### FEDERATED CAUSAL INFERENCE: MULTI-SOURCE ATE ESTIMATION BEYOND META-ANALYSIS

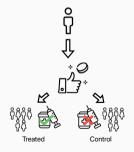
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Joint work with Rémi Khellaf and Julie Josse

2024 IMS International Conference on Statistics and Data Science (ICSDS) Session "Federated Causal Inference Meets Meta-Analysis" December 17, 2024

### Goal of causal inference: measure the effect of a treatment on an outcome

Randomized Controlled Trials (RCTs) are the gold standard but limited scope (stringent eligibility criteria, limited sample size...)



Meta-analysis (aggregating estimated effects from multiple studies) is at the top of the pyramid of evidence



- Meta-analyses still face significant challenges:
  - Data heterogeneity across studies (sample sizes, populations, center effects...)
  - Difficulty to share individual-level data due to data silos and personal data regulations
- **Our work** bridges causal inference and federated learning [Kairouz et al., 2021] to better estimate average treatment effects from decentralized data sources
  - 1. We consider several estimators with varying communication costs
  - 2. We study their statistical performance under various types of data heterogeneity
  - 3. We validate on numerical simulations and provide guidelines for practitioners

- [Xiong et al., 2023] propose to use one-shot aggregation to federate the outcome or propensity score model, but it is unclear when these estimators should be preferred to other methods (e.g., traditional meta-analysis)
- [Vo et al., 2022b] employ a Bayesian framework using Gaussian processes but is restricted to uniform data distributions across sources
- [Vo et al., 2022a, Han et al., 2021, Han et al., 2023, Makhija et al., 2024, Guo et al., 2024] focus on transferring causal estimates from one source to another, while our work aims to estimate causal effects across the joint population

## **PROBLEM SETTING**

#### **REMINDER: CLASSIC RCT FRAMEWORK**

- Estimate effect of treatment W on outcome Y given covariates X, with  $W_i \sim \mathcal{B}(p)$
- Average Treatment Effect (ATE) measured as a risk difference  $\tau = \mathbb{E}[Y_i(1) Y_i(0)]$

Oha Causiatas Trastasat Outcome Datastic Outcome

Obs.	Covariates			Ireatment	Outcome	Potential Outcomes	
i	<i>X</i> <sub>1</sub>	X <sub>2</sub>	<i>X</i> <sub>3</sub>	W	Y	Y(1)	Y(0)
1	2.3	1.5	Μ	1	3.2	3.2	??
2	2.2	3.1	F	0	2.8	??	2.8
3	3.5	2.0	F	1	2.1	2.1	??
:	÷	÷	÷	:	÷	:	÷
<i>n</i> – 1	3.7	2.0	F	0	2.8	??	2.8
n	2.5	1.7	Μ	1	3.2	3.2	??

### OUR SETTING: DECENTRALIZED HETEROGENEOUS RCTS

• We consider K decentralized and potentially heterogeneous RCTs (studies) from different sources and want to estimate the ATE given by  $\tau = \mathbb{E} \left( \mathbb{E}(Y^{(1)} - Y^{(0)} | H) \right)$ 

Source	Obs.	Covariates		Treatment Outcomes				
Н	i	<i>X</i> <sub>1</sub>	X <sub>2</sub>	<i>X</i> <sub>3</sub>	W	Y	Heterogeneity in	
1	1	2.3	1.5	Μ	1	3.2	treatment allocation	
1	2	2.2	3.1	F	0	2.8		
	:	:	:	÷	:	:	(H)	
2	1	4.5	5.0	F	1	4.1		
:	:	:	:	:	:	:		
К	1	3.7	2.0	F	0	2.8	( W ) ( Y	
:	:	÷	÷	÷	:	:		
К	n <sub>K</sub>	2.5	1.7	Μ	0	3.2		

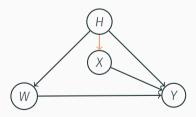
How to estimate  $\tau$  without pooling together individual-level data?

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1	2	2.2	3.1	F	0	2.8
:	÷	÷	÷	÷	:	:
2	1	4.5	5.0	F	1	4.1
:	÷	÷	÷	÷	:	:
K	1	3.7	2.0	F	0	2.8
:	÷	÷	÷	÷	:	:
K	n <sub>K</sub>	2.5	1.7	Μ	0	3.2

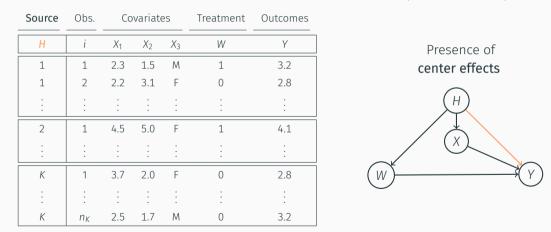
Heterogeneity in covariates distribution



How to estimate  $\tau$  without pooling together individual-level data?

