

## BACKGROUND

- Bayesian causal methods that follow a parametric specification of the joint likelihood of treatment, outcome and covariates, are analytically intractable when face with high-dimensional confounders.
- One possible approach for dimensionality reduction is to model the set of confounders as class indicators in a latent class analysis.
- This approach mimics the treatment assignment process often seen in observational studies with administrative data that contain a large number of variables which are indicative of the patient's disease and health status.
- We aim to provide a Bayesian latent class approach to estimate causal effects.**

## OBJECTIVES

- In this paper, we consider **a causal effect that is confounded by an unobserved, visit specific, latent class** in a longitudinal setting.
- We formulate the joint likelihood of the treatment, outcome and latent class models conditionally on the class indicators, which permits a **full Bayesian causal inference**.

## CAUSAL FRAMEWORK

### 1. NOTATIONS

- $n$  subjects indexed by  $i, i = 1, \dots, n$  and  $k$  number of visits for each subject indexed by  $j, j = 1, \dots, k$ .
- $Y_i, X_{ij}, U_{ij}$  and  $Z_{ij}$  are random variables representing an end-of-study outcome, time-dependent class indicators (a vector of  $p$  elements), time-dependent latent class and time-dependent treatment for  $i$  at visit  $j$ .
- There are  $a$  treatment categories available at each visit and  $c$  number of class memberships at each visit.
- History up to visit  $j$  are denoted as  $\tilde{U}_{ij} = \{U_{i1}, \dots, U_{ij}\}$ ,  $\tilde{X}_{ij} = \{X_{i1}, \dots, X_{ij}\}$  and  $\tilde{Z}_{ij} = \{Z_{i1}, \dots, Z_{ij}\}$ .
- Let  $\theta, \alpha, \beta$  and  $\gamma$  characterize the outcome model, the latent class model, the class indicator model and the treatment model respectively.

