

Inference for Linear Functionals in High-dimensional Linear Models

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High-dimensional linear regression

$$y_{n \times 1} = X_{n \times p} \beta_{p \times 1} + \epsilon_{n \times 1}.$$

- ▶ Number of covariates $p \gg$ sample size n .
- ▶ When $p > n$, $\|\beta\|_0 \leq k$.

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Estimation of β : Basis Pursuit (Chen & Donoho, '94); Lasso (Tibshirani, '96); SCAD (Fan & Li, '01); LARS(Efron, Hastie, Johnstone & Tibshirani, '04) Elastic Net (Zou & Hastie, '05); Adaptive Lasso (Zou, '05); Dantzig Selector (Candès & Tao, '07); Lasso and Dantzig (Bickel, Ritov & Tsybakov, '09); MCP (Zhang '10); scaled Lasso (Sun & Zhang, '10); square-root Lasso (Belloni, Chernozhukov & Wang, '11); ...

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- ▶ $\|\beta\|_2^2$
- ▶ $\beta^\top \Sigma \beta = \text{Var}(X_{i \cdot}^\top \beta)$
- ▶ $\beta_G^\top \Sigma_{G,G} \beta_G = \text{Var}(X_{i,G}^\top \beta_G)$

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3. ℓ_q Accuracy Functionals

- ▶ $\|\widehat{\beta} - \beta\|_2^2$ (Accuracy assessment of $\widehat{\beta}$)
- ▶ $\|\widehat{\beta} - \beta\|_q^q$ for $1 \leq q < 2$.

Overview of talk

- 1 Inference for β_i : Review of De-biasing
- 2 Minimality and Adaptivity
- 3 Uniform Procedure for All loadings
- 4 Further Discussion on Optimality

CI for β_i

- ▶ **Statistics:** Zhang & Zhang '14; van de Geer, Bühlmann, Ritov & Dezeure '14; Javanmard & Montanari '14;
- ▶ **Econometrics:** Chernozhukov, Belloni & Hansen '13; Chernozhukov, Hansen & Spindler '15;

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- ▶ **Econometrics:** Chernozhukov, Belloni & Hansen '13; Chernozhukov, Hansen & Spindler '15;
- ▶ Main idea: **Bias correction.**

$$\hat{\beta} = \arg \min_{\beta \in \mathbb{R}^p} \frac{1}{2n} \|y - X\beta\|_2^2 + \lambda \|\beta\|_1, \text{ with } \lambda \asymp \sqrt{\log p/n}\sigma$$

- ▶ De-biased Estimator:

$$\tilde{\beta}_i = \hat{\beta}_i + \underbrace{\hat{u}^\top \frac{1}{n} X^\top (y - X\hat{\beta})}_{\text{Correction term}} \text{ with } \left(\frac{1}{n} X^\top X \right) \hat{u} \approx e_i.$$

Construction of Projection Direction

Estimation error of $\hat{\beta}_i$: $\hat{\beta}_i - \beta_i = \mathbf{e}_i^\top (\hat{\beta} - \beta)$

$$\hat{u}^\top \frac{1}{n} \mathbf{X}^\top (Y - \mathbf{X}\hat{\beta}) = \hat{u}^\top \hat{\Sigma}(\beta - \hat{\beta}) + \hat{u}^\top \frac{1}{n} \mathbf{X}^\top \epsilon$$

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$$\begin{aligned}\hat{u}^\top \frac{1}{n} \mathbf{X}^\top (\mathbf{Y} - \mathbf{X} \hat{\beta}) &= \hat{u}^\top \hat{\Sigma} (\beta - \hat{\beta}) + \hat{u}^\top \frac{1}{n} \mathbf{X}^\top \epsilon \\ &= -\mathbf{e}_i^\top (\hat{\beta} - \beta) + \underbrace{(\hat{u}^\top \hat{\Sigma} - \mathbf{e}_i^\top)(\beta - \hat{\beta})}_{\text{Remaining Bias}} + \underbrace{\hat{u}^\top \frac{1}{n} \mathbf{X}^\top \epsilon}_{\text{Variance}}\end{aligned}$$

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De-biased estimator

$$\tilde{\beta}_i = \mathbf{e}_i^\top \hat{\beta} + \hat{u}^\top \frac{1}{n} X^\top (Y - X\hat{\beta}).$$

