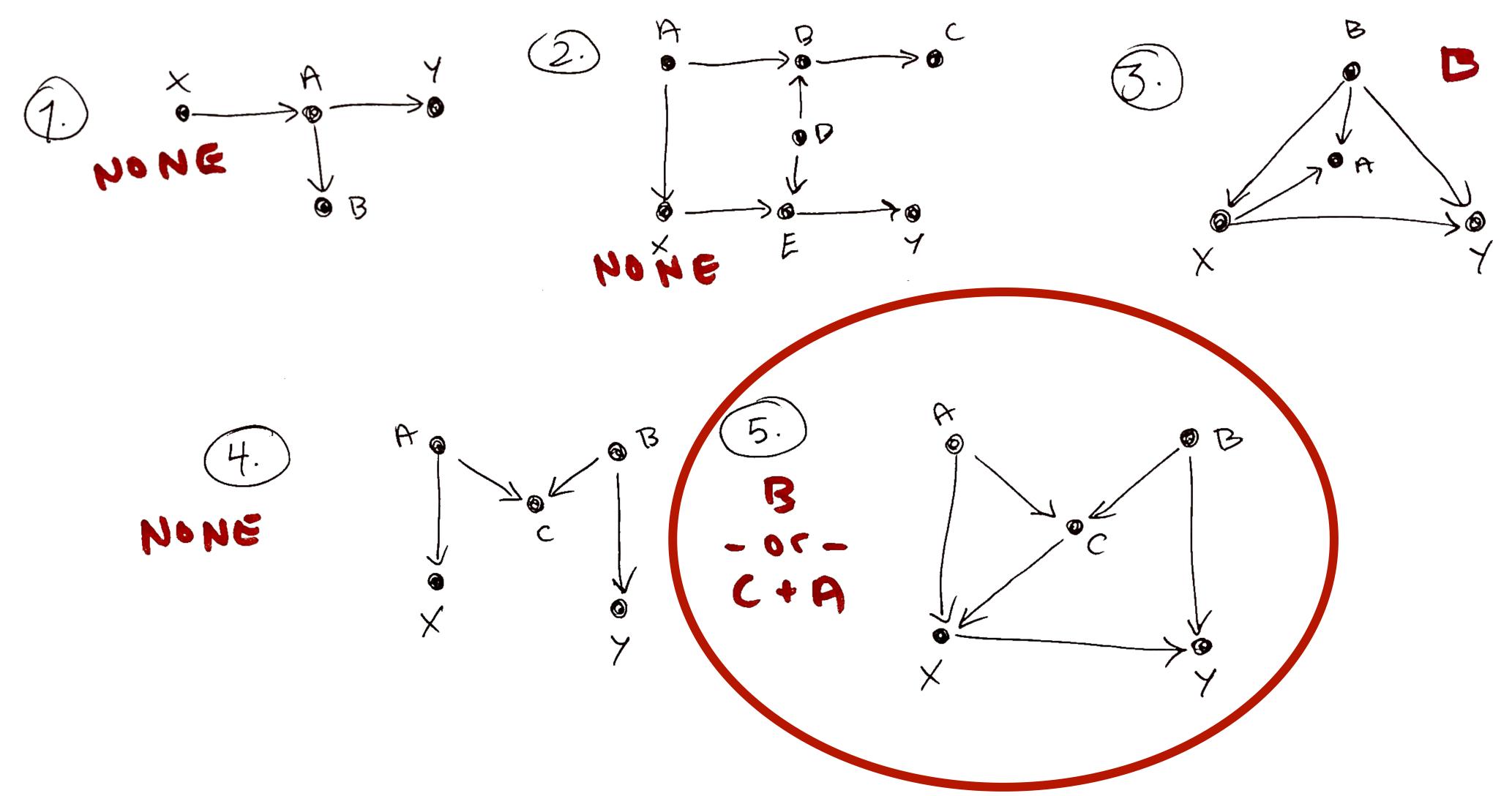








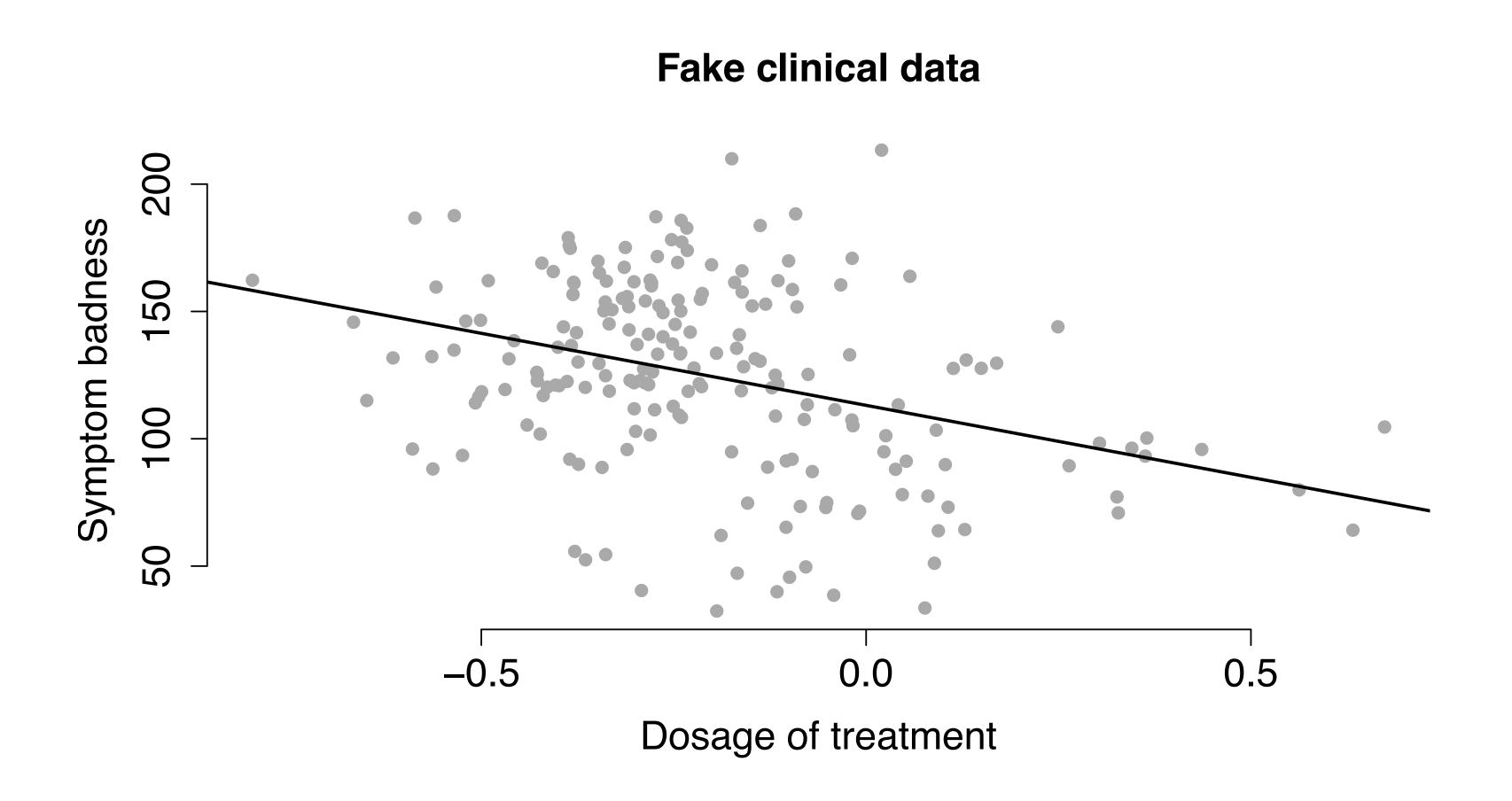




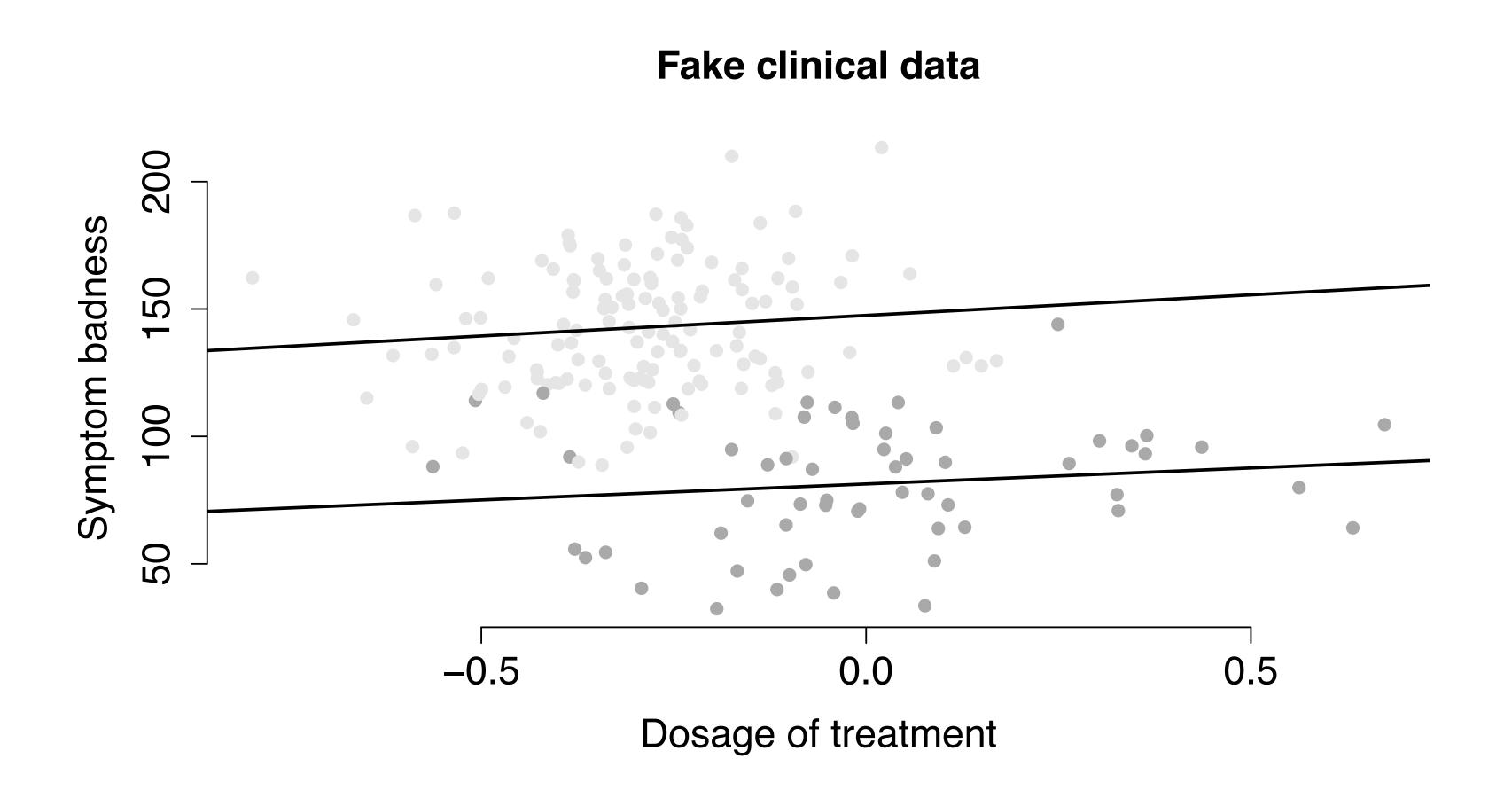
A+B+C also works but this set is not *minimal*, since we can remove something from it and still have a valid adjustment set