### 2. DAG

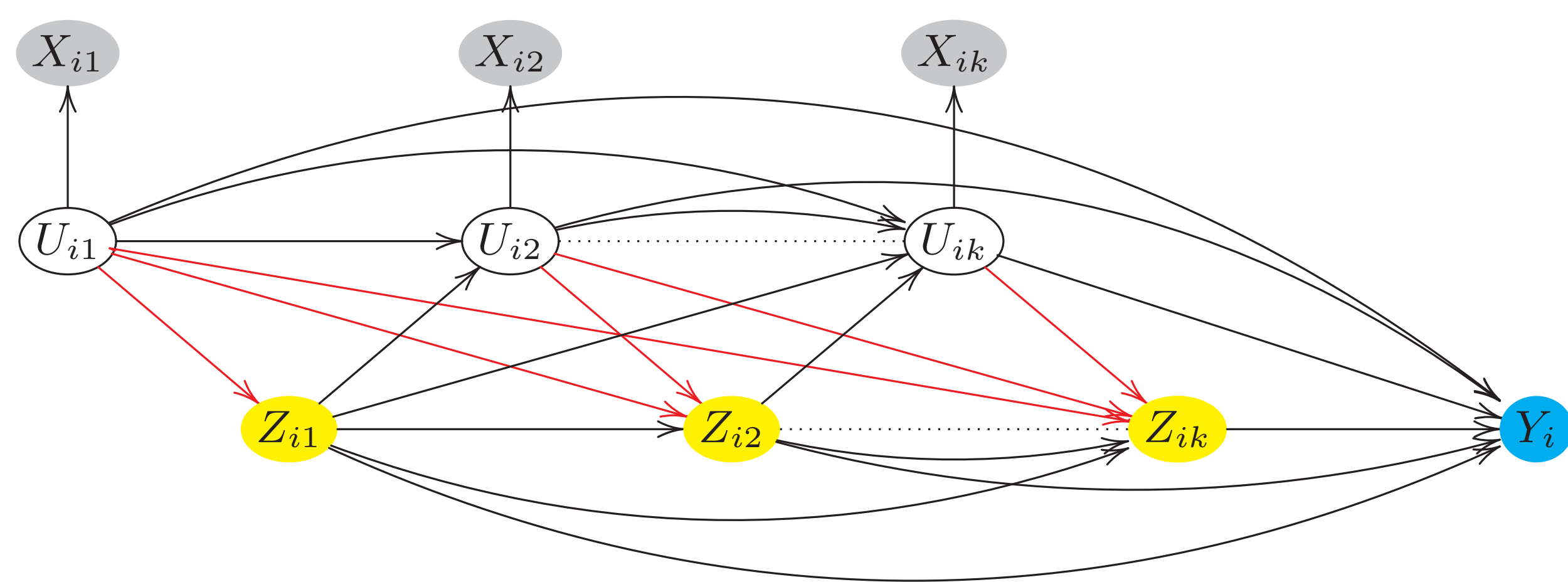


Figure 1. Longitudinal causal DAG between latent class, indicators, treatment and outcome

### 3. ASSUMPTIONS

- Stable unit treatment value assumption
- Consistency,  $Y_i^a | (z_{i1} = a_1, \dots, z_{ik} = a_k) = Y_i | (z_{i1} = a_1, \dots, z_{ik} = a_k)$
- Positivity, at each visit, every possible treatment sequence which is compatible with treatment history up till that visit have positive probability of occurring.
- Sequential latent unconfoundedness**,  $Z_{ij} \perp (U_{ij}^a, X_{ij}^a, Y_i^a) | (U_{ij-1}^a, Z_{ij-1})$ , for  $j = 1, \dots, k$ .
- Conditional independence between class indicators**,  $P(X_{ij} | U_{ij}) = \prod_{l=1}^p P(X_{lij} | U_{ij})$ .

## SIMULATION STUDY

### 1. SIMULATED DATASET

- We simulated 1000 iterations of a simple two-visit ( $k = 2$ ) longitudinal dataset with  $n = 250$  and  $n = 500$ .
- $C_i^a \sim N(10, 3)$  and  $C_i^s$  from a Bernoulli distribution with  $P(C_i^s = 1) = 0.6$
- $Z_{ij}$ , a binary treatment assignment.  $\text{logit}(P(Z_{ij} = 1)) = -1 + u_{ij} - z_{ij-1}$
- $U_{ij}$ , three class memberships with simulated proportion at 42% : 29% : 29% from a Multinomial distribution with
 
$$\log \frac{P(U_{ij} = 2)}{P(U_{ij} = 1)} = 0.5 - 0.1C_i^a + 0.2C_i^s + I(u_{ij-1} = 2) + 0.5I(u_{ij-1} = 3) - z_{ij-1}$$

$$\log \frac{P(U_{ij} = 3)}{P(U_{ij} = 1)} = 0.5 - 0.1C_i^a + 0.2C_i^s + 0.5I(u_{ij-1} = 2) + I(u_{ij-1} = 3) - z_{ij-1}$$
- $X_{ij}$ , simulated conditionally independent given  $U_{ij}$  from Bernoulli distribution.
  - High quality defined as  $P(X_{ij} = 1 | U_{ij} = c) = 0.88$
  - Medium quality defined as  $P(X_{ij} = 1 | U_{ij} = c) = 0.73$
  - Low quality defined as  $P(X_{ij} = 1 | U_{ij} = c) = 0.62$
- $Y_i \sim N(\mu_a, 1)$ , where  $\mu_a = 0.1 + 0.5I(z_{i1} = a_1) + I(z_{i2} = a_2) - 0.2I(u_{i1} = 2) - 0.5I(u_{i1} = 3) - 0.5I(u_{i2} = 2) - I(u_{i2} = 3)$ .