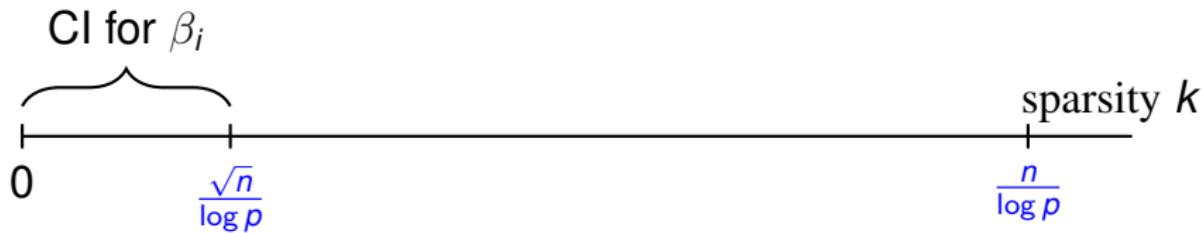
$$\hat{u} = \arg \min_{u \in \mathbb{R}^p} \left\{ \underbrace{u^\top \hat{\Sigma} u}_{\text{Variance}} : \underbrace{\|\hat{\Sigma} u - \mathbf{e}_i\|_\infty}_{\text{Constrained Bias}} \leq \|\mathbf{e}_i\|_2 \lambda_1 \right\}$$

Construction of CI for β_1

$$\tilde{\beta}_i - \beta_i = \underbrace{(\hat{u}^\top \hat{\Sigma} - e_i^\top)(\beta - \hat{\beta})}_{\text{Remaining Bias}} + \underbrace{\hat{u}^\top \frac{1}{n} X^\top \epsilon}_{\text{Variance}}$$

1. Variance $\sqrt{n} \hat{u}^\top \frac{1}{n} X^\top \epsilon \mid X \sim N(0, \hat{u}^\top \hat{\Sigma} \hat{u})$
2. $\sqrt{n} |(\hat{u}^\top \hat{\Sigma} - e_i^\top)(\beta - \hat{\beta})| \leq \sqrt{n} \|\hat{\Sigma} \hat{u} - e_i\|_\infty \|\beta - \hat{\beta}\|_1 \lesssim \frac{k \log p}{\sqrt{n}}$

Ultra-sparse case $k \ll \frac{\sqrt{n}}{\log p} \Rightarrow$ Variance dominates.



CI over $k \lesssim \frac{n}{\log p}$

$$\text{CI}_{\beta_1}(k) = \left[\tilde{\beta}_1 - \rho(k), \quad \tilde{\beta}_1 + \rho(k) \right],$$

with $\rho(k) = \frac{c_\alpha}{\sqrt{n}} \hat{\sigma} + \underbrace{Ck \frac{\log p}{n} \hat{\sigma}}_{\text{Account for remaining bias}}$.

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Minimaxity and Adaptivity (Cai and G., '16)



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For $k \lesssim \frac{n}{\log p}$,

1. Minimax expected length of CI for β_i .
2. Possible regime to construct adaptive CI for β_i .

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1. Minimax expected length of CI for β_i .
2. Possible regime to construct adaptive CI for β_i .

Adaptivity: without knowing the true sparsity k , construct CI as well as we know k .

Optimal expected length

- ▶ **Coverage**: Guaranteed coverage probability.
- ▶ **Precision**: As short as possible.

$$\Theta(k) = \left\{ \theta = (\beta, \Sigma, \sigma) : \|\beta\|_0 \leq k, \frac{1}{M_1} \leq \lambda_{\min}(\Sigma) \leq \lambda_{\max}(\Sigma) \leq M_1, 0 < \sigma \leq M_2 \right\}$$

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- ▶ For $0 < \alpha < 1$, CI has coverage for β_1 over $\Theta(k)$ if

$$\inf_{\theta \in \Theta(k)} \mathbf{P}_\theta(\beta_1 \in \text{CI}) \geq 1 - \alpha.$$

- ▶ For given k , the optimal length over $\Theta(k)$,

$$\mathcal{L}_\alpha^*(\Theta(k)) = \inf_{\substack{\text{CI having coverage} \\ \text{for } \beta_1 \text{ over } \Theta(k)}} \underbrace{\sup_{\theta \in \Theta(k)} \mathbf{E}_\theta \mathbf{L}(\text{CI})}_{\text{Precision}}.$$

Optimal expected length

Theorem 1(Cai and G., '16)

For $k \leq c \min\{p^\gamma, \frac{n}{\log p}\}$ with $0 \leq \gamma < \frac{1}{2}$,

