Modelos gráficos causales: introducción y algunos de nuestros resultados recientes sobre reglas gráficas sobre ajustes eficientes

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Based on

Rotnitzky and Smucler, 2020, Journal of Machine Learning Research, 21 188: 1-86,

Smucler, Sapienza and Rotnitzky, 2021, Biometrika, 109, 1, 49-65.

Smucler and Rotnitzky, 2022. Journal of Causal Inference, 10, 1, 174-189

Guo, Perkovic and Rotnitzky, 2022, https://arxiv.org/abs/2202.11994

Academia Nacional de Ciencias Economicas, 6 de Septiembre, 2022

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- The analysis of a single modern medical study may use
  - mediation analysis (origin in psychology and sociology),
  - instrumental variables (origin in economics and genetics), and
  - marginal structural models (origin in epidemiology and biostatistics).

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- "Causal revolution" in great part due to the emergence and adoption of two formalisms:
  - Counterfactual Models
  - Graphical Models

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- In epidemiology and medical research: responsible for the acceptance and adoption of modern causal analytic techniques because they facilitate encoding complex causal assumptions and reasoning in an intuitive way
- Simple graphical rules exist to explain the potential biases of one's preferred estimation procedure and the possible remedial approaches.

- No graphical rules existed to explain efficiency (variance) in estimation
- In this talk: review graphical models and its use for understanding biases and summarize some of our own work towards filling this gap

#### An adjustment set



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#### Another adjustment set



#### Road map of the talk

- ► Gentle introduction to causal graphical models.
  - Definition and properties
  - Some examples of their use for detecting potential sources of bias

Some of our results on efficient adjustment sets

Rules for comparing adjustment sets for point exposure studies

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- Final remarks





$$\begin{split} V_1 &= f_1(\varepsilon_1) \\ V_2 &= f_2(\varepsilon_2) \\ V_3 &= f_3(\varepsilon_3) \\ V_4 &= f_4(\varepsilon_4) \\ V_5 &= f_5(V_1, \varepsilon_5) \\ \vdots \\ V_{11} &= f_{11}(V_5, V_7, \varepsilon_{11}) \\ V_{12} &= f_{12}(V_{11}, V_4, \varepsilon_{12}) \\ V_{13} &= f_{13}(V_8, V_{10}, V_{12}, \varepsilon_{13}) \\ \end{array}$$

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No omitted common cause assumption formalized as: the  $\varepsilon'_j s$  are mutually independent (Pearl, 1995)



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No omitted common cause assumption formalized as: the  $\varepsilon'_j s$  are mutually independent (Pearl, 1995)



• Graphical model with independent  $\varepsilon'_i s$  is tantamount to:

$$p\left(\mathbf{v}
ight)=\prod_{j}p\left(v_{j}|pa_{\mathcal{G}}\left(v_{j}
ight)
ight)$$

The collection of laws for V that factor like this is called a Bayesian Network B (G).

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# Causal Graphical Models in a nutshell: counterfactual world static intervention



$$\begin{split} V_1 &= f_1(\varepsilon_1) \\ V_2 &= f_2(\varepsilon_2) \\ V_3 &= f_3(\varepsilon_3) \\ V_4 &= f_4(\varepsilon_4) \\ V_5 &= f_5(V_1, \varepsilon_5) \\ \vdots \\ V_{11}^{\upsilon_{11}=0} &= 0 \\ V_{12}^{\upsilon_{11}=0} &= f_{12} \left( V_{11}^{\upsilon_{11}=0}, V_4, \varepsilon_{12} \right) \\ V_{13}^{\upsilon_{11}=0} &= f_{13} \left( V_8, V_{10}, V_{12}^{\upsilon_{11}=0}, \varepsilon_{13} \right) \\ & \varepsilon_{1}, \dots, \varepsilon_{13} \text{ omitted} \\ \text{non- common causes} \end{split}$$

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# Causal Graphical Models in a nutshell: counterfactual world static intervention



Corollary: counterfactual law is identified and given by

 $\rho_{(v_{11}=0)}\left(\mathbf{v}\right) = \prod_{j\neq 11} \rho\left(v_j | pa_{\mathcal{G}}\left(v_j\right)\right) \times I_{\{0\}}\left(v_{11}\right)$ 