Awarning

The analysis and interpretation of your data depends on the assumed causal model

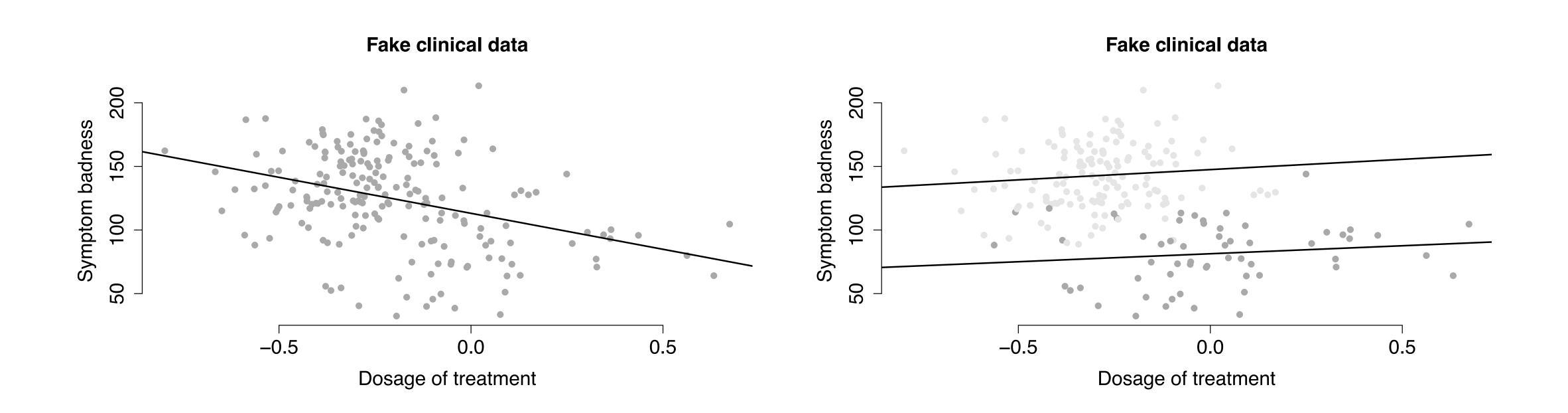


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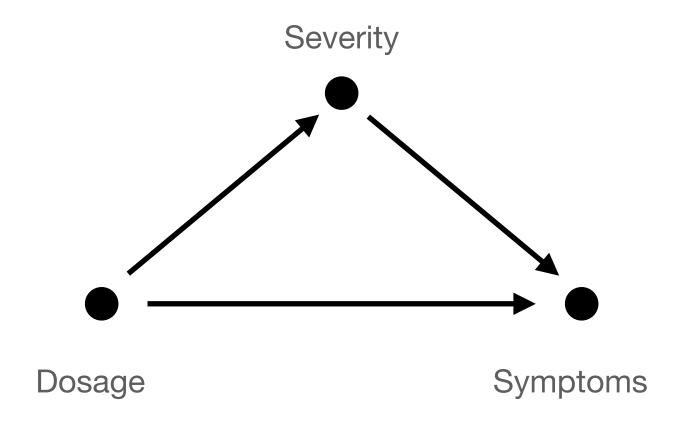


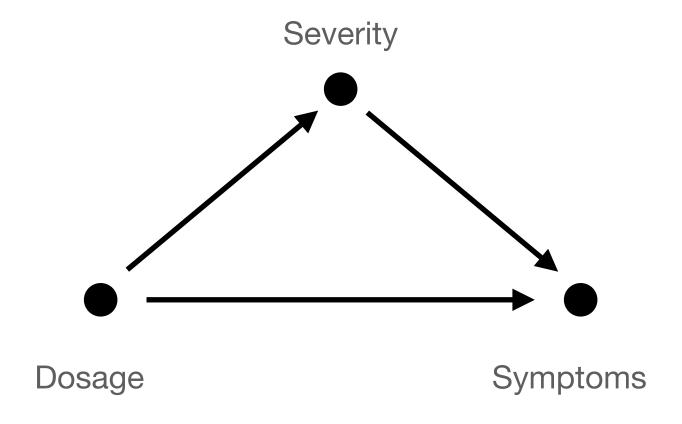
Effect reverses if we condition on disease severity