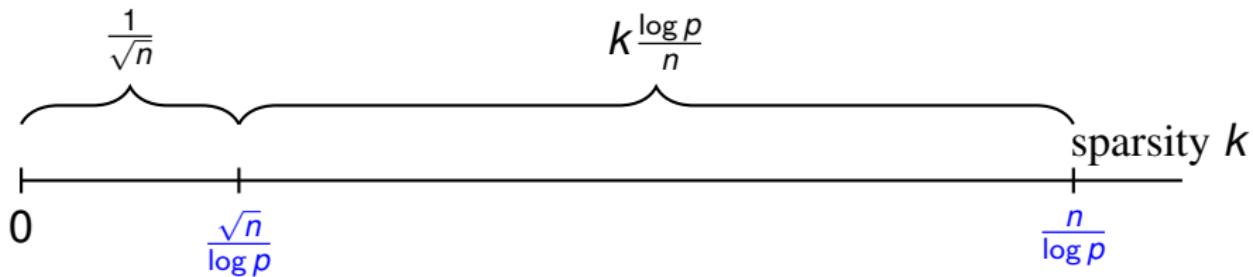
$$L_\alpha^*(\Theta(k)) \asymp \frac{1}{\sqrt{n}} + k \frac{\log p}{n}.$$

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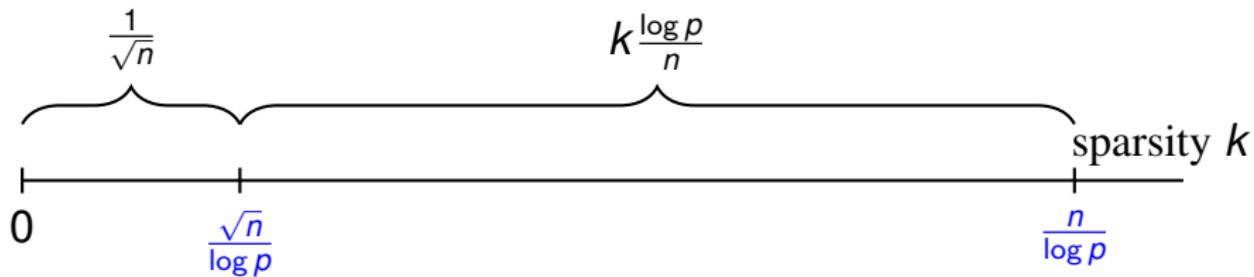


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Cls of length $\frac{1}{\sqrt{n}}$: NO coverage for $\frac{\sqrt{n}}{\log p} \ll k \lesssim \frac{n}{\log p}$.

Adaptive Procedures?

$$\text{Length of CI: } \rho(k) = \frac{c_\alpha}{\sqrt{n}} \hat{\sigma} + Ck \frac{\log p}{n} \hat{\sigma}.$$

Adaptivity \implies

Without knowing k , possible to construct CIs as well as known k ?

Adaptive procedures?

$k(\text{unknown true sparsity}) \leq k_u(\text{known upper bound}), \Theta(k) \subset \Theta(k_u)$

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$$k(\text{unknown true sparsity}) \leq k_u(\text{known upper bound}), \Theta(k) \subset \Theta(k_u)$$

Is it possible to construct CIs for β_1

1. coverage over $\Theta(k_u)$

Adaptive procedures?

k (unknown true sparsity) $\leq k_u$ (known upper bound), $\Theta(k) \subset \Theta(k_u)$

Is it possible to construct CIs for β_1

1. coverage over $\Theta(k_u)$
2. for any $\theta \in \Theta(k)$,

$$\mathbf{E}_\theta \mathbf{L}(\text{CI}) \lesssim \frac{1}{\sqrt{n}} + k \frac{\log p}{n}?$$

Lack of adaptivity

Theorem 2(Cai and G., '16)

For any $\theta = (\beta, I, \sigma) \in \Theta(k)$ and $k \leq k_u \leq \sqrt{p}$,

$$\inf_{\substack{\text{CI having coverage} \\ \text{for } \beta_1 \text{ over } \Theta(k_u)}} \mathbb{E}_\theta L(\text{CI}) \geq c \left(\frac{1}{\sqrt{n}} + k_u \frac{\log p}{n} \right) \sigma.$$

