

ESTIMATION BAYÉSIENNE CIBLÉE DE MODÈLES STRUCTURELS MARGINAUX

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Introduction

- Slides: herbsusmann.com/jmb2021.
- Causal analysis is concerned with estimating *treatment effects*: the effect of an intervention on an outcome.
- We may also be interested in how individual characteristics affect the size of the treatment effect.
- Can we estimate how a set of *treatment effect modifiers* influence the size of the treatment effect?
- Our contribution: asymptotically efficient non-parametric Bayesian estimator

Cheatsheet

- Data structure
 - $W \in \mathbb{R}^d$: vector of covariates
 - $A \in \{0, 1\}$: binary indicator of treatment
 - $Y \in \mathbb{R}$: outcome
 - Observed data: $O = (W, A, Y) \sim P_0$

- Conditional mean:

$$\bar{Q}_P(a, w) = \mathbb{E}_P[Y \mid A = a, W = w]$$

- Conditional treatment effect:

$$\tilde{Q}_P(w) = \bar{Q}_P(1, w) - \bar{Q}_P(0, w)$$

- Treatment effect modifiers
 - Let $V = (V_1, \dots, V_k) \subset W$.
 - Let $X = (1, V_1, \dots, V_k)$. ($X \in \mathbb{R}^p$, with $p = k + 1$.)
- Marginal structural model:

$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - X^\top \beta \right)^2 \right]$$

Data Structure

Data structure:

- $W \in \mathbb{R}^d$: vector of covariates
- $A \in \{0, 1\}$: binary indicator of treatment
- $Y \in \mathbb{R}$: outcome

Observed data:

- We assume that the observed data are n i.i.d. draws of $O = (W, A, Y)$ from a distribution P_0 .

Average Treatment Effect

- Suppose we are interested in summarizing the effect of the treatment on the outcome over the entire population.
- Notation

$$\bar{Q}_P(a, w) = \mathbb{E}_P[Y \mid A = a, W = w]$$

$$\tilde{Q}_P(w) = \bar{Q}_P(1, w) - \bar{Q}_P(0, w)$$

- Average Treatment Effect (ATE):

$$\begin{aligned}\psi_P^{ATE} &= \mathbb{E}_P[\tilde{Q}_P(W)] \\ &= \mathbb{E}_P[\bar{Q}_P(1, W) - \bar{Q}_P(0, W)] \\ &= \mathbb{E}_P[\mathbb{E}_P[Y \mid A = 1, W] - \mathbb{E}_P[Y \mid A = 0, W]]\end{aligned}$$

Treatment Effect Modification

- What if we are interested in whether the size of the effect is influenced by the covariates?
- Naive approach: estimate the conditional treatment effect for every value of W :

$$\begin{aligned}\tilde{Q}_P(w) &= \bar{Q}_P(1, w) - \bar{Q}_P(0, w) \\ &= \mathbb{E}_P[Y \mid A = 1, W = w] - \mathbb{E}_P[Y \mid A = 0, W = w]\end{aligned}$$

- Two problems:
 - Could be difficult to estimate for every w , especially if there are strata with few observations.
 - How do we make sense of the results?

Marginal Structural Models

- Alternative approach: summarize the relationship between covariates and treatment effects via a low-dimensional *working model*.
- Let $V = (V_1, \dots, V_k) \subseteq W$ be a set of possible *treatment effect modifiers*.
- Let $X = (1, V_1, \dots, V_k)$.
 - $X \in \mathbb{R}^p$, where $p = k + 1$.
- Define $\mathbf{B}(P) \in \mathbb{R}^p$ as the solution to the optimization problem

$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - X^\top \beta \right)^2 \right]$$

Marginal Structural Models

$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\boxed{\tilde{Q}_P(W)} - X^\top \beta \right)^2 \right]$$

↓
treatment effect for W

Marginal Structural Models

$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - \boxed{X^\top \beta} \right)^2 \right]$$

linear working model

Marginal Structural Models

$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - X^\top \beta \right)^2 \right]$$

↓
squared-error risk

Marginal Structural Models

$$\boxed{\mathbf{B}(P)} = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - X^\top \beta \right)^2 \right]$$



defined in terms of P

Semi-parametric inference

- Define $\beta_0 \equiv \mathbf{B}(P_0)$
- Estimation: we assume only that P_0 falls within the non-parametric model \mathcal{M} .
- We wish to construct regular, asymptotically normal and efficient estimator of β_0 .
- We start with a frequentist estimator, which we develop a Bayesian version of.

