

# Semi-supervised D-Learning for Optimal Individual Treatment Regimes

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# Outline

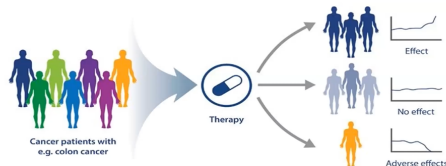
- 1 Introduction
- 2 Methodology
- 3 Asymptotic properties
- 4 Numerical Simulations
- 5 Real Data Analysis: MIMIC-IV
- 6 Conclusions

# Precision Medicine

- **Heterogeneity**: different patients respond differently to the same treatment.
  - ▶ Positive treatment effects;
  - ▶ Side effects.
- **One-size-fits-all** → **Precision Medicine**
- **Advantages**:
  - ▶ Improve patient adherence;
  - ▶ Reduce unnecessary treatments and side effects;
  - ▶ Promote recovery;
  - ▶ Enhance quality of care and quality of life;
  - ▶ Optimize allocation of medical resources;
  - ▶ Lower overall healthcare costs;
  - ▶ .....

## Current Medicine

One Treatment Fits All



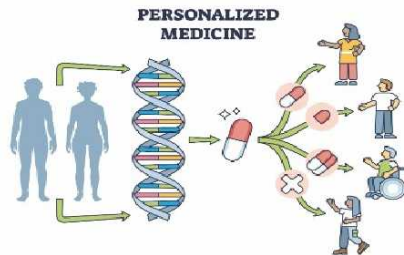
## Future Medicine

More Personalized Diagnostics



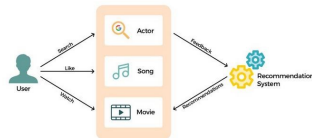
# Precision Medicine - Personalized Decision-Making

- **Goal:** Find the optimal mapping from individual characteristics  $\mathbf{X} \in \mathcal{X}$  to treatments  $A \in \mathcal{A}$ , i.e.  $d^{opt}(\mathbf{X})$ , to maximize the expected clinical outcome  $E[Y^*(d(\mathbf{X}))]$ .
  - ▶ **X:** demographics, clinical features, genetic information, environmental factors, etc.;
  - ▶ **A:** drug choice, dosage, surgery, specific dietary or exercise recommendations, etc.;
  - ▶ **Y:** biomarker levels, survival time, disease progression or remission status, quality of life scores, etc.
- **Applications:**
  - ▶ **Disease management:** Recommend the optimal drug dosage based on patient characteristics to optimize treatment efficacy;
  - ▶ **Smart health monitoring:** Use wearable devices and biosensors for personalized health management;
  - ▶ **Personalized medical intervention:** Combine multimodal data to predict disease risk and enable early intervention.



# Personalized Decision-Making - Beyond Precision Medicine

- **Computer Science:** context-aware recommender systems that improve accuracy by incorporating time, location, and social context.



- **Finance:** provide personalized investment advice and wealth management plans based on consumption habits and risk preferences.

- **Public Management:** improve the overall effectiveness of policies through personalized interventions targeting individuals with high social connectivity.



# Traditional Methods

- **Q-learning** (Qian and Murphy, 2011; Watkins, 1989; Watkins and Dayan, 1992)

Define the Q-function:  $Q(\mathbf{x}, a) := E[Y|\mathbf{X} = \mathbf{x}, A = a]$ , and specify a model  $Q(\mathbf{X}, A; \beta)$ .

$$\hat{d}^{opt}(\mathbf{X}) = \arg \max_{a \in \mathcal{A}} Q(\mathbf{X}, a; \hat{\beta}),$$

where  $\hat{\beta} = \arg \min_{\beta} \frac{1}{n} \sum_{i=1}^n (Y_i - Q(\mathbf{X}_i, A_i; \beta))^2$ .

- **A-learning** (Blatt et al., 2004; Murphy, 2003; Robins, 2004)

Define the contrast function:  $C(\mathbf{X}) = Q(\mathbf{X}, 1) - Q(\mathbf{X}, 0)$ , then  $d^{opt}(\mathbf{X}) = I(C(\mathbf{X}) \geq 0)$ .