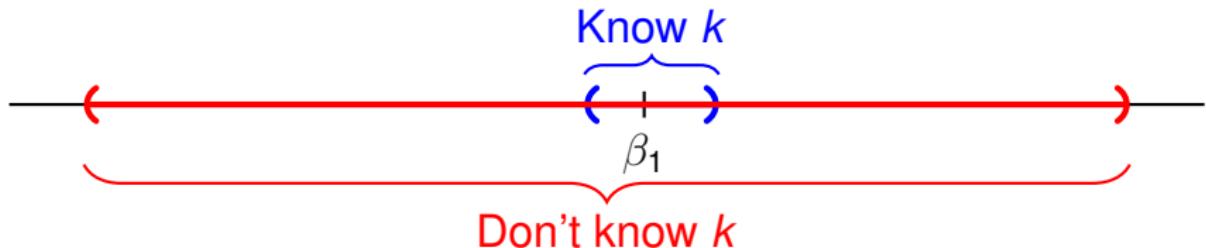
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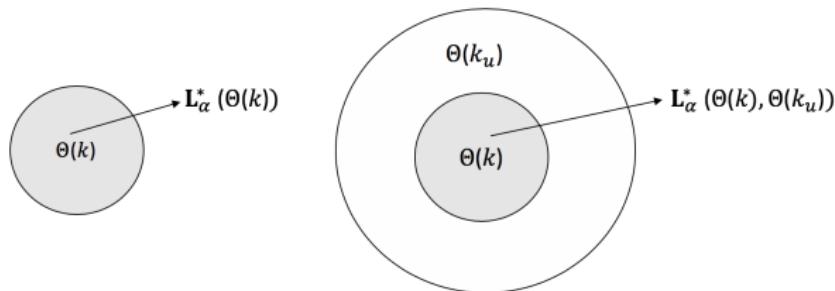
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For $\frac{\sqrt{n}}{\log p} \lesssim k_u \lesssim \frac{n}{\log p}$,

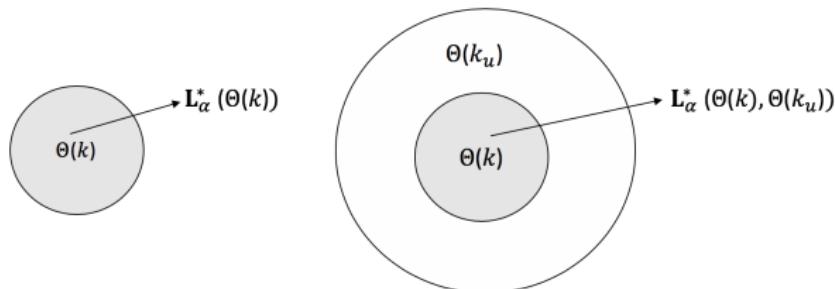


General Adaptation Benchmark



$$L_{\alpha}^{*}(\Theta(\textcolor{blue}{k}), \Theta(\textcolor{red}{k}_u)) = \inf_{\substack{\text{CI having coverage} \\ \text{for } \beta_1 \text{ over } \Theta(\textcolor{red}{k}_u)}} \sup_{\theta \in \Theta(\textcolor{blue}{k})} \mathbb{E}_{\theta} L(\text{CI})$$

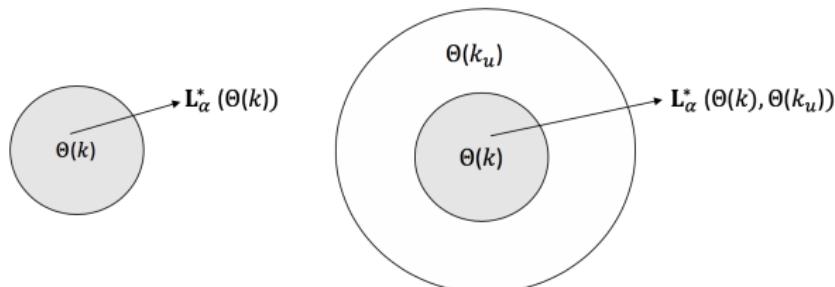
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$$L_\alpha^*(\Theta(\textcolor{blue}{k}), \Theta(\textcolor{red}{k}_u)) \gg L_\alpha^*(\Theta(\textcolor{blue}{k})) \implies \text{Impossible adaptive CI.}$$

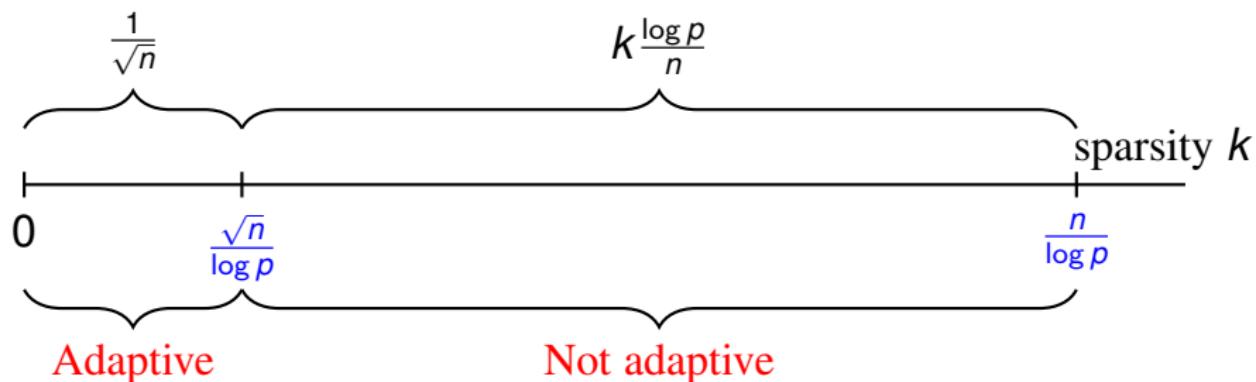
Summary of CI for β_1

- ▶ First constructed CI for β_1 over $k \lesssim \frac{n}{\log p}$.