Semi-parametric inference

- We can write any regular estimator of β_0 as:

$$\hat{\beta}_n = \beta_0 + \frac{1}{n} \sum_{i=1}^n IC_{P_0}(O_i) + o_p(n^{-1/2}),$$

where IC_{P_0} is called an *influence function* of the parameter β_0 .

- The influence function with the smallest variance is called the *efficient influence function* (EIF). The EIF of β_0 is given by:

$$D_P(O) = M^{-1} \left[\frac{2A - 1}{g_P(A, W)} (Y - \bar{Q}_P(A, W)) + \tilde{Q}_P(W) - \beta^\top X \right] X,$$

with $g_P(a, w) \equiv \mathbb{P}_P[A = a | W = w]$ and $M = \mathbb{E}_P[X^\top X]$.

- The semi-parametric efficiency bound for estimating β_0 is given by $\text{var}_{P_0}(D_{P_0}(O))$

Targeted Maximum Likelihood Estimation

- It turns out we can construct an estimator that achieves this efficiency bound!
- *Targeted Maximum Likelihood Estimation (TMLE)* (van der Laan and Rose, 2011, 2018)
- Plug-in estimator of the form

$$\beta^{TMLE} = \mathbf{B}(P_n^0(\epsilon))$$

where $\{P_n^0(\epsilon) : \epsilon \in \mathbb{R}^p\}$ is a *fluctuation* of an initial estimator P_n^0 of the pieces of P_0 relevant to β .

- The TMLE estimator is designed to solve the EIF of the target parameter: $\mathbb{E}_{P_n}[D_{P_n^*}(O)] = 0$.

Bayesian Targeted Maximum Likelihood Estimation

- Can we make this procedure Bayesian?
- Core insight: the fluctuation submodel $\{P_n^0(\epsilon) : \epsilon \in \mathbb{R}^p\}$ defines a proper likelihood. So we can use Bayesian inference to estimate ϵ ! (Diaz et al., 2011; Díaz et al., 2020)
- Basic application of Bayes rule: posterior distribution of ϵ is given by

$$\Pi_{\epsilon}(\epsilon \mid O_1, \dots, O_n) \propto \pi_{\epsilon}(\epsilon) \prod_{i=1}^n p_{\epsilon}(O_i \mid \epsilon)$$

Bayesian Targeted Maximum Likelihood Estimation

- In practice, we probably don't have any prior information for ϵ .
- Instead, we can put a prior on β , which we map to a prior on ϵ :

$$\pi_{\epsilon}(\epsilon) = \pi_{\beta}(\mathbf{B}(P_n^0(\epsilon))) |\det(J(\epsilon))|,$$

where J is the Jacobian of the transformation $\mathbf{B}(P_n^0(\epsilon))$.

Bayesian Targeted Maximum Likelihood Estimation

- Sampling techniques such as Markov-Chain Monte Carlo can be used to draw a set of samples from the posterior distribution of ϵ .
- Then, we can map each of these samples of ϵ to a β , yielding a set of samples from the posterior distribution of β .
- Credible intervals can be constructed using the empirical quantiles of the posterior samples.

Bayesian Targeted Maximum Likelihood Estimation

Theorem

Under certain conditions, the posterior distribution converges in total variation:

$$\sup_{\mathbb{B}^p} \left| \Pi_{\beta} \{ \sqrt{n}(\bar{\beta} - \beta) \in \mathbb{B}^p \mid O_n \} - N_{\sqrt{n}(\tilde{\beta} - \beta), P_n D_n^\top D_n}(\mathbb{B}^p) \right| \rightarrow 0$$

where \mathbb{B}^p is a subset of \mathbb{R}^p , $N_{\mu, \Sigma}$ is the multivariate normal distribution with mean vector μ and covariance matrix Σ , D_n is the targeted estimate of the EIF, $\bar{\beta} \sim \Pi_{\beta}$, and $\tilde{\beta}$ is the targeted MLE.