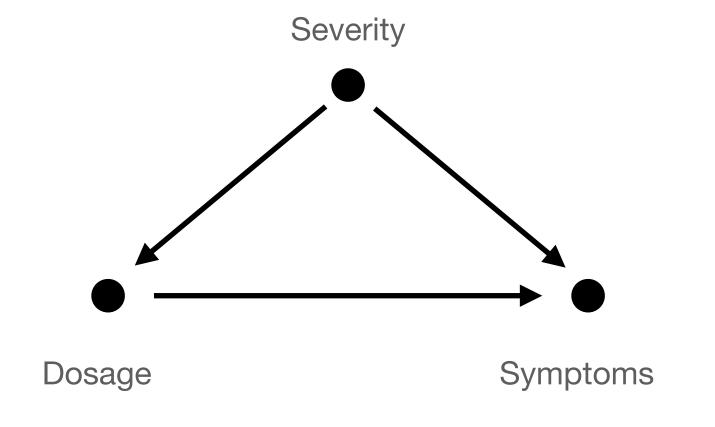
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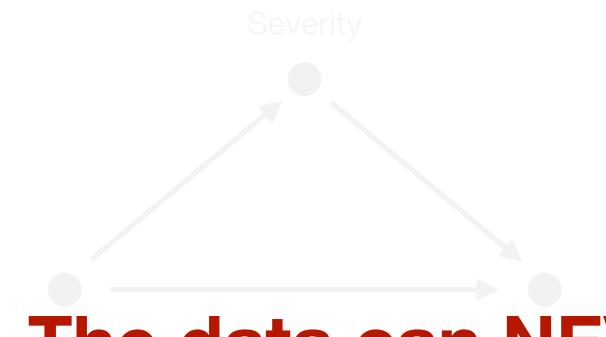
"Simpson's paradox": which is it, is the treatment good or bad???



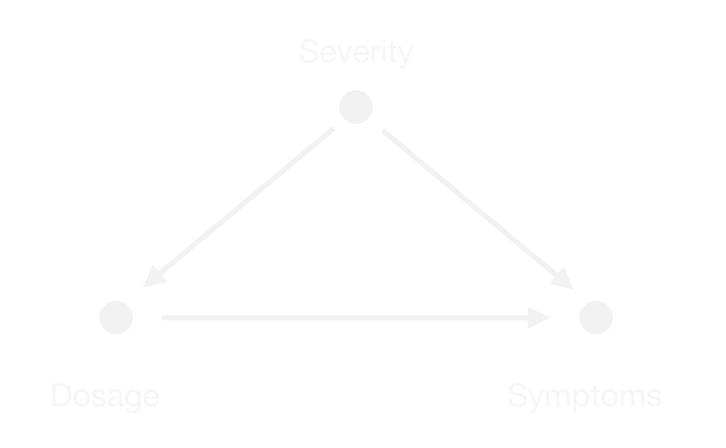




Drug acts on symptoms. Different doses given to severe and non-severe cases. Back-door path present: adjust.



The data can NEVER tell you which is right.



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The data can NEVER tell you which is right.

Be very skeptical of claims like "we found an effect but only for women" or similar.



Drug acts on symptoms. Different doses given to severe and non-severe cases. Back-door path present: adjust.

Severity

Drug acts by reducing both symptoms and disease severity. No back-door path: don't adjust.

The data can NEVER tell you which is right.

Be very skeptical of claims like "we found an effect but only for women" or similar.

Could be a collider, a mediator, or a confounding factor.