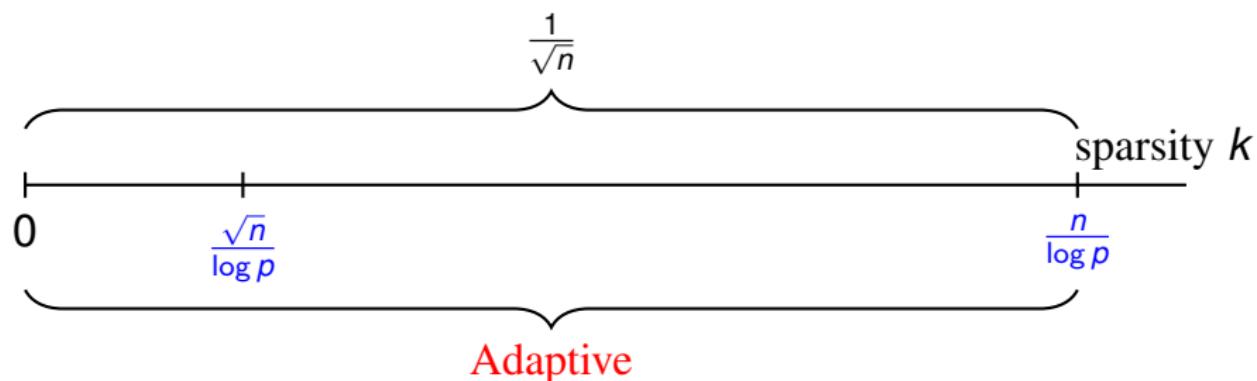


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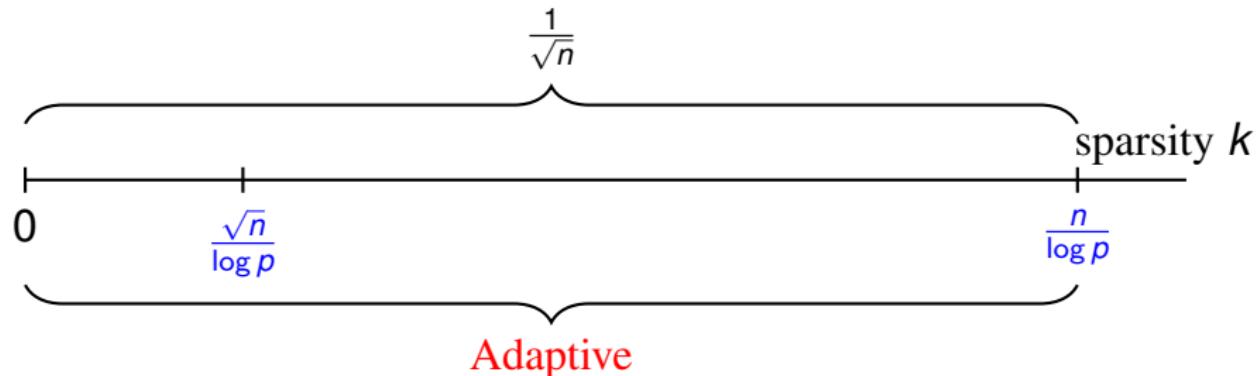


Comparison with known Σ



- ▶ CI for β_1 was constructed in Javanmard & Montanari '15.

Comparison with known Σ



- ▶ CI for β_1 was constructed in Javanmard & Montanari '15.
- ▶ Technical difference: **unknown** covariance structure between X_{i1} and X_{i2}, \dots, X_{ip} .

Four scenarios

Table: Confidence Intervals for $\eta^T \beta$

	Known Σ	Unknown Σ
Sparse Loading η (e.g., β_1)	✓	✓
Dense Loading η (e.g., $\sum_{i=1}^p \beta_i$)	?	?

