The incredible utility of causal models UMC Methods Meeting

Oisín Ryan

Department of Methodology and Statistics Utrecht University

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Should we prescribe the drug? app.wooclap.com/CBRFDD

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Statistical phenomena where a relationship which is present when aggregating over the population may be reversed or absent when looking at sub-populations

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Berksons Paradox

Two phenomena which are statistically *independent* in the general population are statistically *dependent* in a sub-population

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The relationship between a categorical exposure and a continuous outcome is reversed when we condition on a third variable

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You're asking a question that statistical inference alone is not equipped to answer

Estimand

Estimator

Estimate

Estimator

Estimate



Estimand

Estimand



Estimator

Estimate

Prepare Chocolate Cake Batter

Preheat oven to 350 degrees, and prepare Yo's Ultimate Chocolate Cake batter. Prepare your pans with parchment. Pour 2 ¼ lbs into each 7" round pan, 1 ½ lbs into your 6" round pan, and divide the remaining batter evenly between your 5" round pans.

Bake Cakes

Bake your 7⁴ round cakes for 50 minutes, your 6⁴ round cake for 40 minutes, and your 5⁴ round cakes for 30 minutes, or until a toothpick comes out clean. Set aside to cool completely in their pars on a wire rack.

O Prepare Fillings & Simple Syrup

Prepare your dark chocolate ganache, italian meringue buttercream, and simple syrup. Set aside until you're ready to decorate.

6 Level Cakes

Remove your cooled cakes from their pans and level them with a ruler and serrated knife.

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Credit to Peter Tennant @PWGTennant

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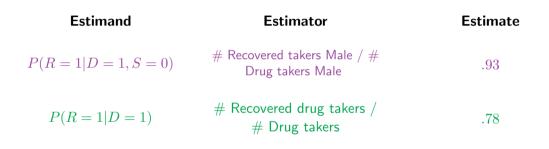
Conditional Probabilities:

P(R = r | D = d, S = s)

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Marginal Probabilities:

P(R=r|D=d)



Two different sets of estimands yield two different sets of estimates

• No paradox there!

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We are not interested in either of these estimands for their own sake

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We are interested in a **causal effect**

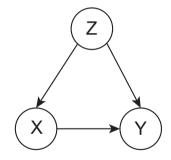
- Does taking the drug cause recovery?
- Causal Estimand
- But we have no way of expressing this in the language of statistical inference

Causal Inference

A causal graph is a diagram representing (our beliefs about) which variables share causal relations with each other

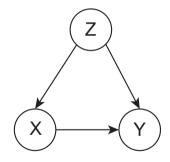
A causal graph is a diagram representing (our beliefs about) which variables share causal relations with each other

- The arrow X \rightarrow Y represents our belief that X is a direct cause of Y
- We omit an arrow if expert knowledge tells us that one variable does not directly cause another. The *absence* of an arrow is a strong statement



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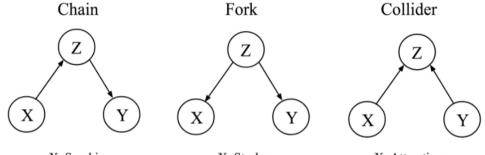


Directed Acyclic Graph (DAG) or Bayesian Network

This machinery is useful for three important and closely related reasons:

- 1 Causal models map causal dependencies onto statistical dependencies
 - *Regardless* of distributions and functional forms

Graphs and statistical dependencies



- X: Smoking Z: Tar Y: Cancer
- X ⊥ Y X ⊥ Y | Z

- X: Storks Z: Environment Y: Babies
- X ⊭ Y X ⊥ Y | Z

- X: Attractiveness Z: Being Single Y: Intelligence

This machinery is useful for three important and closely related reasons:

- 1 Causal models map causal dependencies onto statistical dependencies
 - *Regardless* of distributions and functional forms
- 2 Causal models allow us to define **causal effects** in the language of interventions and probabilities

The **do-operator** do(X = x) represents a "surgical intervention", forcing X to x.