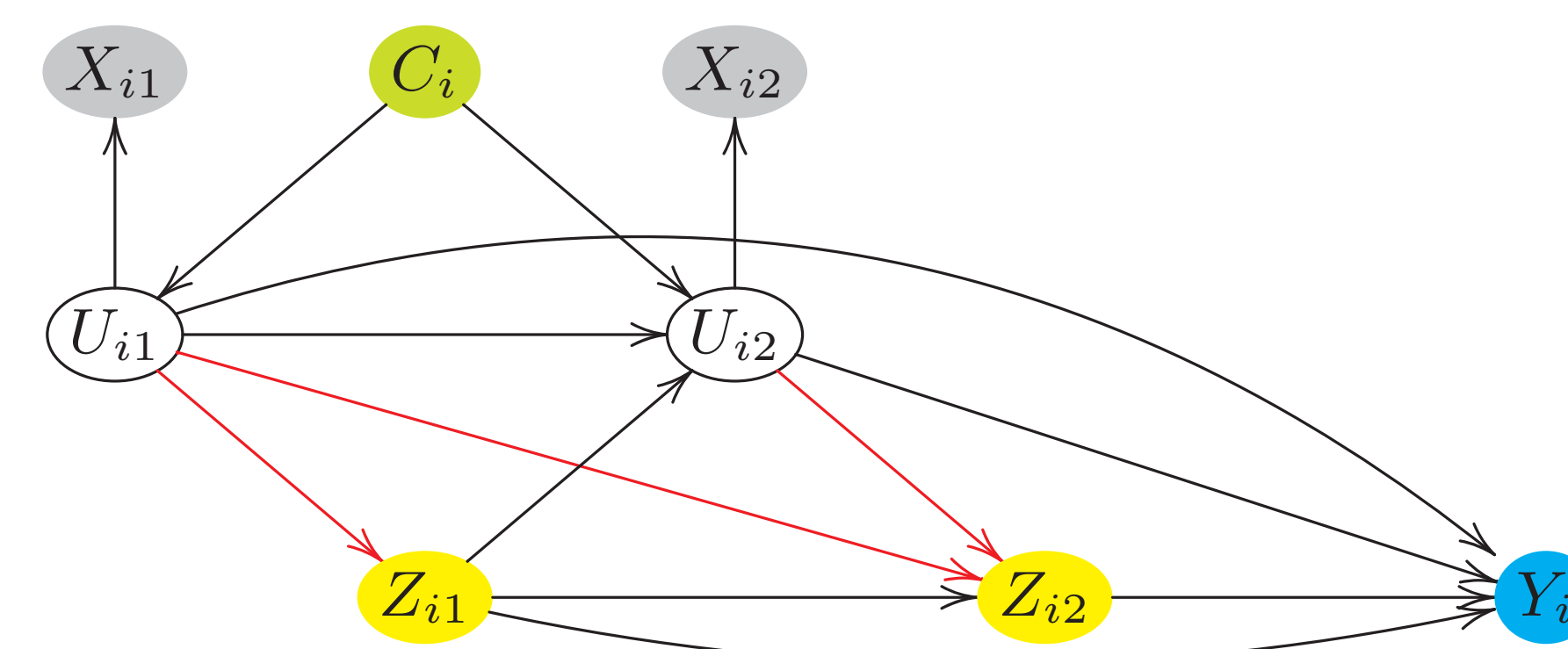


Figure 2. Causal diagram of the simulation dataset.

**Table 1. Simulation results from 1000 iterations.** Parameter of interest ( $\mu_{11} - \mu_{00}$ ), treatment effect evaluated between always treated and never treated. RB: Relative Bias; ESE: Empirical Standard Error; ASE: Average Standard Error; CP: Coverage Probability; U1 and U2 represent the average proportion of correct imputation under Bayesian estimation.