Exact Loading: Sparse and Dense

We calibrate the sparsity levels as

$$k = p^\gamma, \quad k_u = p^{\gamma_u} \quad \text{for} \quad 0 \leq \gamma < \gamma_u \leq \frac{1}{2},$$

We consider exact loadings.

$$\max_{\{i: \eta_i \neq 0\}} |\eta_i| / \min_{\{i: \eta_i \neq 0\}} |\eta_i| \leq C_0,$$

$$\|\eta\|_0 = p^{\gamma_\eta} \quad \text{for} \quad 0 \leq \gamma_\eta \leq 1.$$

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$$\|\eta\|_0 = p^{\gamma_\eta} \quad \text{for} \quad 0 \leq \gamma_\eta \leq 1.$$

- (E1) x_{new} is called *exact sparse* if $\gamma_\eta \leq \gamma$;
- (E2) x_{new} is called *exact dense* if $\gamma_\eta > 2\gamma$;

CI for $\sum_{i=1}^p \beta_i$ (Cai and G., '16)

1. Centering at Lasso estimator

$$\text{CI}_{\sum \beta_i}(k) = \left[\sum_{i=1}^p \hat{\beta}_i - Ck \sqrt{\frac{\log p}{n}} \hat{\sigma}, \quad \sum_{i=1}^p \hat{\beta}_i + Ck \sqrt{\frac{\log p}{n}} \hat{\sigma} \right],$$

► NOT using de-biased estimator: **Inflation of variance!**

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- ▶ NOT using de-biased estimator: Inflation of variance!
- 2. $\text{CI}_{\sum \beta_i}(k)$ achieves optimal expected length $k \sqrt{\frac{\log p}{n}}$.
- 3. NOT possible to construct adaptive CI.
 - ▶ Without knowing k , CI must be longer than $k \sqrt{\frac{\log p}{n}}$.

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3. NOT possible to construct adaptive CI.
 - Without knowing k , CI must be longer than $k \sqrt{\frac{\log p}{n}}$.
4. The information Σ is NOT useful.

Confidence intervals for $\eta^\top \beta$

	Known Σ	Unknown Σ
Sparse Loading η	$\frac{\ \eta\ _2}{\sqrt{n}}$	$\ \eta\ _2 \left(\frac{1}{\sqrt{n}} + \frac{k \log p}{n} \right)$
Dense Loading η		$\ \eta\ _\infty k \sqrt{\frac{\log p}{n}}$

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	Known Σ	Unknown Σ
Sparse Loading η	$k \lesssim \frac{n}{\log p}$	$k \ll \frac{\sqrt{n}}{\log p}$
Dense Loading η		Impossible

Tony Cai and Zijian Guo. *Confidence intervals for high-dimensional linear regression: Minimax rates and adaptivity*. AOS, 2017.

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Beyond Minimaxity

The minimax results for **dense η** are pessimistic.

Let's put the minimaxity aside first.

A practical question: Inference procedure for $\eta^\top \beta$?

1. Works for all η .
2. Requires no knowledge of sparsity.

Literature for $\eta^T \beta$

Cai and Guo (2017)	η is sparse
Athey, Imbens, Wager (2018)	$\ \eta\ _2$ is bounded
Zhu and Bradic (2018)	Certain sparse η

Susan Athey, Guido W Imbens, and Stefan Wager. *Approximate residual balancing: debiased inference of average treatment effects in high dimensions*. JRSSB, 2018.

Yinchu Zhu and Jelena Bradic. *Linear hypothesis testing in dense high-dimensional linear models*. JASA, 2018.

A uniform procedure for all $x_{\text{new}} \in \mathbb{R}^P$

Revisit β_i

$$\begin{aligned}\widehat{\boldsymbol{u}}^\top \frac{1}{n} \boldsymbol{X}^\top (\boldsymbol{Y} - \boldsymbol{X} \widehat{\boldsymbol{\beta}}) &= \widehat{\boldsymbol{u}}^\top \widehat{\boldsymbol{\Sigma}} (\boldsymbol{\beta} - \widehat{\boldsymbol{\beta}}) + \widehat{\boldsymbol{u}}^\top \frac{1}{n} \boldsymbol{X}^\top \boldsymbol{\epsilon} \\ &= -\mathbf{e}_i^\top (\widehat{\boldsymbol{\beta}} - \boldsymbol{\beta}) + (\widehat{\boldsymbol{\Sigma}} \widehat{\boldsymbol{u}} - \mathbf{e}_i)^\top (\boldsymbol{\beta} - \widehat{\boldsymbol{\beta}}) + \widehat{\boldsymbol{u}}^\top \frac{1}{n} \boldsymbol{X}^\top \boldsymbol{\epsilon}\end{aligned}$$