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• We consider K decentralized and potentially heterogeneous RCTs (studies) from different sources and want to estimate the ATE given by  $\tau = \mathbb{E} \left( \mathbb{E}(Y^{(1)} - Y^{(0)} | H) \right)$ 



How to estimate  $\tau$  without pooling together individual-level data?

• For now, same linear outcome model for all studies:

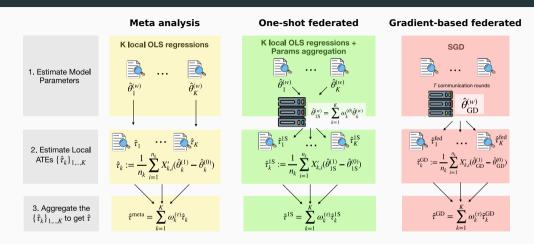
$$\forall k: \quad Y_{k,i}^{(w)} = c^{(w)} + X_{k,i}\beta^{(w)} + \varepsilon_{k,i}^{(w)}, \quad \text{with } \mathbb{E}\left[X_k^\top \varepsilon_{k,i}^{(w)}\right] = 0, \mathbb{V}\left(\varepsilon_{k,i}^{(w)} \mid X_k\right) = \sigma^2$$

- Standard assumptions (consistency, positivity, unconfoundedness)
- Ideal baseline: estimator  $\hat{\tau}_{\text{pool}} = \frac{1}{n} \sum_{i=1}^{n} X'_i(\hat{\theta}^{(1)}_{\text{pool}} \hat{\theta}^{(0)}_{\text{pool}})$  on pooled data, where

$$\hat{\theta}_{ ext{pool}}^{(w)} = (\hat{c}_{ ext{pool}}^{(w)}, \hat{eta}_{ ext{pool}}^{(w)}) = ({X'}^{(w)^{ op}} {X'}^{(w)})^{-1} {X'}^{(w)^{ op}} Y^{(w)}$$
 is the OLS estimator and  ${X'}^{(w)} = [1, X^{(w)}]$ 

+  $\hat{\tau}_{\rm pool}$  always has lower variance than the simple difference-in-means estimator [Benkeser et al., 2021, Lei and Ding, 2021] Federated Estimators

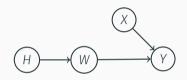
#### THREE TYPES OF FEDERATED ESTIMATORS



- Meta and one-shot require local sample size  $n_k^{(w)} \ge d$  for k, w
- Aggregation: sample size weights (SW) or inverse variance weights (IVW)

### **COMPARISON OF THE ESTIMATORS**

HOMOGENEOUS SETTING



- The source membership variable *H* only affects the treatment allocation scheme
- Let  $W_{k,i} \sim \mathcal{B}(p_k)$

### SUMMARY OF RESULTS

Estimators are unbiased but differ by their asymptotic variance and communication costs:

Estimator	Notation	$\mathbb{V}^{\infty}$	Com. rounds	Com. cost
Meta-SW	$\hat{ au}_{Meta-SW}$	$\frac{\sigma^2}{n} \sum_{k=1}^{K} \frac{\rho_k}{p_k(1-p_k)} + \frac{1}{n} \ \beta^{(1)} - \beta^{(0)}\ _{\Sigma}^2$	1	0(1)
Meta-IVW	$\hat{ au}_{Meta-IVW}$	$\Big(\sum_{k=1}^{K} \left(\sigma^2 \frac{n\rho_k}{p_k(1-p_k)} + \frac{1}{n_k} \ \beta^{(1)} - \beta^{(0)}\ _{\Sigma}^2\right)^{-1}\Big)^{-1}$	1	O(1)
1S-SW	$\hat{ au}_{ ext{1S-SW}}$	$V_{\rm pool}$	2	0(d)
1S-IVW	$\hat{ au}_{ extsf{1S-IVW}}$	$V_{\rm pool}$	2	$O(d^2)$
GD	$\hat{\tau}_{\mathrm{GD}}$	$V_{\rm pool}$	T + 1	O(Td)
Pool	$\hat{ au}_{pool}$	$V_{\text{pool}} = \frac{\sigma^2}{n} \frac{1}{p(1-p)} + \frac{1}{n} \ \beta^{(1)} - \beta^{(0)}\ _{\Sigma}^2$	_	_

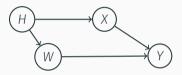
with  $\rho_k = \mathbb{P}(H = k) = \mathbb{E}\left[\frac{n_k}{n}\right]$  and  $p = \sum_{k=1}^{K} \frac{n_k}{n} p_k$ 

### NUMERICAL ILLUSTRATION (K = 5 and d = 10)



### COMPARISON OF THE ESTIMATORS

**HETEROGENEOUS DISTRIBUTIONS** 



- Distributional shift across sources:  $H \not \perp X \implies \tau_k \neq \tau_{k'}$
- Global ATE is given by  $\tau = \sum_{k=1}^{K} \rho_k \tau_k$  with  $\rho_k = \mathbb{P}(H = k) = \mathbb{E}\left[\frac{n_k}{n}\right]$