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Causal Graphical Models in a nutshell: counterfactual world, deterministic dynamic intervention



$$\begin{split} V_{1} &= f_{1}\left(\varepsilon_{1}\right) \\ V_{2} &= f_{2}\left(\varepsilon_{2}\right) \\ V_{3} &= f_{3}\left(\varepsilon_{3}\right) \\ V_{4} &= f_{4}\left(\varepsilon_{4}\right) \\ V_{5} &= f_{5}\left(V_{1}, \varepsilon_{5}\right) \\ \vdots \\ V_{11}^{g} &= g\left(V_{9}\right) \\ V_{12}^{g} &= f_{12}\left(V_{11}^{g}, V_{4}, \varepsilon_{12}\right) \\ V_{13}^{g} &= f_{13}\left(V_{8}, V_{10}, V_{12}^{g}, \varepsilon_{13}\right) \end{split}$$

 $\varepsilon_1,\ldots,\varepsilon_{13}$  omitted non- common causes

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Corollary: counterfactual law is identified and given by

 $p_{g}(\mathbf{v}) = \prod_{j \neq 11} p\left(v_{j} | pa_{\mathcal{G}}(v_{j})\right) \times I_{\{g(v_{9})\}}(v_{11})$ 

# Causal Graphical Models in a nutshell: counterfactual world, random dynamic intervention



$$\begin{split} V_1 &= f_1\left(\varepsilon_1\right) \\ V_2 &= f_2\left(\varepsilon_2\right) \\ V_3 &= f_3\left(\varepsilon_3\right) \\ V_4 &= f_4\left(\varepsilon_4\right) \\ V_5 &= f_5\left(V_1, \varepsilon_5\right) \\ \vdots & & & \\ V_{11}^{\pi} &= g\left(V_9, U_{11}\right) \\ V_{12}^{\pi} &= f_{12}\left(V_{11}^{\pi}, V_4, \varepsilon_{12}\right) \\ V_{13}^{\pi} &= f_{13}\left(V_8, V_{10}, V_{12}^{\pi}, \varepsilon_{13}\right) \end{split}$$

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# Causal Graphical Models in a nutshell: counterfactual world, random dynamic intervention



$$V_{1} = f_{1} (\varepsilon_{1})$$

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$$V_{5} = f_{5} (V_{1}, \varepsilon_{5})$$

$$\vdots$$

$$V_{11}^{\pi} = g (V_{9}, U_{11})$$

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Corollary: counterfactual law is identified and given by

$$p_{\pi}\left(\mathbf{v}
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ight)$$

# Precursors, review papers in economics and an important reference

- Pearl's causal graphical model precursors:
  - In biology: Sewall Wright's *linear* structural equations models with normal errors (geneticist) → path analysis
  - ► In economics: Haavelmo's simultaneous structural equations model → allows non-recursiveness (simultaneous causation) and assumes parametric equations.
- For a review contrasting Pearl's and Haavelmo's models see Heckman and Pinto (2015). Causal Analysis After Haavelmo, Economic Theory.
- See also Imbens (2020) Potential Outcome and Directed Acyclic Graph Approaches to Causality: Relevance for Empirical Practice in Economics. Journal of Economic Literature
- For a unifying approach to potential outcomes and causal graphical models see T.S. Richardson, J.M. Robins (2013). Single World Intervention Graphs (SWIGs): A Unification of the Counterfactual and Graphical Approaches to Causality, In Foundations and Trends in Machine Learning, ISBN 13: 9781601988102

### Causal graphical models

#### Causal graphical models

a. Factual world. The law p of  $\mathbf{V} = (V_1, ..., V_J)$  belongs to Bayesian Network  $\mathcal{B}(\mathcal{G})$ , i.e. it factorizes as

$$p\left(\mathbf{v}
ight)=\prod_{j=1}^{J}p\left(v_{j}|p \mathsf{a}_{\mathcal{G}}\left(v_{j}
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where  $pa_{\mathcal{G}}(V_j)$  are the parents of  $V_j$  in  $\mathcal{G}$ .