Dosage

Symptoms

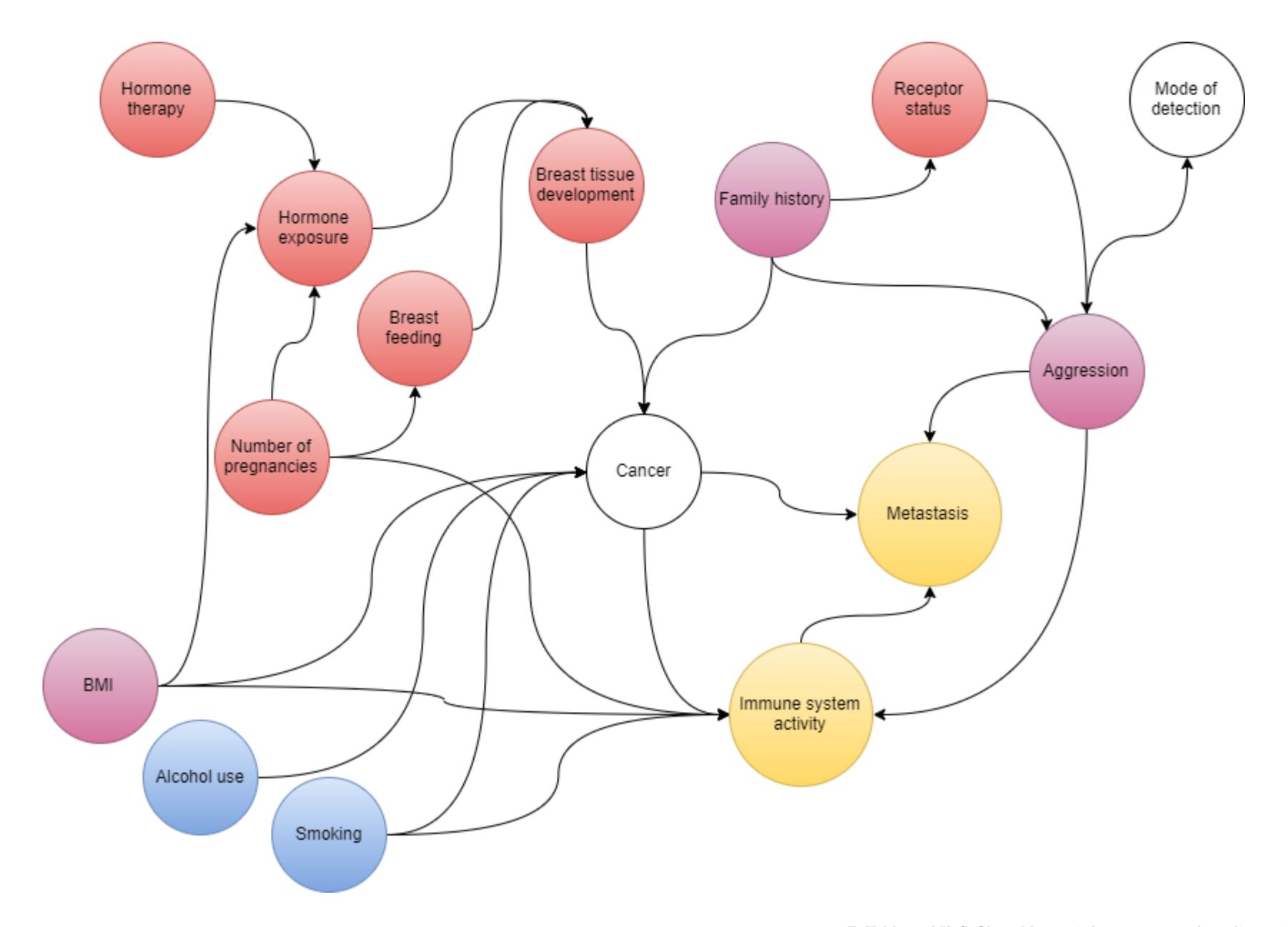
The data can NEVER tell you which is right.

Be very skeptical of claims like "we found an effect but only for women" or similar.

Could be a collider, a mediator, or a confounding factor.

Adjustment without some kind of domain knowledge (which you can encode as a DAG) practically impossible to interpret

