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# **Selective Randomization Inference for Adaptive Experiments**

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## **Collaborators**



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 $,Z_i, X_i)_{i=1,...,n}$   $\longrightarrow$ 

 $\rightarrow$  Inference without modelling- or i.i.d. data-assumptions



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- P-value:  $\mathbb{P}(\ T(Z^*, Y(\cdot), X) \leq T(Z, Y(\cdot), X) | Y(\cdot), X, Z),$ where  $Z^* \stackrel{D}{=} Z$  and  $Z^* \perp\!\!\!\perp Z \mid Y(\cdot), X$

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## **Example**











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## **Graphical Model**

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$$
\boxed{X, Y(\ \cdot \ )}
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- Selective choice:  $S_k$
- Short-hand:  $W = (R, X_R, Y_R(\cdot))$



 $X_{R_2}$ 

*<u>Stage 2</u>* 

 $R<sub>2</sub>$ 

 $Y_{R_2}(\cdot)$ 







• **Assumption (A1):** *K* ∏ *k*=1

 $q(z | w) := \prod P(Z_k = z_k | R_{[k]} = r_{[k]}, X_{R_{[k]}} = x_{R_{[k]}}, Y_{R_{[k-1]}} = y_{R_{[k-1]}}, Z_{[k-1]} = z_{[k-1]})$  is known.  $\mathbb{P}(Z_k = z_k \mid R_{[k]} = r_{[k]}, X_{R_{[k]}} = x_{R_{[k]}}, Y_{R_{[k-1]}} = y_{R_{[k-1]}}, Z_{[k-1]} = z_{[k-1]})$ 

- **Assumption (A1):** *K* ∏ *k*=1  $\mathbb{P}(Z_k = z_k \mid R_{[k]} = r_{[k]}, X_{R_{[k]}})$
- **Assumption (A2):**  $Z_k \perp\!\!\!\perp Y_{R_{[k]}}(\cdot) \mid R_{[k]}.$



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- Assumption (A3):



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$$
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 $, Y_{R_k}(\cdot) \perp Z_{[k-1]} | W_{[k-1]}$ , *Sk*−<sup>1</sup> ∀ *k* ∈ [*K*]

$$
P, X_{R_{[k]}}, Y_{R_{[k-1]}}, Z_{[k-1]} \qquad \forall k \in [K]
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- Null hypothesis:  $Y_i(1) Y_i(0) = 0$  for all/subset of units
- Condition on  $W$  and compare observed value of statistic  $T(Z, W)$  against values  $T(Z^*,W)$  under alternative treatment assignments  $Z^*$ .
- $\mathbb{P}(T(Z^*,W) \leq T(Z,W) \mid W,Z)$ , where  $Z^* \stackrel{D}{=} Z$  and  $Z^* \perp Z \mid W$

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- Is there a problem when the experiment is adaptive?

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	- Data splitting (Cox, 1975):

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P_{sel} = \mathbb{P}(T(Z^*, W) \leq T(Z, W) \mid W, Z, S(Z^*) = S(Z))
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- Formula for the selective randomization distribution:

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\mathbb{P}(Z = z \mid W = w, S(Z) = s) = \frac{\mathbf{1}\{S(z) = s\} \cdot q(z \mid w)}{\sum_{z'} \mathbf{1}\{S(z') = s\} \cdot q(z' \mid w)}
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• Formula for p-value:

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P_{sel} = \frac{\sum_{z^*} \mathbf{1} \{ T(z^*, W) \le T(Z, W) \} \cdot \mathbf{1} \{ S(z^*) = S(Z) \} \cdot q(z^* | W)}{\sum_{z^*} \mathbf{1} \{ S(z^*) = S(Z) \} \cdot q(z^* | W)}
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## **Computation**

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## **Computation**

and compute

$$
\hat{P}_M := \frac{1 + \sum_{j=1}^M \mathbb{1}\{T(z_j^*, W) \le T(Z, W)\}}{1 + M}.
$$

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• Monte Carlo approximation: Generate  $M$  feasible samples  $(z_j^*)_{j=1}^M$ , i.e.  $S(z_j^*) = S(Z)$ , *M*  $J_{j=1}^{M}$ , i.e.  $S(z_{j}^{*}) = S(Z)$ 

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• Rejection sampling, Markov Chain Monte Carlo (MCMC)

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• Monte Carlo approximation: Generate  $M$  feasible samples  $(z_j^*)_{j=1}^M$ , i.e.  $S(z_j^*) = S(Z)$ , and compute *M*  $J_{j=1}^{M}$ , i.e.  $S(z_{j}^{*}) = S(Z)$ 

$$
P_{sel} = \mathbb{P}(T(Z^*, W) \le T
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- Confidence intervals:
	- test  $Y_i(1) Y_i(0) = \tau$  for different  $\tau$
	- (1 − *α*) confidence interval:  $C_{1-\alpha} = \{ \tau : P_{\text{sel}}(\tau) \ge \alpha \}$

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- Data carving: non-adaptive hold-out units

$$
P_{sel} = \mathbb{P}(T(Z^*, W) \le T
$$

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- 2 stages, 2 treatments  $Z_i \in \{0,1\}$ , 2 groups  $X_i \in \{$ low, high $\}$
- Potential outcomes:  $Y_i(0) = Y_i(1) \sim N(0,1)$  i.i.d.
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- First stage: 100 patients, Second stage: 40 patients
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- Selection variable:

$$
S = \begin{cases} \text{only low,} & \Delta < \Phi^{-1}(0.2), \\ \text{only high,} & \Delta > \Phi^{-1}(0.8), \\ \text{both,} & \text{otherwise,} \end{cases}
$$

recruit 40 from group  $X_i =$  low recruit 40 from group  $X_i =$  high recruit 20 from each group

















- **RT 2nd: Wild but has low power** RT 2nd **RT**
- Selective RT: **valid and more powerful**.
- Rejection sampling and MCMC lead to very similar approximations. R<br>O.C.<br>C.C. rejection probability

#### • RT: **no type-I error control** MCMC RS





 $AC$ 

nd?

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• Type-I error control in every subgroup



 $AC$ 

nd?

- Type-I error control in every subgroup
- Gain in power when there is a lot of "randomness left"



## **Conclusion**

- Experiments with adaptive treatments, recruitment and null hypothesis
- Visualization via DAGs
- 
- Computability under general assumptions
- Approximation via rejection sampling or MCMC

#### **• Key idea: Conditioning randomization p-value on the selection information**

# Thanks for your attention!



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## **Hold-out Units**