We can use this to define our causal estimand

Causal Effect of Drug-Taking on Recovery:

$$CE = P[R \mid do(D = 1)] - P[R \mid do(D = 0)]$$

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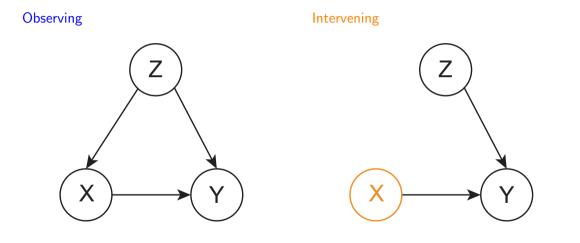
Causal Effect of Drug-Taking on Recovery:

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Inference problem: "Seeing" is not always the same as "doing"

Observing \neq **Intervening**: $P[Y \mid X = x]$ is not **generally** the same as $P[Y \mid do(X = x)]$

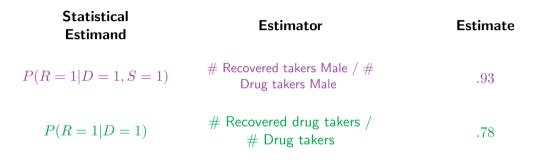
Two versions of the causal system

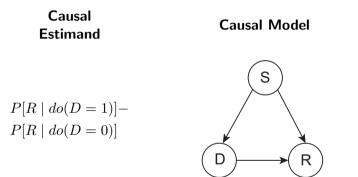


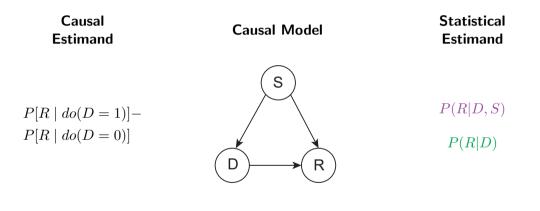
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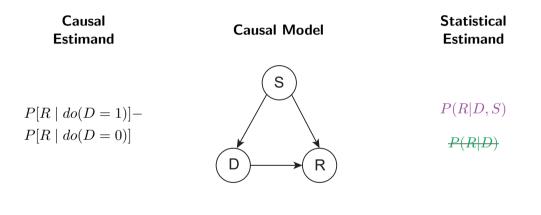
- 1 Causal models map causal dependencies onto statistical dependencies
 - *Regardless* of distributions and functional forms
- 2 Causal models allow us to define **causal effects** in the language of interventions and probabilities
- 3 Causal models tell us which when and how statistical estimands can act as causal estimands

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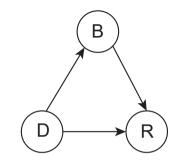






Post-Treatment Blood Pressure:

- Statistical information is exactly the same
- The drug works in part by decreasing blood pressure
- We should **not** condition on blood pressure



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Example from Pearl, Glymour & Jewell, 2016

Causal Inference in a nutshell

Causal Estimand

Causal Model





Statistical Estimand



Estimator

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Level Calves
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Simpsons Paradox

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Absolutely not a paradox.

• Confusion comes from a lack of clarity regarding our **causal estimand** and **causal model**

Statistical information *alone* cannot provide the answer

• Different DAGs can produce the exact same statistical dependencies in observational data

Causal models provide immediate conceptual clarity

• Miguel Hernan: Draw your assumptions before your conclusions!

Beyond Toy Examples

Causal Inference in Practice

Two broad frameworks for causal modeling

- Graphical Causal Models (Structural Causal Models)
- Potential Outcomes framework

Many tools available in these frameworks

- Causal Mediation analysis
- Cyclic Causal Models
- Instrumental Variables / Mendellian Randomization
- Synthetic Control & Interrupted Time Series
- "Target Trial" emulation



Miguel Hernán 🤣 @_MiguelHernan

1/

Six years ago we emulated a #TargetTrial of screening colonoscopy for colorectal #cancer using observational #Medicare data: pubmed.ncbi.nlm.nih.gov/27669524/

Today we publish preliminary findings from a truly randomized trial: nejm.org/doi/full/10.10...

Their main results side by side \P 1.75. Age 55-64 0.05 Age 70-74 v --- No screening 80-1.50----- Colonoscopy 70-1.25. 0.04 60-1.00-Invited group 0.03 0.75-0.98 (95% CI. 0.86-1.0 50-40-0.02 -Usual-care group 30-1.20 (95% CI, 1.10-1.29) 0.01 20 4 5 6 7 8 9 0 1 2 Vente Years since Randomization Annals of Int Med and 3 others

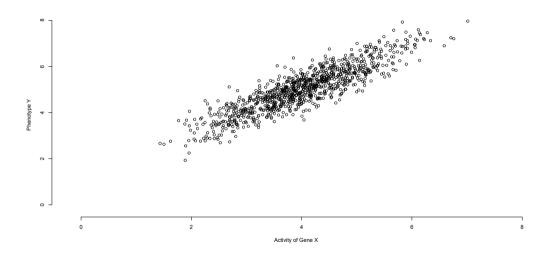
Statistical Learning / Machine Learning / "Data Science"

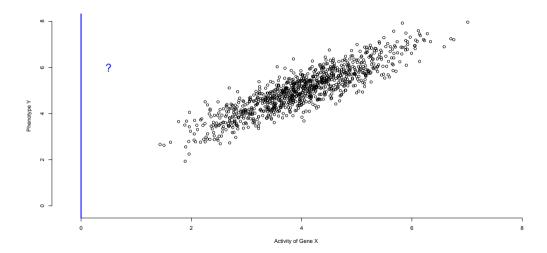
- Incredibly useful tools for learning *certain kinds* of representations of probability densities
- Usually: Those that minimize out-of-sample prediction error. But this is actually quite specific
- If I observe a new data point in *exactly the same circumstances* as I observed my training set, what types of values are likely to (co-)occur?