**Doubly robust A-learning:** Let  $\nu(\mathbf{X}) = E[Y|\mathbf{X}]$  and  $\pi(\mathbf{X}) = E[A|\mathbf{X}]$ , with corresponding estimators  $\hat{\nu}(\mathbf{X})$  and  $\hat{\pi}(\mathbf{X})$ . Specify a model for the contrast function  $C(\mathbf{X}; \theta)$ , then

$$\hat{\theta} = \arg \min_{\theta} \frac{1}{n} \sum_{i=1}^n \{Y_i - \hat{\nu}(\mathbf{X}_i) - [A_i - \hat{\pi}(\mathbf{X}_i)]C(\mathbf{X}_i; \theta)\}^2.$$

# Traditional Methods

- **Direct search methods** (Chu et al., 2023; Zhang et al., 2012)

Denote value function  $V(d(\mathbf{X})) := E[Y(d(\mathbf{X}))]$ , then  $d^{\text{opt}}(\mathbf{X}) = \arg \max_{d(\mathbf{X}) \in \mathcal{D}} V(d(\mathbf{X}))$ .

- ▶ IPW-based estimator:  $\hat{V}_{IPW}(d(\mathbf{X})) = P_n \left[ \frac{I(A=d(\mathbf{X}))}{\hat{\pi}(\mathbf{X}, A)} Y \right]$ .
- ▶ AIPW-based estimator:  $\hat{V}_{AIPW}(d(\mathbf{X})) = P_n \left[ \frac{I(A=d(\mathbf{X}))}{\hat{\pi}(\mathbf{X}, A)} Y - \frac{I(A=d(\mathbf{X})) - \hat{\pi}(\mathbf{X}, A)}{\hat{\pi}(\mathbf{X}, A)} \hat{Q}(\mathbf{X}, d(\mathbf{X})) \right]$ , where  $Q(\mathbf{X}, d(\mathbf{X})) = Q(\mathbf{X}, 1)I(d(\mathbf{X}) = 1) + Q(\mathbf{X}, 0)I(d(\mathbf{X}) = 0)$ .

- **D-learning** (Qi et al., 2020; Qi and Liu, 2018; Shah et al., 2023)

$$d^{\text{opt}}(\mathbf{X}) = \text{sign}\{E[Y|\mathbf{X}, A = 1] - E[Y|\mathbf{X}, A = -1]\} = \text{sign} \left\{ E \left[ \frac{AY}{\pi(A, \mathbf{X})} \middle| \mathbf{X} \right] \right\} := \text{sign}\{f^{\text{opt}}(\mathbf{X})\}.$$

$$f^{\text{opt}}(\mathbf{X}) \in \arg \min_{f(\mathbf{X}) \in \mathcal{F}} E \left[ \frac{1}{\pi(\mathbf{X}, A)} (2AY - f(\mathbf{X}))^2 \right].$$

# Challenges

In the literature, **linear decision classes** are particularly favored by researchers due to their **simple structure** and **good interpretability** (Chu et al., 2023; Fan et al., 2017; Li et al., 2025).

- **Misspecification** → Suboptimal decisions (Maronge et al., 2023; Zhao et al., 2012).
  - ▶ Song et al. (2017) proposed a novel method to estimate optimal ITR under a semiparametric additive single-index model and the link function was estimated by B-spline. But it suffers from model mis-specification.
  - ▶ Qi and Liu (2018) handled the nonlinear ITR by kernel-based and machine learning methods.
- **Scarcity of labeled data** → Underutilization of large amounts of unlabeled data.



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- **Scarcity of labeled data** → Underutilization of large amounts of unlabeled data.

**Semi-supervised learning:** Leverage information from both labeled and unlabeled data to enhance estimation efficiency and robustness.