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where  $p_{a_{\mathcal{G}}}(V_j)$  are the parents of  $V_j$  in  $\mathcal{G}$ .

b. Counterfactual world. For any  $\mathbf{A} = (A_1, ..., A_s) \subset \mathbf{V}$ , the distrib. of the data when a regime that assigns  $a_t$  to  $A_t$  with prob.  $\pi_t(a_t|\mathbf{Z}_t)$  is implemented in the population (where  $\mathbf{Z}_t$  are non-descendants of  $A_t$ ), is

$$p_{\pi}\left(\mathbf{v}
ight) = \prod_{V_{i}\in\mathbf{V}\setminus\mathbf{A}} p\left(v_{j}|pa_{\mathcal{G}}\left(v_{j}
ight)
ight) imes \prod_{t=1}^{s} \pi_{t}\left(a_{t}|\mathbf{z}_{t}
ight)$$

So,  $p_{\pi}$  is **identified** from p

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► Bayesian Network B (G) : collection of laws p for V that factorize as

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 $A \perp\!\!\!\perp_{\mathcal{G}} B \mid C \ (A \text{ and } B \text{ are d-separated by } C \text{ in } \mathcal{G})$ 

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• **d-separation:** a sound and complete graphical rule for determining whether a conditional independence holds **under any**  $p \in \mathcal{B}(\mathcal{G})$ .

 $A \perp\!\!\!\perp_{\mathcal{G}} B \mid C \ (A \text{ and } B \text{ are d-separated by } C \text{ in } \mathcal{G})$ 

Theorem (Geiger, Verma & Pearl, 1990):

 $A \perp\!\!\!\perp_{\mathcal{G}} B \mid C \Leftrightarrow$ A is cond. indep. of B given C under any  $p \in \mathcal{B}(\mathcal{G})$ 

### d-separation

- A, B single vertices,  $C \subset V \setminus \{A, B\}$
- ▶ a path from A to B is blocked by C if either

(1) at least one non-collider is in C





(2)  $\exists$  at least one collider, such that neither itself nor its descendants is in C





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• A set A is d-separated from another set B by  $C \subset V \setminus \{A, B\}$  if all  $A_j \in A$  and  $B_k \in B$  are d-separated by C, in which case we write

 $A \perp \!\!\!\perp_{\mathcal{G}} B \mid C$
#### Road map of the talk

- ► Gentle introduction to causal graphical models.
  - Definition and properties
  - Some examples of their use for detecting potential sources of bias

Some of our results on efficient adjustment sets

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# Two potential sources of bias in your causal analysis

NOT conditioning on common causes Conditioning on a common effect





**Confounding bias** 

Berkson's bias

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# Berkson bias

Two variables that are marginally independent will typically be dependent if we condition on a common effect of both variables. (Berkson, 1946) **Example** 



Suppose P(gene 1) = P(gene 2) = 0.02, genes are marginally independent and Disease if and only if at least one of the two genes is present, i.e.

