

Nonparametric-efficient Causal Mediation Analysis for Stochastic Interventions Nima Hejazi, Mark van der Laan, and Iván Díaz

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OVERVIEW & MOTIVATIONS

- We present a decomposition of the population intervention effect, defined through stochastic interventions.
- Population intervention effects provide a generalized framework in which a variety of interesting causal contrasts can be defined, including effects for continuous and categorical exposures.
- We propose estimators of these direct and indirect effects:
 - substitution (G-computation), weighted (IPW), and
 - efficient estimators based on flexible regression techniques (one-step, TMLE).

SOFTWARE IMPLEMENTATION

- Our efficient estimators are asymptotically linear under a condition requiring $n^{1/4}$ consistency of certain regression functions.
- The efficient estimators are asymptotically normal with estimable variance, thereby allowing for the construction of confidence intervals and hypothesis tests.
- The medshift R package [2] implements these estimators and leverages state-of-theart machine learning in the procedure.
- tlverse plug: https://tlverse.org

STOCHASTIC POPULATION INTERVENTION (IN) DIRECT EFFECTS

- Consider $O = (W, A, Z, Y) \sim P_0 \in \mathcal{M}$, where W is a set of baseline covariates, A an intervention, Z a mediator between A and outcome, and Y the outcome, with no assumptions on model \mathcal{M} .
- We may decompose the PIE in terms of a population intervention direct effect (PIDE) and a population intervention indirect effect (PIIE):

$$\psi(\delta) = \underbrace{\mathbb{E}\{Y(g,q) - Y(g_{\delta},q)\}}_{\text{PIDE}} + \underbrace{\mathbb{E}\{Y(g_{\delta},q) - Y(g_{\delta},q_{\delta})\}}_{\text{PIDE}}.$$
(2)

• The causal parameter is identified by the observed data parameter [?]:

$$\theta(\delta) = \int m(a, z, w) g_{\delta}(a \mid w) p(z, w) d\nu(a, z, w). \tag{3}$$

• Let $\eta = (g, m, e, \phi)$. The efficient influence function for $\theta(\delta)$ in the nonparametric model M is $D_{n,\delta}^{Y}(o) + D_{n,\delta}^{A}(o) + D_{n,\delta}^{Z,W}(o) - \theta(\delta)$, where

$$D_{\eta,\delta}^{Y}(o) = \frac{g_{\delta}(a \mid w)}{e(a \mid z, w)} \{ y - m(z, a, w) \}; D_{\eta,\delta}^{Z,W}(o) = \int m(z, a, w) g_{\delta}(a \mid w) d\kappa(a),$$

$$D_{\eta,\delta}^{A}(o) = \frac{g_{\delta}(a \mid w)}{g(a \mid w)} \left\{ \phi(a, w) - \int \phi(a, w) g_{\delta}(a \mid w) d\kappa(a) \right\}.$$

and
$$\phi_0(a, w) = \mathbb{E}\left\{\frac{g(A|W)}{e(A|Z,W)}m(Z, A, W) \mid A = a, W = w\right\}.$$

CONSTRUCTION OF NONPARAMETRIC-EFFICIENT ESTIMATORS

• To avoid entropy conditions on the initial estimators, we rely on cross-fitting [3, 1] — denote by $\hat{\eta}_i$ the estimator of $\eta = (g, m, e, \phi)$, obtained by training the corresponding prediction algorithm using only data in the sample \mathcal{T}_i . Further, let j(i) denote the index of the validation set which contains observation i. The estimator is thus defined as:

$$\hat{\theta}\delta = \frac{1}{n} \sum_{i=1}^{n} D_{\hat{\eta}_{j(i)}, \delta}(O_i) = \frac{1}{n} \sum_{i=1}^{n} \left\{ D_{\hat{\eta}_{j(i)}, \delta}^{Y}(O_i) + D_{\hat{\eta}_{j(i)}, \delta}^{A}(O_i) + D_{\hat{\eta}_{j(i)}, \delta}^{Z, W}(O_i) \right\}. \tag{1}$$

• TMLE!

RESULTS & DISCUSSION

- samples; however, inefficient TMLE with HAL has bias not converging at $n^{-\frac{1}{2}}$.
- All estimators approx. unbiased in large
 Fitting Π with HAL or GLM, efficient TMLE has lower variance than the inefficient.

REFERENCES

- [1] V. Chernozhukov, D. Chetverikov, M. Demirer, E. Duflo, C. Hansen et al., "Double machine learning for treatment and causal parameters," arXiv preprint arXiv:1608.00060, 2016.
- [2] N. S. Hejazi and I. Díaz, medshift: Causal mediation analysis for stochastic interventions in R, 2019 r package version 0.0.8. [Online]. Available: https://github.com/nhejazi/medshift
- [3] W. Zheng and M. J. van der Laan, "Cross-validated targeted minimum-loss-based estimation," in Targeted Learning. Springer, 2011, pp. 459–474

BUT WAIT, THERE'S MORE!

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- Pre-print of our original paper: https://