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- 1 Introduction
- 2 Methodology**
- 3 Asymptotic properties
- 4 Numerical Simulations
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# Methodology

## Notations $(\mathbf{X}, A, Y)$

$\mathbf{X} = (1, X_1, \dots, X_p) \in \mathcal{X} \subseteq \mathbb{R}^{p+1}$ :  $p$ -dimensional **covariates**  $\mathbf{X}^- = (X_1, \dots, X_p)$  including the interception term, with bounded support  $\mathcal{X}$ , and positive definite variance  $\text{Var}(\mathbf{X}^-)$ ;

$A \in \mathcal{A} = \{-1, 1\}$ : the binary **treatment** indicator;

$Y \in \mathcal{Y} \subseteq \mathbb{R}$ : the **outcome** variable, larger values are better.

## Observations $\mathcal{L} \cup \mathcal{U}$

$\mathcal{L} = \{(\mathbf{X}_i, A_i, Y_i) : i = 1, 2, \dots, n\}$ :  $n$  iid **labeled** observations;

$\mathcal{U} = \{\mathbf{X}_i : i = n + 1, n + 2, \dots, n + N\}$ :  $N$  iid **unlabeled** observations.

# Methodology

## Semi-supervised assumptions

- a.  $\mathcal{L} \perp \mathcal{U}$ ;
- b. Observations in  $\mathcal{L}$  and  $\mathcal{U}$  potentially follow the same distribution;
- c.  $\frac{n}{N} \rightarrow 0$  as  $n, N \rightarrow \infty$ .

## Identifiability Assumptions

Let  $Y^*(a)$  be the potential outcome.

- a. SUTVA:  $Y = \sum_{a \in \mathcal{A}} Y^*(a) I(A = a)$ ;
- b. Ignoreability:  $A \perp \{Y^*(-1), Y^*(1)\} \mid \mathbf{X}$ ;
- c. Positivity:  $0 < P(A = a | \mathbf{X} = \mathbf{x}) < 1$ .

# Methodology

**Optimal ITR:**  $d^{\text{opt}}(\mathbf{X}) = \arg \max_d V(d) = \arg \max_d E \left[ \frac{YI(A=d(\mathbf{X}))}{\pi(A, \mathbf{X})} \right].$

**D-learning - Equivalent representation of optimal ITR**

$$d^{\text{opt}}(\mathbf{X}) = \text{sign}\{E[Y|\mathbf{X}, A = 1] - E[Y|\mathbf{X}, A = -1]\} = \text{sign} \left\{ E \left[ \frac{AY}{\pi(A, \mathbf{X})} \middle| \mathbf{X} \right] \right\} := \text{sign}\{f^{\text{opt}}(\mathbf{X})\}.$$

**Structural Nested Mean Model (SNMM)**

$$Y = \mu_0(\mathbf{X}) + A\delta(\mathbf{X}) + e, E[e] = 0 \implies f^{\text{opt}}(\mathbf{X}) = 2\delta(\mathbf{X}).$$

**Linear decision function class:**  $f \in \mathcal{F} = \{f(\mathbf{X}) = \mathbf{X}^T \beta : \beta \in \mathbb{R}^{p+1}\}.$

**Supervised estimation:**  $\hat{\beta} = \arg \min_{\beta} P_n \left\{ \frac{AY}{\pi(A, \mathbf{X})} - \mathbf{X}^T \beta \right\}^2.$

Assume  $\pi(1, \mathbf{X}) = 0.5$  in a RCT setting. Define  $\beta_0$  as the solution to the equation  $E[\mathbf{X}(2AY - \mathbf{X}^T \beta)] = 0$ . The supervised estimator is obtained by solving

$$P_n \mathbf{X}(2AY - \mathbf{X}^T \beta) = 0.$$

## Misspecification of linear decision classes $\implies$ Semi-supervised D-learning (SSDL)

SSDL estimator based on fully nonparametric imputation:  $\hat{\beta}_{np}$

Let  $m(\mathbf{X}) = E[2AY|\mathbf{X}] = E[2AY|\mathbf{X}^-]$ , and its corresponding kernel estimator is