$$X = 1 - (1 - A) (1 - Y)$$

Then,

```
P (Gene 1|Disease, Not Gene 2) = 1
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P (Gene 1|Disease, Gene 2) = P (Gene 1| Gene 2) = 0.02

So, Gene 1 and Gene 2 are negatively correlated conditional on having the disease.

# M bias



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**Example**: sequentially randomized trial of the effect of High vs Low dose of Highly Active Antiretroviral Therapy (HAART) at months 0 and 3 on Viral Load (high vs low) at month 6. (Assume in the graph all variables are binary)



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Causal sharp null hypothesis  $H_0^{causal}$  that  $(A_0, A_1)$  has no causal effect on Y is represented by the graph



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1. **Regression controlling for** *L* **fails:** Suppose we fit a saturated (and hence correctly specified) logistic regression model

and to test  $H_0^{causal}$  we test the null hypothesis

$$H_0: (\gamma_0, \gamma_1, \gamma_2, \gamma_3, \eta_2, \eta_3) = (0, 0, 0, 0, 0, 0)$$

The test does not preserve the  $\alpha$ - level because  $H_0^{causal} \neq H_0$  since the path  $Y - U - L - A_0$  is open when we condition on the collider L.

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The test does not preserve the  $\alpha$ - level because  $H_0^{causal} \neq H_0$  since the path  $Y - U - L - A_0$  is open when we condition on the collider L.

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Causal sharp null hypothesis  $H_0^{causal}$  that  $(A_0, A_1)$  has no causal effect on Y is represented by the graph



2. Regression that does not control for *L* also fails: Suppose we fit a saturated (and hence correctly specified) logistic regression model

logitPr 
$$(Y = 1 | A_0, A_1) = A_1 (\alpha_0 + \alpha_1 A_0) + (\nu_0 + \nu_1 A_0)$$

and to test  $H_0^{causal}$  we test the null hypothesis

$$H_0^*: (\alpha_0, \alpha_1, \nu_1) = (0, 0, 0)$$

The test does not preserve the  $\alpha$ - level because  $H_0^{causal} \neq H_0^*$  since the path  $Y - U - L - A_1$  is open when we fail to condition on *L*.

# Identification

- Y<sub>a₀,a₁</sub>: potential outcome when everybody in the study population takes treatment A₀ = a₀, A₁ = a₁.
- Result (Robins, 1986): under the causal graphical model represented by the graph



the probability  $\Pr(Y_{a_0,a_1} = 1)$  of high viral load in the counterfactual world in which everybody receives treatment  $A_0 = a_0, A_1 = a_1$  is identified and given by

$$\Pr(Y_{a_0,a_1} = 1) = \sum_{l=0}^{1} \Pr(Y = 1 | A_0 = a_0, A = a_1, L = l) \Pr(L = l | A_0 = a_0)$$

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Counterfactual law.

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• Then for  $Y = V_J$ ,

$${E_\pi \left[ Y 
ight]} = \int y \prod\limits_{j: V_j \in {f V} ackslash A} {p\left( {{v_j}} 
ight|{f pa_{\mathcal G}}\left( {{v_j}} 
ight)} 
ight) imes \pi \left( {f a} 
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► Then for 
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,  

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▶ But under the Bayesian Network E<sub>π</sub>(Y) is equal to many other functionals

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# Adjustment formula and adjustment sets

Adjustment formula:

$$\underbrace{E_{\pi}[Y]}_{\text{intervention mean}} = \underbrace{\sum_{a=0}^{1} \int E[Y|A = a, \mathbf{L} = \mathbf{I}] \pi(a|\mathbf{z}) p_{\mathbf{L}}(\mathbf{I}) d\mathbf{I}}_{\text{g-functional}}$$
$$= E_{p} \left[ \frac{\pi(A|\mathbf{Z})}{p(A|\mathbf{L})} Y \right]$$

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where  $\textbf{Z} \subset \textbf{L} \subset \textbf{V}$ 

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where  $\textbf{Z} \subset \textbf{L} \subset \textbf{V}$ 

- Definition: A Z- adjustment set for a single trx A and outcome Y is any L disjoint with A and Y such that