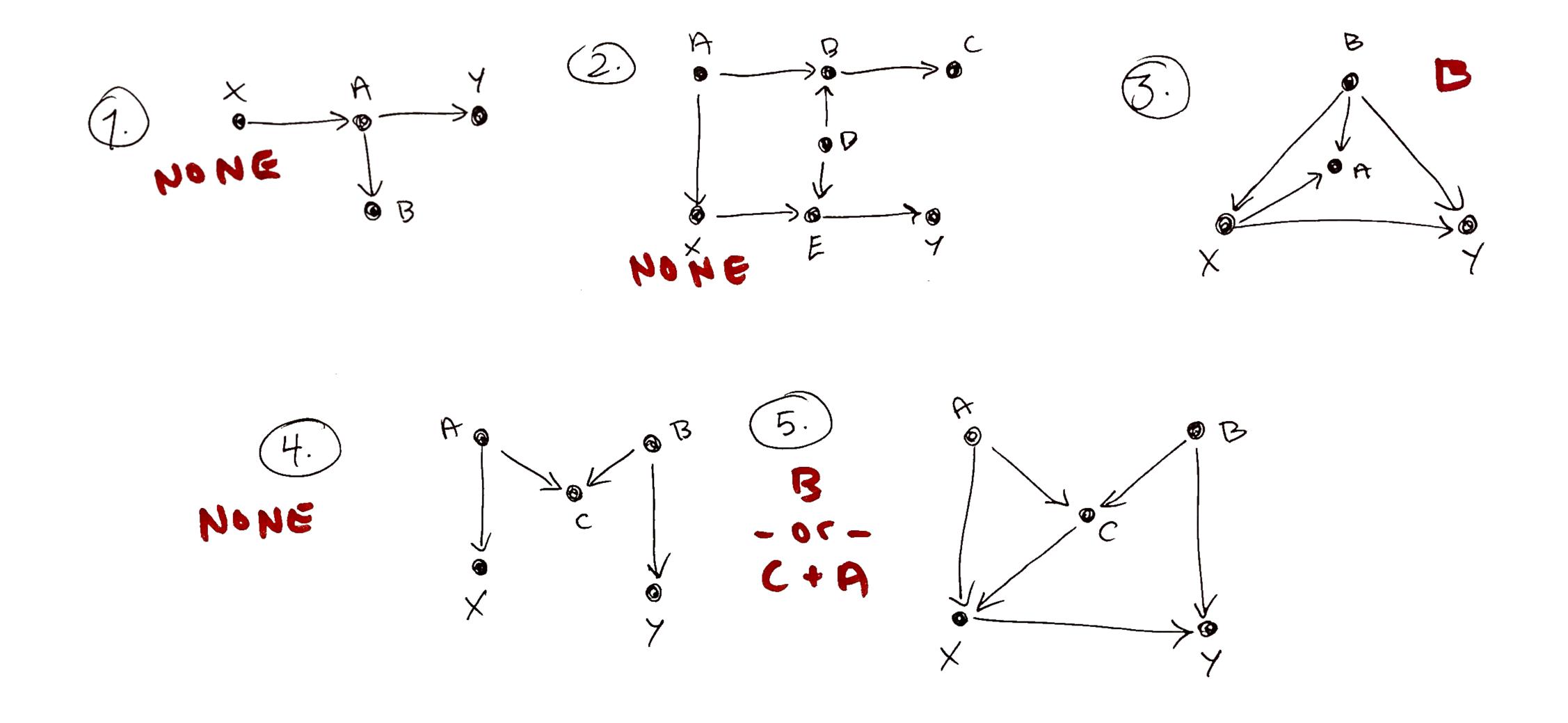
Real DAGs can be complicated



E. Holsbø and K. S. Olsen. Metastatic breast cancer and pre-diagnostic blood gene expression profiles—the norwegian women and cancer (nowac) post-genome cohort. Frontiers in Oncology, 10, 2020.

There are automated tools that tell you what to adjust for to close back-door paths

One such tool: DAGitty — http://www.dagitty.net/



Go to dagitty.net, start the online browser mode. Input these DAGs and see that you get the adjustment sets you expect


```
dag {
A [pos="-2.200,-1.520"]
B [pos="1.400,-1.460"]
D [outcome,pos="1.400,1.621"]
E [exposure,pos="-
2.200,1.597"]
Z [pos="-0.300,-0.082"]
A -> E
A -> Z [pos="-0.791,-1.045"]
B -> D
B -> Z [pos="0.680,-0.496"]
E -> D
}
```

Save your DAG by copying this text. Can be pasted into the same box when you want to continue

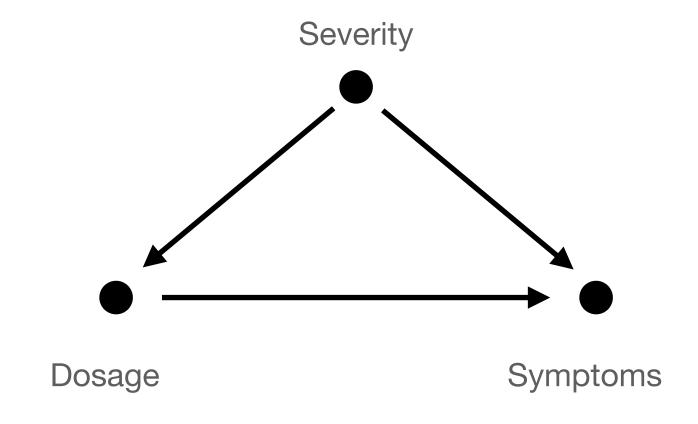
Once you have a diagram, computers will help you do a lot of the analysis. Frees up time to think about the science

Broadly speaking: adjustment for Z to close back door paths is simply to add it as a predictor in the regression model

Big assumption: the mathematical form of the regression model is correct

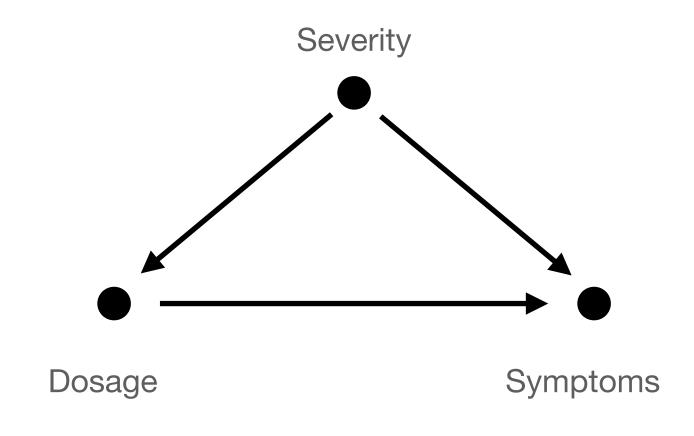
Basic linear model (symptom is continuous: blood pressure maybe)

Enough to include the adjustment variable as a predictor:



Basic linear model (symptom is continuous: blood pressure maybe)

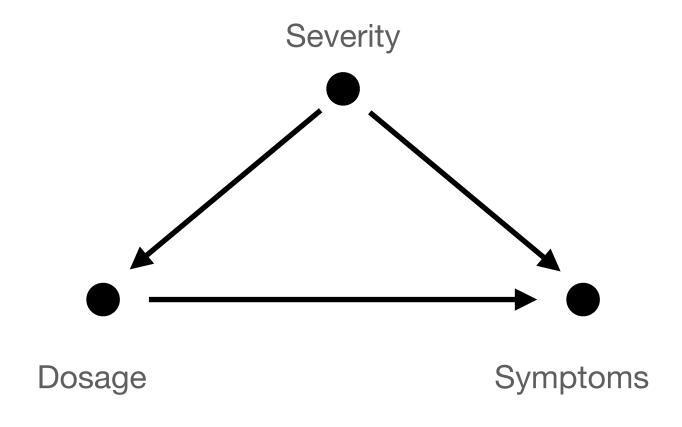
Enough to include the adjustment variable as a predictor:



symptom =
$$\beta_0 + \beta_1 dosage + \beta_2 severity$$

Basic linear model (symptom is continuous: blood pressure maybe)