- +  $\hat{ au}_{ ext{meta-IVW}}$  is biased because inverse variance weights give biased estimates of the  $ho_k$
- $\cdot \mathbb{V}^{\infty}(\hat{\tau}_{\text{pool}}) \!=\! \mathbb{V}^{\infty}(\hat{\tau}_{\text{GD}}) \!=\! \mathbb{V}^{\infty}(\hat{\tau}_{\text{1S-IVW}}) \!\leq\! \mathbb{V}^{\infty}(\hat{\tau}_{\text{meta-SW}})$
- $\hat{\tau}_{1S-SW}$  is robust to heterogeneous covariances  $\{\Sigma_k\}_k$  but has larger variance for different means  $\{\mu_k\}_k$

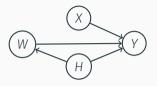
 $X_k \sim \mathcal{N}(\mu_k, \Sigma_k)$ 



### COMPARISON OF THE ESTIMATORS

PRESENCE OF CENTER EFFECTS

#### HETEROGENEITY FROM CENTER EFFECTS



- Studies may have different baselines in individual outcomes due to varying practices or organizational contexts (e.g. hospital specialized in oncology)
- We model this by a fixed effect of the source H onto the outcome Y:

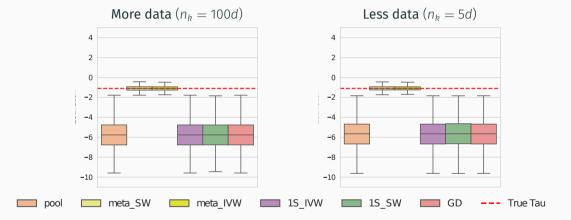
$$Y_{k,i}^{(w)} = c^{(w)} + \frac{h_k}{h_k} + X_{k,i}\beta^{(w)} + \varepsilon_i(w)$$

(Note: the CATEs  $\mathbb{E}[Y(1) - Y(0)|X, H]$  remain the same across sources)

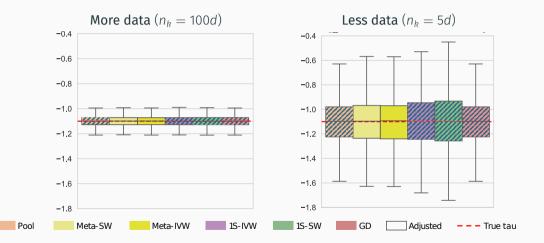
• Caution: *H* is now a confounder!

- +  $\hat{ au}_{
  m meta-SW}$  and  $\hat{ au}_{
  m meta-IVW}$  naturally account for the center effects
- Other federated estimators are biased and need to be adjusted
  - One-shot estimators: share and aggregate only the covariates coefficients  $\hat{\beta}_k$ , while keeping intercepts local
  - GD estimators: add H as an additional covariate

#### NUMERICAL ILLUSTRATION



### NUMERICAL ILLUSTRATION



**CONCLUSION & PERSPECTIVES** 

#### SUMMARY: DECISION DIAGRAM FOR PRACTITIONERS

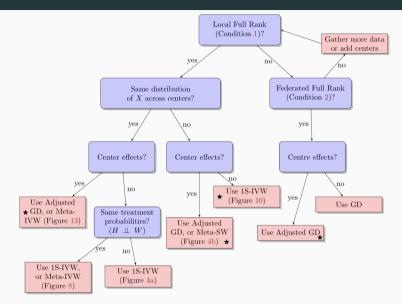


Figure 6: Decision Diagram for Practitionners. The sign  $\bigstar$  denotes scenarios where the DM estimator is biased.

- Extend to observational studies (e.g., federated IPW and AIPW) and nonlinear models
- Handle covariate mismatch across sources
- · Consider non-collapsible causal measures (e.g., odds ratio)
- Provide robust privacy guarantees (differential privacy)

# THANK YOU FOR YOUR ATTENTION! QUESTIONS?

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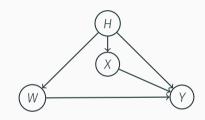
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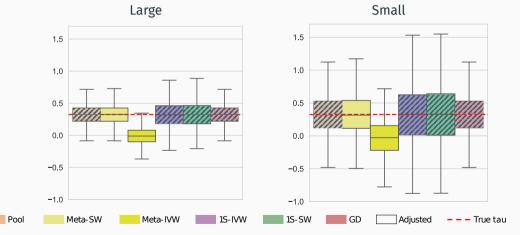
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### FULL HETEROGENEITY - NUMERICAL ILLUSTRATION

Different  $h_k, p_k, \mu_k, \Sigma_k$ 



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