  - $\mathbf{Z} \subset \mathbf{L}$  and,
  - Under the causal graphical model, for any regime  $\pi(A|\mathbf{Z})$ ,  $E_{\pi}[Y]$  is equal to the corresponding adjustment formula.

#### Adjustment formula and adjustment sets

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  - $\mathbf{Z} \subset \mathbf{L}$  and,
  - Under the causal graphical model, for any regime  $\pi(A|\mathbf{Z})$ ,  $E_{\pi}[Y]$  is equal to the corresponding adjustment formula.
- ► If Z = Ø, then we say L is a static adjustment set.

Generalized adj. criterion for static (i.e. Z = Ø) treatments (Shpitzer. et. al., 2010, Perkovic et. al., 2015, 2018): L is static adj. set iff

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  - L is neither a mediator, nor descendant of Y or of a mediator

L blocks all non-causal paths between A and Y.

- ▶ Generalized adj. criterion for static (i.e. Z = Ø) treatments (Shpitzer. et. al., 2010, Perkovic et. al., 2015, 2018): L is static adj. set iff
  - L is neither a mediator, nor descendant of Y or of a mediator
  - L blocks all non-causal paths between A and Y.
- Result (Smucler and Rotnitzky, 2020):

Class of all Z – adj sets =  $\{L : L \text{ is a static adj. set and } Z \subset L\}$ 

# Static adjustment set



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#### Another static adjustment set



# An invalid Z-adjustment , Z= previous injury



# A valid Z-adjustment set, Z= previous injury



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▶ **Recall:** a **Z**- adj. set **L** satisfies that for any regime  $\pi(A|\mathbf{Z})$ , the counterfactual mean  $E_{\pi}(Y)$  is equal to

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#### Questions that we addressed:.

- Given two adjustment sets, are there graphical rules to determine which one yields an estimator with smaller variance?
- Is there a universally optimal adjustment set and, if so, what graphical rules determine it?

# Related literature

- Henckel, Perkovic and Maathuis (2019) provided graphical rules
  - for comparing certain pairs of static adjustment sets
  - ▶ for determining the globally optimal static adjustment set
- Also, Kuroki and Miyakawa, 2003 and Kuroki and Cai 2004.
- These works assume:
  - causal graphical linear model, i.e.  $V_j = \beta_j^T pa_G(V_j) + \varepsilon_j, \{\varepsilon_j : j\}$  indep.

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- Works connected with efficiency implications of inclusion of overadjustment and precision variables in regression and in semip. estimation of ATE:
  - Linear regression: Cochran (1968)
  - Non-linear regression: Mantel and Haenszel (1959), Breslow (1982), Gail (1988), Robinson and Jewell (1991), Neuhaseuser and Becher (1997) and De Stavola and Cox, (2008).
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#### Supplementing adjustment sets with precision variables.

▶ Lemma 1. Suppose B is a Z-adj. set and G, disjoint with B, satisfies

 $A \perp \!\!\!\perp_{\mathcal{G}} \mathbf{G} \mid \mathbf{B}$ 

then,  $\mathbf{G} \cup \mathbf{B}$  is also a  $\mathbf{Z}$ -adj. set and for all  $p \in \mathcal{B}(\mathcal{G})$  and all regimes  $\pi(A|\mathbf{Z})$ 

 $\sigma_{\pi,\mathbf{G}\cup\mathbf{B}}^{2}\left(\boldsymbol{p}\right)\leq\sigma_{\pi,\mathbf{B}}^{2}\left(\boldsymbol{p}\right)$ 

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$$\sigma_{\pi,\mathbf{B}}^{2}\left(p\right) - \sigma_{\pi,\mathbf{G}\cup\mathbf{B}}^{2}\left(p\right) = E\left[\left\{\frac{1}{P\left(A=a|\mathbf{B}\right)} - 1\right\} var\left\{E\left(Y|A=a,\mathbf{G},\mathbf{B}\right)|A=a,\mathbf{B}\right\}\right]$$



#### Deleting overadjustment variables

**Lemma 2.** Suppose  $\mathbf{G} \cup \mathbf{B}$  is a  $\mathbf{Z}$ -adj. set and  $\mathbf{B}$  satisfies

 $Y \perp \!\!\!\perp_{\mathcal{G}} \mathbf{B} \mid \mathbf{G}, A$ 

If  $Z \subset G$ , then G is *also a* Z-*adj. set* and for all  $p \in \mathcal{B}(\mathcal{G})$  and all regimes  $\pi(A|Z)$ 