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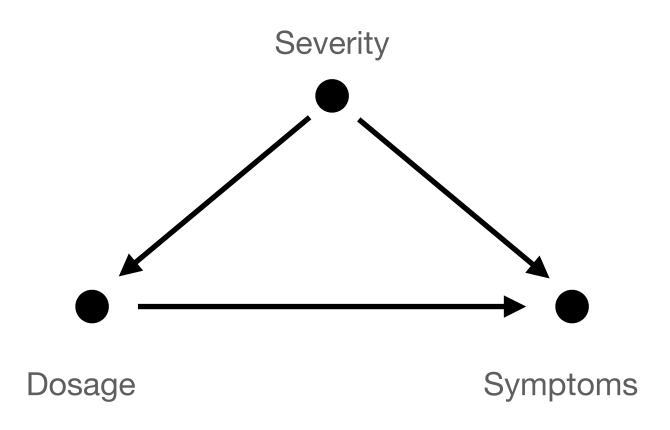


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 β_1 is an estimate of the causal effect of dosage on symptom

Basic linear model (symptom is continuous: blood pressure maybe)

Enough to include the adjustment variable as a predictor:



symptom =
$$\beta_0 + \beta_1 dosage + \beta_2 severity$$

 β_1 is an estimate of the causal effect of dosage on symptom

 β_2 does not have a straight-forward interpretation: Simply there to "deconfound"

Average treatment effect

The β_1 of linear regression is an estimate of the **average treatment** effect:

$$\mathbb{E}[Y_1-Y_0]$$

Average treatment effect

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$$\mathbb{E}[Y_1-Y_0]$$

Which reads "average difference between giving someone the treatment and not giving them the treatment"

Logistic regression and similar (eg. binary outcome)

$$\log \frac{p}{1-p} = \beta_0 + \beta_1 x + \beta_2 z \qquad p = \frac{1}{1 + e^{-(\beta_0 + \beta_1 x + \beta_2 z)}}$$

Adjustment is still to add Z as predictor, BUT interpretation more tricky.

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Adjustment is still to add Z as predictor, BUT interpretation more tricky.

We know that e^{β_1} is an estimate of odds ratio, but it is not the "average causal" odds ratio. We say that the odds ratio is *noncollapsible*.

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It's always possible to "stratify-and-average:"

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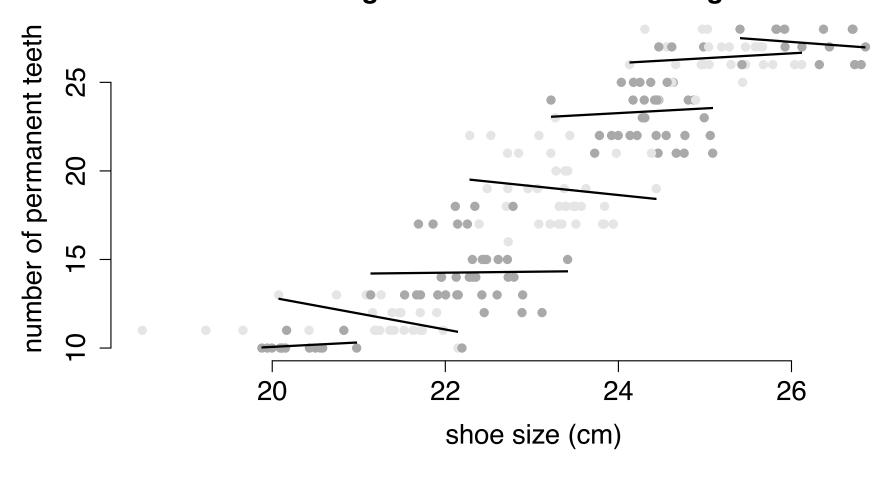
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teeth vs shoe size, regressions conditional on age



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Slightly annoying to do by hand, but there is software

> Eur J Epidemiol. 2016 Jun;31(6):563-74. doi: 10.1007/s10654-016-0157-3. Epub 2016 May 14.