Bias-corrected estimator

$$\widetilde{\beta}_{1,i} = \mathbf{e}_i^\top \widehat{\boldsymbol{\beta}} + \widehat{\boldsymbol{u}}^\top \frac{1}{n} \boldsymbol{X}^\top (\boldsymbol{Y} - \boldsymbol{X} \widehat{\boldsymbol{\beta}}).$$

$$\widehat{\boldsymbol{u}} = \arg \min_{\boldsymbol{u} \in \mathbb{R}^p} \left\{ \underbrace{\boldsymbol{u}^\top \widehat{\boldsymbol{\Sigma}} \boldsymbol{u}}_{\text{Variance}} : \underbrace{\|\widehat{\boldsymbol{\Sigma}} \boldsymbol{u} - \mathbf{e}_i\|_\infty}_{\text{Constrained Bias}} \leq \|\mathbf{e}_i\|_2 \lambda_1 \right\}$$

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Challenges for Dense Loadings

Dense η :

Feasible Set: $\left\| \widehat{\Sigma} u - \eta \right\|_{\infty} \leq \|\eta\|_2 \lambda_1$

$$\|\eta\|_2 \lambda_1 \geq \|\eta\|_{\infty} \Rightarrow \widehat{u} = 0!$$

Example: If η is decaying as $\eta_j \asymp j^{-\delta}$, then $\|\eta\|_2 \asymp p^{\frac{1}{2}-\delta}$.

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Example: If η is decaying as $\eta_j \asymp j^{-\delta}$, then $\|\eta\|_2 \asymp p^{\frac{1}{2}-\delta}$.

Bias-corrected estimator=plug-in estimator,

$$\widetilde{\eta^T \beta} = \eta^T \widehat{\beta} + \widehat{u}^T \frac{1}{n} X^T (Y - X \widehat{\beta}) = \eta^T \widehat{\beta}.$$

Curse of dimensionality from dense η .

New Projection Direction

$$\hat{u} = \arg \min_{u \in \mathbb{R}^p} u^\top \hat{\Sigma} u$$

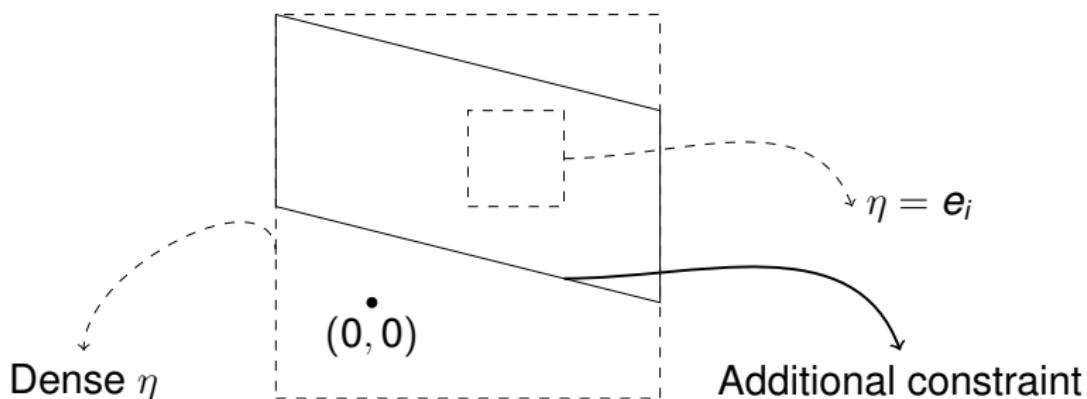
subject to $\left\| \hat{\Sigma} u - \eta \right\|_\infty \leq \|\eta\|_2 \lambda_1$

$$\left| \eta^\top \hat{\Sigma} u - \|\eta\|_2^2 \right| \leq \|\eta\|_2^2 \lambda_1$$

The proposed estimator for $\eta^\top \beta$ is

$$\widehat{\eta^\top \beta} = \eta^\top \hat{\beta} + \hat{u}^\top \frac{1}{n} X^\top (Y - X \hat{\beta}) \quad (1)$$

Additional Constraint and Feasible Set



- ▶ Small dashed: $\eta = e_i$.
- ▶ Large dashed: dense η **without** additional constraint.
- ▶ Solid parallelogram: dense η **with** additional constraint.

$$\left| \eta^\top \widehat{\Sigma} u - \|\eta\|_2^2 \right| \leq \|\eta\|_2^2 \lambda_1$$

Bias-Variance Tradeoff

Bias and Variance Tradeoff.