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If  $Z \subset G$ , then G is *also a* Z-*adj. set* and for all  $p \in B(G)$  and all regimes  $\pi(A|Z)$ 

$$\sigma_{\pi,\mathbf{G}}^{2}\left(\boldsymbol{p}\right) \leq \sigma_{\pi,\mathbf{G}\cup\mathbf{B}}^{2}\left(\boldsymbol{p}\right)$$

• In particular, for the static regime  $\pi$  that sets A to a,

$$\sigma_{\pi,\mathbf{G}\cup\mathbf{B}}^{2} - \sigma_{\pi,\mathbf{G}}^{2} = E\left[var\left(Y|A=a,\mathbf{G}\right)\left\{\frac{1}{P\left(A=a|\mathbf{B},\mathbf{G}\right)} - \frac{1}{P\left(A=a|\mathbf{G}\right)}\right\}\right]$$



#### Comparing two arbitrary adjustment sets

▶ Corollary: Suppose that G and B are two Z-adj. sets such that

 $A \perp\!\!\perp_{\mathcal{G}} (\mathbf{G} \backslash \mathbf{B}) \mid \mathbf{B}$ 

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$$\begin{array}{l} Y \ \ \amalg _{\mathcal{G}} \ \ (\mathbf{B} \backslash \mathbf{G}) \ \mid \mathbf{G}, A \end{array}$$
  
Then, for all  $p \in \mathcal{B} \left( \mathcal{G} \right)$  and all regimes  $\pi \left( A | \mathbf{Z} \right)$   
 $\sigma_{\pi, \mathbf{G}}^2 \left( p \right) \leq \sigma_{\pi, \mathbf{B}}^2 \left( p \right)$ 

Proof:

gain due to supplementation with precision component  $\mathbf{G} \setminus \mathbf{B}$ 

gain due to deletion of noisy component  $\mathbf{B} \setminus \mathbf{G}$ 

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#### Not all adjustment sets are comparable



- $(O_1, W_2)$  is preferable to  $(O_2, W_1)$  if green association stronger than brown, and blue association weaker than red
- $(O_2, W_1)$  is preferable to  $(O_1, W_2)$  if brown association stronger than green, and red association weaker than blue

• but...  $(O_1, O_2)$  is more efficient than both

#### Optimal adjustment set

▶ Theorem: (Henckel, et. al. (2019)). The set

**0** = non-descendants of A that are parents of Y or of vertices in the causal path bw A and Y

is a  $\mathit{static}$  adjustment set. Furthermore, for any other static adjustment set  $\boldsymbol{\mathsf{L}},$ 

 $A \perp\!\!\perp_{\mathcal{G}} (\mathbf{O} \backslash \mathbf{L}) \mid \mathbf{L}$ 

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 Corollary (Rotnitzky and Smucler, 2020): O is the globally optimal static adjustment set.

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 Corollary (Rotnitzky and Smucler, 2020): O is the globally optimal static adjustment set.

► Lemma (Smucler, Sapienza and Rotnitzky, 2021): O ∪ Z is the globally optimal Z - adjustment set

# Globally optimal static adjustment set



# Optimal personalized adjustment set



#### Road map of the talk

- Gentle introduction to causal graphical models.
  - Definition and properties
  - Some examples of their use for detecting potential sources of bias

Some of our results on efficient adjustment sets

Rules for comparing adjustment sets for point exposure studies

- Summary of other results
- Final remarks

## Our contributions

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 In Smucler, Sapienza and Rotnitzky (2021) we characterize sufficient conditions for an optimal observable adjustment set to exist

#### Time dependent treatments

▶ Suppose  $A_1$  and  $A_2$  are two treatments,  $A_1 \in \operatorname{nd}_{\mathcal{G}}(A_2)$ . Under a causal graphical model represented by a graph G, the mean of  $Y_{a_0,a_1}$  when the static regime that sets  $A_0$  to  $a_0$  and  $A_1$  to  $a_1$  is

$$\begin{split} E\left(Y_{a_{0},a_{1}}\right) &= E\left\{\frac{I_{a_{0}}\left(A_{0}\right)}{p\left(a_{0}\right|pa_{\mathcal{G}}\left(A_{0}\right)\right)}\frac{I_{a_{1}}\left(A_{1}\right)}{p\left(a_{1}\right|pa_{\mathcal{G}}\left(A_{1}\right)\right)}Y\right\} \\ &= E\left\{E\left[E\left[Y\right|a_{0},a_{1},pa_{\mathcal{G}}\left(A_{0}\right),pa_{\mathcal{G}}\left(A_{1}\right)\right]|a_{0},pa_{\mathcal{G}}\left(A_{0}\right)\right]\right\} \end{split}$$