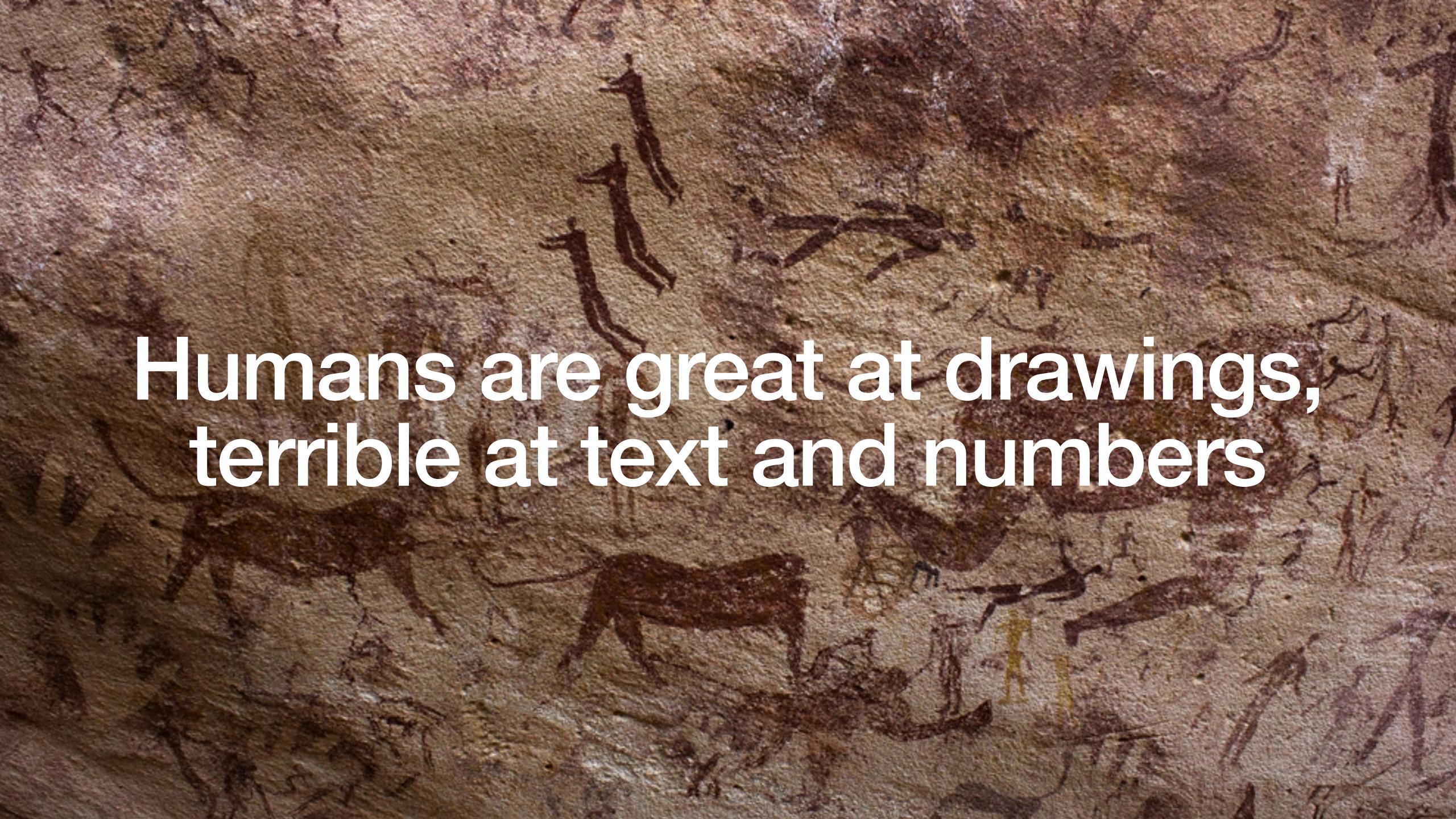
Regression standardization with the R package stdReg

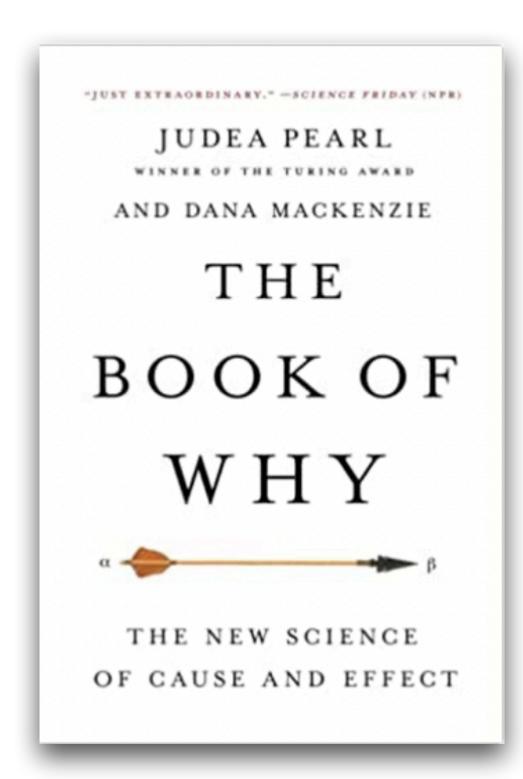
Arvid Sjölander 1

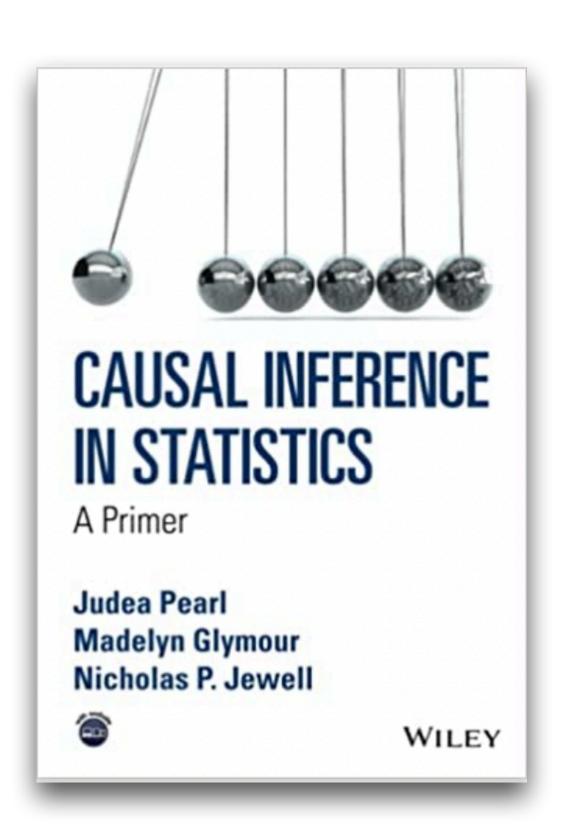


Last slides coming up!









Thank you.