$$\hat{m}(\mathbf{X}_j) = \frac{(nh^p)^{-1} \sum_{i=1}^n H_h(\mathbf{X}_i^-, \mathbf{X}_j^-) \times 2A_i Y_i}{(nh^p)^{-1} \sum_{i=1}^n H_h(\mathbf{X}_i^-, \mathbf{X}_j^-)},$$

where  $H_h(u, v) = H(\frac{u-v}{h})$  with kernel function  $H: \mathbb{R}^p \rightarrow \mathbb{R}$  and bandwidth  $h$ . Then  $\hat{\beta}_{np}$  be obtained by the solution of

$$\frac{1}{N} \sum_{j=n+1}^{n+N} \mathbf{X}_j (\hat{m}(\mathbf{X}_j) - \mathbf{X}_j^T \beta) = 0.$$

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$$\frac{1}{N} \sum_{j=n+1}^{n+N} \mathbf{X}_j (\hat{m}(\mathbf{X}_j) - \mathbf{X}_j^T \beta) = 0.$$

## Curse of dimensionality $\implies$ Projection (dimensionality reduction) + Refitting (debias)

## SSDL estimator based on semiparametric imputation: $\hat{\beta}_{sp}$

**Projection:** Define  $m(\mathbf{X}^T \beta) = E[2AY | \mathbf{X}^T \beta]$ , and its corresponding kernel estimator is

$$\hat{m}(\mathbf{X}_j^T \beta) = \frac{(nh)^{-1} \sum_{i=1}^n K_h(\mathbf{X}_i^T \beta, \mathbf{X}_j^T \beta) * 2A_i Y_i}{(nh)^{-1} \sum_{i=1}^n K_h(\mathbf{X}_i^T \beta, \mathbf{X}_j^T \beta)},$$

where  $K_h(u, v) = K(\frac{u-v}{h})$  with kernel function  $K : \mathbb{R} \rightarrow \mathbb{R}$  and bandwidth  $h$ .

**Refitting:** Define  $\theta_0$  as the solution of  $E[\mathbf{X}(2AY - m(\mathbf{X}^T \beta_0) - \mathbf{X}^T \theta)] = 0$ , and  $\hat{\theta}$  is estimated by the estimating equation that

$$P_n \mathbf{X}(2AY - \hat{m}(\mathbf{X}^T \hat{\beta}) - \mathbf{X}^T \theta) = 0.$$

Denote the imputation function after refitting as  $\nu(\mathbf{X}; \beta, \theta) = m(\mathbf{X}^T \beta) + \mathbf{X}^T \theta$ , and its semiparametric (SP) estimator is

$$\hat{\nu}(\mathbf{X}; \hat{\beta}, \hat{\theta}) = \hat{m}(\mathbf{X}^T \hat{\beta}) + \mathbf{X}^T \hat{\theta}.$$

Then  $\hat{\beta}_{sp}$  can be obtained by solving

$$\frac{1}{N} \sum_{j=n+1}^{n+N} \mathbf{X}_j (\hat{\nu}(\mathbf{X}_j; \hat{\beta}, \hat{\theta}) - \mathbf{X}_j^T \beta) = 0.$$



## Overfitting? $\implies$ $\mathbb{K}$ -fold cross-validation (CV)

SSDL estimator based on semiparametric imputation with  $\mathbb{K}$ -fold CV:  $\hat{\beta}_{sp,\mathbb{K}}$

Let  $\mathcal{L}_k$  be the  $k$ -th random disjoint partition of  $\mathcal{L}$  with sample size  $n_{\mathbb{K}} = \frac{n}{\mathbb{K}}$  and index set  $\mathcal{I}_k$  for  $k \in \{1, \dots, \mathbb{K}\}$ . Let the set excluding the  $k$ -th partition be  $\mathcal{L}_k^- = \mathcal{L} - \mathcal{L}_k$  with sample size  $n_{\mathbb{K}}^- = n - n_{\mathbb{K}}$  and index set  $\mathcal{I}_k^-$ .