Simulation

- Data generating process P_0 :

$$W_1, W_2, W_3, V \sim N_4(0, \mathbf{I}_4)$$

$$A|W_1, W_2, W_3, V \sim \text{Bernoulli}(\text{logit}^{-1}(W_1 + W_2 + W_3))$$

$$Y|A, W_1, W_2, W_3, V \sim N(W_1 + W_2 + W_3 + 3A + 1.5AV, 0.1)$$

- True average treatment effect: $\Psi_{P_0}^{ATE} = 3$.
- $X = (1, V)^\top$.
- Optimization problem:

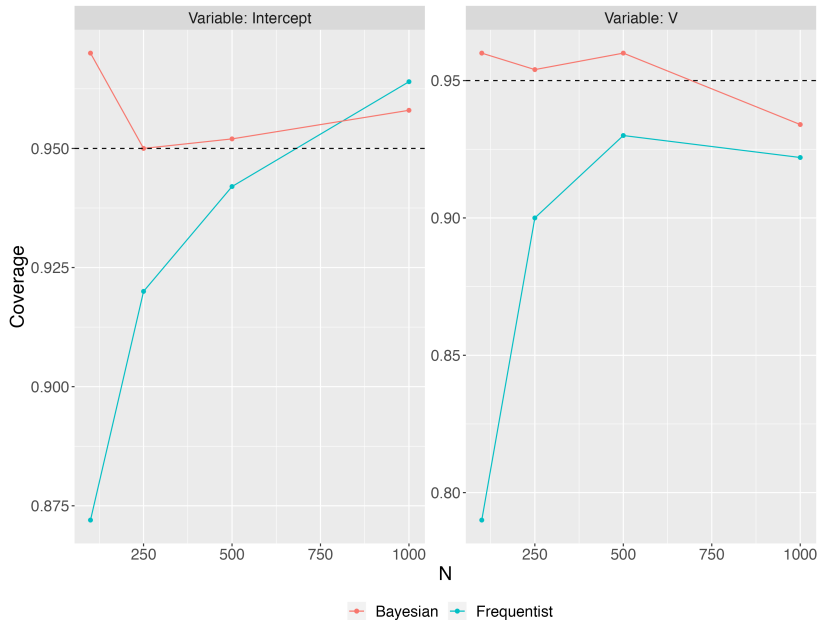
$$\mathbf{B}(P) = \arg \min_{\beta \in \mathbb{R}^p} \mathbb{E}_P \left[\left(\tilde{Q}_P(W) - X^\top \beta \right)^2 \right]$$

- Target parameter: $\beta_0 \equiv \mathbf{B}(P_0)$
- $\beta_0 \in \mathbb{R}^2$: intercept term and linear effect of V on treatment effect
- For this data generating process, $\beta_0 = (0, 1.5)^\top$.

Simulation

- Simulate 500 datasets for $n = (100, 250, 500, 1000)$.
- Estimate nuisance parameters g_P and \bar{Q}_P with correctly specified regressions.
- Flat priors on β in Bayesian TMLE.
- Frequentist TMLE implemented in *R*, Bayesian TMLE implemented in *R* and *Stan*.
- Julia implementation with automatic differentiation

Simulation



Conclusions

- Marginal Structural Models provide a method for summarizing impact of *treatment effect modifiers*.
- TMLE provides a framework for efficient estimation of Marginal Structural Model parameters within a semi-parametric model.
- Bayesian TMLE brings benefits of both frequentist and Bayesian approaches
- Ongoing work: does the Bayesian estimator inherit other useful properties of the frequentist TMLE? Does incorporation of prior information improve finite-sample performance?

References

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