▶ Definition:  $\mathbf{L} = (\mathbf{L}_0, \mathbf{L}_1) \subset \mathbf{V}$  is a static time dependent adjustment set relative to trxs  $(A_0, A_1)$  and outcome Y in G iff for all  $P \in \mathcal{B}(\mathcal{G})$ ,

$$E(Y_{a_0,a_1}) = E\{E[E[Y|a_0, a_1, \mathbf{L}_0, \mathbf{L}_1]|a_0, \mathbf{L}_0]\}$$

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## Time dependent treatments

Example:



- $X_0$  is a time 0 adjustment set  $(= \mathbf{L}_0)$
- $X_1$ , U and  $(X_1, U)$  are time 1 adjustment sets  $(= L_1)$
- In Rotnitzky and Smucler, 2020, we derived rules for comparing static time dependent adjustment sets and showed by example that an optimal adjustment set need not exist.

# Study design.

- Assign cost to each graph variable and find the adjustment set leading to smallest estimation variance:
  - subject to a cost constraint  $\rightarrow$  a universal solution does not exist



► among adjustment sets of minimum cost → for point exposure we provide the universal solution in Smucler and Rotnitzky, 2022, and graphical rules for finding it

Semip. efficient estimation vs optimal non-parametric adjusted estimation



• The interventional mean  $E(Y^a)$  is

$$E\left[E\left(Y|A=a,V,W\right)\right] = \int E\left(Y|A=a,V=v,W=w\right)\underbrace{p\left(v\right)p\left(w\right)}_{=p\left(v,w\right)}dvdw$$

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- Optimal non-parametric adjusted estimator ignores restrictions on the marginal law of covariates, i.e. that V and W are marginally independent.
- Semiparametric efficient (SE) exploits these restrictions and can be much much more efficient than optimally adjusted NP estimator.

However ... in some graphs the optimally adjusted estimator is efficient



• With discrete data the MLE of  $p_a(y)$  under  $\mathcal{G}$  is

 $\widehat{p}_{a,MLE}\left(y\right) = \sum_{m,o} \mathbb{P}_{n}\left(y|m,a\right) \mathbb{P}_{n}\left(m|a,o\right) \mathbb{P}_{n}\left(o\right)$ 

Surprisingly,  $\hat{p}_{a,MLE}(y)$  is asym. equivalent to the MLE of  $p_a(y)$  under  $\mathcal{G}^*$  is

$$\widetilde{p}_{a,MLE}(y) = \sum_{o} \mathbb{P}_{n}(y|o,a) \mathbb{P}_{n}(o)$$

In Rotnitzky and Smucler (2000) we characterized the graphs in which the optimally adjusted estimator is semiparametric efficient

# Graph reduction for semiparametric efficient estimation of a counterfactual mean

- ▶ In Guo, Perkovic and Rotnitzky, 2022, we derived the following.
- Given a graph G we derived an algorithm that outputs another graph G\*over a subset of the variables in G such that
  - ▶ the semiparametric variance bound for estimation of  $E(Y_a)$  in model  $\mathcal{B}(\mathcal{G})$  and in model  $\mathcal{B}(\mathcal{G}^*)$  agree
  - $\mathcal{G}^*$  is the smallest such possible graph in the sense that all variables in  $\mathcal{G}^*$  are informative. More precisely, the efficient influence function for  $E(Y_a)$  is a function of every variable in  $\mathcal{G}^*$  for at least one P in  $\mathcal{B}(\mathcal{G}^*)$

#### Final remarks

- **Estimation via adjustment vs semip.** efficient estimation:
  - Usual variance/bias trade-off: adjustment relies on less model assumptions
  - Equally or perhaps even more importantly: efficient estimation requires estimation of each cond. density  $p(V_j | pa_{\mathcal{G}}(V_j))$ . Even debiased, influence-function based, i.e. one-step estimation, will hardly control the estimation bias of these densities.

# Open problems

- Inference about the functional returned by the ID algorithm when no observable adj. set exists
  - Some special cases have been studied, e.g. the generalized front door formula, (Fulcher, et. al. 2019). General theory for an arbitrary functional not yet available.

Optimal adj. sets and efficient estimation for other parameters e.g., trx effect on the treated, and natural direct and indirect effects

#### THANKS!

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