Denote the OLS and nonparametric imputation estimator under dimension reduction based on  $\mathcal{L}_k^-$  as  $\hat{\beta}_k$  and  $\hat{m}_k(\mathbf{X}^T \beta)$  respectively. Then  $\hat{\theta}_{\mathbb{K}}$  is obtained by solving

$$\frac{1}{n} \sum_{k=1}^{\mathbb{K}} \sum_{i \in \mathcal{I}_k} \mathbf{X}_i (2A_i Y_i - \hat{m}_k(\mathbf{X}_i^T \hat{\beta}_k) - \mathbf{X}_i^T \theta) = 0,$$

and the semiparametric imputation function estimation is  $\hat{\nu}(\mathbf{X}; \hat{\beta}_k, \hat{\theta}_{\mathbb{K}}) = \frac{1}{\mathbb{K}} \sum_{k=1}^{\mathbb{K}} \hat{m}_k(\mathbf{X}^T \hat{\beta}_k) + \mathbf{X}^T \hat{\theta}_{\mathbb{K}}$ . Then  $\hat{\beta}_{sp,\mathbb{K}}$  can be obtained by solving

$$\frac{1}{N} \sum_{j=n+1}^{n+N} \mathbf{X}_j (\hat{\nu}(\mathbf{X}_j; \hat{\beta}_k, \hat{\theta}_{\mathbb{K}}) - \mathbf{X}_j^T \beta) = 0.$$

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# Asymptotic properties

## Theorem 1

Under certain regularity conditions, we have

$$n^{\frac{1}{2}}(\hat{\beta}_{sp} - \beta_0) = \frac{1}{\sqrt{n}} \sum_{i=1}^n \varphi(\mathbf{Z}_i) + O_p(r_{n,N}),$$

where the influence function  $\varphi(\mathbf{Z}_i) = E[\mathbf{X}\mathbf{X}^T]^{-1}\{\mathbf{X}_i[2A_iY_i - \nu(\mathbf{X}_i; \beta_0, \boldsymbol{\theta}_0)]\}$  and  $r_{n,N} = O_p\left(\frac{n}{N}\right)^{\frac{1}{2}} + O_p(b_n)$  with  $b_n = n^{-\frac{2q-3}{2(2q+1)}}$ .

Thus  $n^{\frac{1}{2}}(\hat{\beta}_{sp} - \beta_0) \xrightarrow{d} N_{p+1}(\mathbf{0}, \Sigma)$  with positive definite  $(p+1) \times (p+1)$  matrix  $\Sigma = E[\mathbf{X}\mathbf{X}^T]^{-1}E\{\mathbf{X}\mathbf{X}^T[2AY - \nu(\mathbf{X}; \beta_0, \boldsymbol{\theta}_0)]^2\}E[\mathbf{X}\mathbf{X}^T]^{-1}$ .

## Theorem 2

When the fold of CV is fixed and satisfies  $\mathbb{K} \geq 2$ , under the conditions same as Theorem 1, we have

$$n^{\frac{1}{2}}(\hat{\beta}_{sp,\mathbb{K}} - \beta_0) = \frac{1}{\sqrt{n}} \sum_{i=1}^n \varphi(\mathbf{Z}_i) + O_p(\gamma_{n,N}),$$

where  $\gamma_{n,N} = O_p\left(\frac{n}{N}\right)^{\frac{1}{2}} + O_p(a_{n_{\mathbb{K}-}})$  with  $a_n = n^{-\frac{q}{2q+1}}\sqrt{\log n}$ . Thus  $n^{\frac{1}{2}}(\hat{\beta}_{sp,\mathbb{K}} - \beta_0) \xrightarrow{d} N_{p+1}(\mathbf{0}, \Sigma)$ .