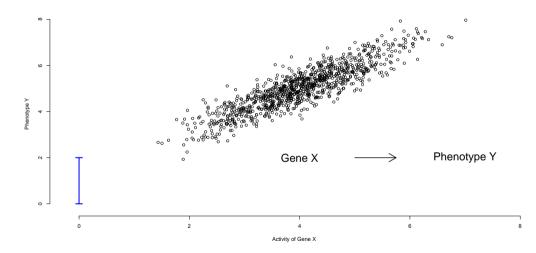
But is this what we actually want in most cases?

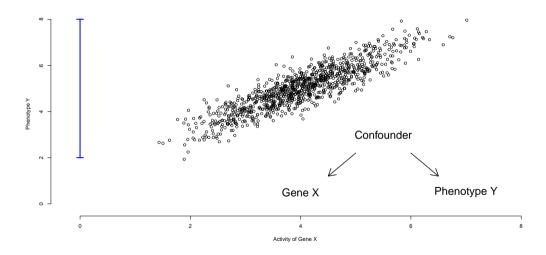
- Which treatment assignment would be best for this individual?
- What would happen if I change some policy, e.g., make more funding available for after-school programs?
- How do I make decisions which avoid gender bias?
- How do I make sure my ML model doesn't learn to classify data points based on some "undesirable" factor?

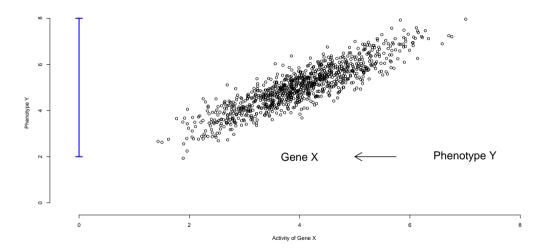
I would argue that answering these questions requires us to learn a *causal model*, not a purely statistical or predictive one

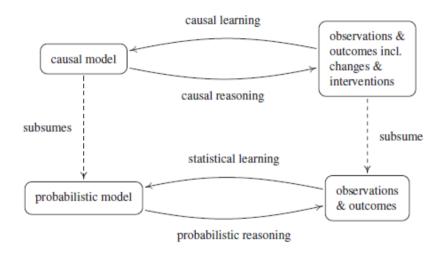














Toward Causal Representation Learning

This article reviews fundamental concepts of causal inference and relates them to crucial open problems of machine learning, including transfer learning and generalization, thereby assaying how causality can contribute to modern machine learning research.

By Bernhard Schölkopf[®], Francesco Locatello⁹, Stefan Bauer[®], Nan Rosemary Ke, Nal Kalchbrenner, Anirudh Goyal, and Yoshua Bengio⁹

Conclusions

Inappropriate reliance on (advanced) statistical modeling or statistical learning techniques, with no clear link to causal estimands or models

• Paradoxes, confusion, poor decisions result

Causal modeling can be powerful in reshaping how we approach statistical modeling

- Judea Pearl, Don Rubin, Jamie Robins, Miguel Hernan, Angrist & Imbens
- Controlling for as many variables as possible is **an obviously terrible idea** when estimating causal effects
- Minimizing OOS PE / classification error does not in any obvious way licence causal inferences (counterfactual predictions)

Researchers make causal inferences based on observational data all the time

• Better to be explicit and open about this so we can move forward

Thanks! (o.ryan@uu.nl | oisinryan.org)

- Statistical network analysis as ad-hoc causal discovery (e.g. Ryan, Bringmann, Schuurman, 2022)
- Causal models of dynamical systems from cross-sectional data (Ryan* & Dablander*, pre-print 2022)
- Estimating the effect of a hypothetical language training program on study success using administrative data. A pre-registered causal inference approach (Spit, Andringa, Ryan. In-principle acceptance)
- Causal and statistical estimands (e.g. Haslbeck*, Ryan*, Dablander* 2021)
- Constructing theories (Haslbeck*, Ryan*, Robinaugh*, Waldorp, Borsboom, 2021)
- Synthetic Control and Interrupted Time Series with large-scale administrative data (in progress)
- Causal Discovery of cyclic models (in progress)
- Teaching materials and workshops