Setting	Estimator	Mean	RB	ESE	ASE	CP	U1	U2
$n = 250$	Naive	1.18	-29.38	0.24	0.24	46.1		
10 indicators	Adjust	1.47	-12.21	0.22	0.23	86.4		
high quality	MSMs Sand.	1.61	-3.47	0.27	0.28	95.8		
	Bayes	1.67	0.12	0.22	0.23	96.4	0.70	0.70
$n = 250$	Naive	1.18	-29.38	0.24	0.24	46.1		
10 indicators	Adjust	1.38	-17.07	0.24	0.24	75.0		
med quality	MSMs Sand.	1.48	-11.22	0.28	0.28	87.5		
	Bayes	1.68	0.80	0.26	0.25	95.9	0.60	0.60
$n = 250$	Naive	1.18	-29.38	0.24	0.24	46.1		
10 indicators	Adjust	1.28	-23.48	0.24	0.24	61.2		
low quality	MSMs Sand.	1.32	-20.99	0.26	0.26	71.5		
	Bayes	1.19	-28.76	0.25	0.24	47.6	0.39	0.39
$n = 500$	Naive	1.18	-29.59	0.16	0.17	13.6		
10 indicators	Adjust	1.47	-12.14	0.15	0.16	78.4		
high quality	MSMs Sand.	1.62	-3.09	0.18	0.20	95.2		
	Bayes	1.68	0.45	0.15	0.16	96.6	0.78	0.78
$n = 500$	Naive	1.18	-29.59	0.16	0.17	13.6		
10 indicators	Adjust	1.38	-17.68	0.15	0.16	54.4		
med quality	MSMs Sand.	1.46	-12.31	0.17	0.19	82.2		
	Bayes	1.68	0.70	0.17	0.17	96.4	0.68	0.69
$n = 500$	Naive	1.18	-29.59	0.16	0.17	13.6		
10 indicators	Adjust	1.27	-24.12	0.16	0.17	29.8		
low quality	MSMs Sand.	1.30	-22.33	0.17	0.18	41.8		
	Bayes	1.31	-21.73	0.28	0.18	35.2	0.40	0.41

## BAYESIAN CAUSAL INFERENCE WITH LATENT CLASS

### 1. Causal parameter of interest - Average potential outcome (APO)

$$E[Y_i^a] = \sum_{u_{ik}=1}^C \dots \sum_{u_{i1}=1}^C E(y_i | \tilde{z}_i = \tilde{a}, \tilde{u}_i = \tilde{c}_k, \theta) \left[ \prod_{j=1}^k P(u_{ij} = c_j | \tilde{z}_{ij-1} = \tilde{a}_{j-1}, \tilde{u}_{ij-1} = \tilde{c}_{j-1}, \alpha_j) \right] \quad (1)$$

### 2. Joint likelihood

$$P(y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i | \theta, \alpha, \beta, \gamma) = P(y_i | \tilde{z}_i, \tilde{u}_i = \tilde{c}_k, \theta) \left[ \prod_{j=1}^k P(z_{ij} | \tilde{z}_{ij-1}, \tilde{u}_{ij} = \tilde{c}_j, \gamma_j) \times P(u_{ij} = c_j | \tilde{u}_{ij-1}, \tilde{z}_{ij-1}, \alpha_j) P(x_{ij} | u_{ij} = c_j, \beta_j) \right] \quad (2)$$

### 3. Posterior Distribution

$$P(\theta, \alpha, \beta, \gamma | y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i) = \frac{P(y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i | \theta, \alpha, \beta, \gamma) P_0(\theta, \alpha, \beta, \gamma)}{\int_{\{\theta, \alpha, \beta, \gamma\}} P(y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i | \theta, \alpha, \beta, \gamma) P_0(\theta, \alpha, \beta, \gamma) d\theta d\alpha d\beta d\gamma} \quad (3)$$

We assume a prior independence  $P_0(\theta, \alpha, \beta, \gamma) = P_0(\theta)P_0(\alpha)P_0(\beta)P_0(\gamma)$ .

### 4. Posterior Predictive Inference on APO

We predict the potential outcome of a given treatment sequence for a new observation drawn from data distribution  $\{Y, Z, U, X\}$ ,

$$P(\tilde{y}_i^a | y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i) = \int_{\{\theta, \alpha, \beta, \gamma\}} P(\tilde{y}_i^a | y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i) P(\theta, \alpha, \beta, \gamma | y_i, \tilde{z}_i, \tilde{u}_i, \tilde{x}_i) d\theta d\alpha d\beta d\gamma. \quad (4)$$

## CONCLUSION & FUTURE WORKS

- Based on the simulation study, Bayesian latent class approach is preferred when we have medium to high quality class indicators.**
- Full Bayesian specification permits imputation on class membership; Even when memberships are not well predicted, we can still obtain relative unbiased APO estimate.
- Our Bayesian approach can be easily implemented in common MCMC software.
- Future works to investigate the trade off between estimation accuracy and minimum number of high quality indicators.**

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