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# Numerical Simulations

**Data generation:**  $Y = \mu_0(\mathbf{X}) + A\delta(\mathbf{X}) + e$ ;  $P(A = 1) = P(A = -1) = 0.5$ ,  $e \sim N(0, 0.5^2)$ ,  
 $\mathbf{X} = (1, X_1, X_2, \dots, X_p)'$ :  $X_i$  ( $i = 1, \dots, p$ )  $\sim U(-5, 5)$ .

## Model setting:

- Linear (Lin):  $\delta(\mathbf{X}) = 20\mathbf{X}^T\alpha$ ,
- Nonlinear 1 (NL1):  $\delta(\mathbf{X}) = 0.2(\mathbf{X}^T\alpha)^3$ ,
- Nonlinear 2 (NL2):  $\delta(\mathbf{X}) = \mathbf{X}^T\alpha + 0.2(\mathbf{X}^T\alpha)^3 + \sin(\mathbf{X}^T\alpha)$ .
- Quadratic:  $\mu_0^Q(\mathbf{X}) = \mathbf{X}^T\omega_1 + (\mathbf{X}^T\omega_2)^2$ ,
- Cubic:  $\mu_0^C(\mathbf{X}) = 0.1(\mathbf{X}^T\omega_2)^3$ .

## Parameters setting:

- $\alpha = \alpha^{(a)} = (0, -\mathbf{1}_{p/2}^T, \mathbf{1}_{p/2}^T)^T$ ,
- $\alpha = \alpha^{(b)} = (0, \mathbf{1}_p^T)^T$
- $\omega_1 = (0, \mathbf{1}_{p/2}^T, -\mathbf{1}_{p/2}^T)^T$ ,
- $\omega_2 = (0, \mathbf{1}_p^T)^T$ .

Table: The average of RE and PCD for  $p = 10$

|         | $\alpha = \alpha^{(a)}$         |        |                             |        | $\alpha = \alpha^{(b)}$         |        |                             |        |
|---------|---------------------------------|--------|-----------------------------|--------|---------------------------------|--------|-----------------------------|--------|
|         | Quadratic $\mu_0^Q(\mathbf{X})$ |        | Cubic $\mu_0^C(\mathbf{X})$ |        | Quadratic $\mu_0^Q(\mathbf{X})$ |        | Cubic $\mu_0^C(\mathbf{X})$ |        |
| Lin     | RE                              | PCD    | RE                          | PCD    | RE                              | PCD    | RE                          | PCD    |
| SUP     | 1                               | 96.20% | 1                           | 92.69% | 1                               | 96.85% | 1                           | 94.28% |
| NP      | 0.19                            | 92.64% | 0.38                        | 87.75% | 0.19                            | 92.99% | 0.38                        | 88.45% |
| SP      | 0.99                            | 96.16% | 1.00                        | 92.64% | 1.06                            | 96.94% | 1.10                        | 94.55% |
| SP.CV   | 0.98                            | 96.14% | 0.98                        | 92.64% | 0.94                            | 96.81% | 0.92                        | 94.18% |
| KRLS    | 1.01                            | 96.20% | 1.07                        | 92.79% | 1.02                            | 96.80% | 1.06                        | 94.30% |
| KRLS.CV | 0.94                            | 96.10% | 0.95                        | 92.54% | 0.94                            | 96.71% | 0.95                        | 94.14% |
| NL1     | RE                              | PCD    | RE                          | PCD    | RE                              | PCD    | RE                          | PCD    |
| SUP     | 1                               | 96.60% | 1                           | 95.60% | 1                               | 96.84% | 1                           | 96.24% |
| NP      | 0.20                            | 92.51% | 0.24                        | 91.24% | 0.19                            | 91.87% | 0.21                        | 89.81% |
| SP      | 3.05                            | 97.97% | 1.76                        | 96.40% | 3.12                            | 98.46% | 2.10                        | 97.63% |
| SP.CV   | 3.24                            | 97.99% | 1.89                        | 96.57% | 2.94                            | 98.23% | 1.62                        | 97.27% |
| KRLS    | 0.91                            | 97.28% | 0.98                        | 96.06% | 0.86                            | 97.56% | 0.94                        | 96.62% |
| KRLS.CV | 1.54                            | 97.08% | 1.18                        | 95.76% | 1.52                            | 97.34% | 1.16                        | 96.40% |

**Table:** The average of SE and CP of proposed estimators with  $p = 10$

| SE(CP %)           | Quadratic $\mu_0^Q(\mathbf{X})$ |              |              | Cubic $\mu_0^C(\mathbf{X})$ |              |              |
|--------------------|---------------------------------|--------------|--------------|-----------------------------|--------------|--------------|
|                    | SP                              | SP.CV        | SP.DCV       | SP                          | SP.CV        | SP.DCV       |
| Lin $\alpha^{(a)}$ | 5.300 (93.5)                    | 5.539 (94.6) | 5.728 (95.2) | 10.48 (93.8)                | 10.90 (94.6) | 11.28 (95.6) |
| Lin $\alpha^{(b)}$ | 4.931 (91.8)                    | 5.525 (93.5) | 5.711 (94.1) | 9.297 (89.5)                | 10.69 (92.2) | 11.06 (93.0) |
| NL1 $\alpha^{(a)}$ | 5.590 (81.1)                    | 7.214 (92.5) | 7.473 (93.5) | 10.75 (89.1)                | 12.10 (94.3) | 12.53 (95.1) |
| NL1 $\alpha^{(b)}$ | 4.827 (73.2)                    | 7.415 (90.4) | 7.682 (91.4) | 8.494 (81.6)                | 11.98 (90.4) | 12.41 (91.2) |
| NL2 $\alpha^{(a)}$ | 5.579 (81.1)                    | 7.204 (92.5) | 7.462 (93.5) | 10.74 (89.1)                | 12.09 (94.4) | 12.52 (95.1) |
| NL2 $\alpha^{(b)}$ | 4.820 (73.2)                    | 7.406 (90.4) | 7.671 (91.4) | 8.490 (81.6)                | 11.98 (90.4) | 12.40 (91.2) |



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# Real Data Analysis: MIMIC-IV

**Subjects:** 9,052 adult patients with sepsis who were first admitted to the ICU in the MIMIC-IV data.

**Outcome:** lactate clearance within 48 hours of ICU admission.

**Treatments:**  $A = -1$ : intravenous fluid resuscitation;  $A = 1$ : vasopressor therapy.

**Covariates:** age (years), admission weight (kilograms), blood urea nitrogen (BUN) amount (mg/dL), creatinine (mg/dL), white blood cell count (WBC) count ( $K/\mu L$ ), and heart rate (HR) (bpm)

**Sample size:** labeled dataset:  $n=184$  samples; unlabeled dataset:  $N=7,623$  samples.

Table: Treatment recommendation

| Treatment                    | Methods |      |      |       |      |         |
|------------------------------|---------|------|------|-------|------|---------|
|                              | SUP     | NP   | SP   | SP.CV | KRLS | KRLS.CV |
| A=-1: IV Fluid Resuscitation | 5113    | 5113 | 5112 | 5132  | 5150 | 5142    |
| A=1: Vasopressors            | 2694    | 2694 | 2695 | 2675  | 2657 | 2665    |

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# Conclusions

- We propose a novel **semi-supervised D-learning framework** for estimating optimal individualized treatment regime (ITR).
- Theoretical results:
  - ▶ Estimators converge asymptotically to normality at  $\sqrt{n}$  rate, depending on labeled sample size.
  - ▶ Under model misspecification, semi-supervised estimator achieves lower asymptotic variance than supervised estimator.
- Numerical studies:
  - ▶ When linear decision model is correct, unlabeled data does not improve efficiency.
  - ▶ When misspecified, semi-supervised estimator significantly improves performance.
  - ▶ Remains robust with increasing covariate dimension  $p$ ; outperforms fully nonparametric methods and KRLS-based estimators.
- Our method mitigates the curse of dimensionality, and maintains robustness and efficiency in multi-dimensions.

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# Thank you for listening.

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