- ▶ Minimizing variance with bias constrained.

$$\left| (\widehat{\Sigma} \widehat{u} - \eta)^\top (\beta - \widehat{\beta}) \right| \leq \|\widehat{\Sigma} \widehat{u} - \eta\|_\infty \|\beta - \widehat{\beta}\|_1$$

- ▶ Minimizing variance with bias and variance constrained.

$$\widehat{u} = \arg \min_{u \in \mathbb{R}^p} u^\top \widehat{\Sigma} u$$

$$\text{subject to } \left\| \widehat{\Sigma} u - \eta \right\|_\infty \leq \|\eta\|_2 \lambda_1$$

$$\left| \eta^\top \widehat{\Sigma} u - \|\eta\|_2^2 \right| \leq \|\eta\|_2^2 \lambda_1$$

Enhancing Variance Lemma

Lemma 1 (Cai, Cai, G. (2018)).

Under regularity conditions, we have

$$c_0 \frac{\|\eta\|_2}{\sqrt{n}} \leq \sqrt{\frac{1}{n} \hat{u}^\top \hat{\Sigma} \hat{u}} \leq C_0 \frac{\|\eta\|_2}{\sqrt{n}}$$

- ▶ Lower bound does not hold without the additional constraint
- ▶ Additional constraint leads to a dominating variance

Theory

Theorem 2 (Cai, Cai, G. (2018)).

Under regularity conditions and $\|\beta\|_0 \leq c\sqrt{n}/\log p$, then

$$\frac{1}{\sqrt{\mathbf{V}}} (\widehat{\eta^\top \beta} - \eta^\top \beta) \xrightarrow{d} N(0, 1) \quad (2)$$

Theory

Theorem 2 (Cai, Cai, G. (2018)).

Under regularity conditions and $\|\beta\|_0 \leq c\sqrt{n}/\log p$, then

$$\frac{1}{\sqrt{V}} (\widehat{\eta^\top \beta} - \eta^\top \beta) \xrightarrow{d} N(0, 1) \quad (2)$$

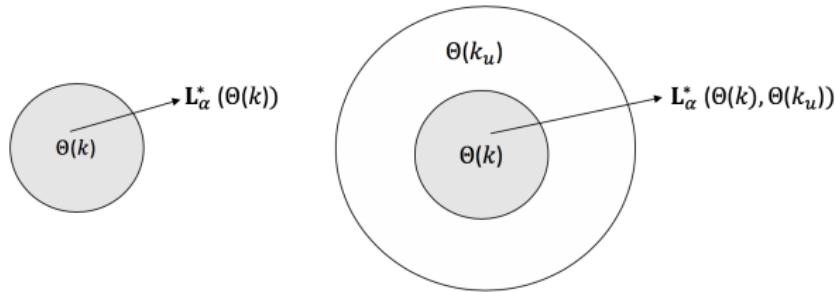
$V \asymp \frac{\|\eta\|_2}{\sqrt{n}}$ depends on η .

Works if $\|\beta\|_0 \leq c\sqrt{n}/\log p$.

Overview of talk

- 1 Inference for β_i : Review of De-biasing
- 2 Minimality and Adaptivity
- 3 Uniform Procedure for All loadings
- 4 Further Discussion on Optimality

Adaptive Optimal



$$L_\alpha^*(\Theta(\textcolor{blue}{k}), \Theta(\textcolor{red}{k}_u)) = \inf_{\substack{\text{CI having coverage} \\ \text{for } \beta_1 \text{ over } \Theta(\textcolor{red}{k}_u)}} \sup_{\theta \in \Theta(\textcolor{blue}{k})} \mathbb{E}_\theta L(\text{CI})$$

Adaptive optimal: a procedure achieving $L_\alpha^*(\Theta(\textcolor{blue}{k}), \Theta(\textcolor{red}{k}_u))$.

Review of Exact Loading

We calibrate the sparsity levels as

$$k = p^\gamma, \quad k_u = p^{\gamma_u} \quad \text{for } 0 \leq \gamma < \gamma_u \leq 1,$$

$$c_0 \leq \max_{\{i: \eta_i \neq 0\}} |\eta_i| / \min_{\{i: \eta_i \neq 0\}} |\eta_i| \leq C_0,$$

$$\|\eta\|_0 = p^{\gamma_\eta} \quad \text{for } 0 \leq \gamma_\eta \leq 1.$$

(E1) x_{new} is called *exact sparse* if $\gamma_\eta \leq 2\gamma$;

(E2) x_{new} is called *exact dense* if $\gamma_\eta > 2\gamma$;

Possibility of Adaptive Testing

Suppose that $k \leq k_u \lesssim \frac{\sqrt{n}}{\log p}$,

	$\gamma, \gamma_u, \gamma_\eta$	$L_\alpha^*(\Theta(k))$	Rel	$L_\alpha^*(\Theta(k), \Theta(k_u))$	Adpt
(E1)	$\gamma_\eta \leq 2\gamma$	$\frac{\ \eta\ _2}{\sqrt{n}}$	\asymp	$\frac{\ \eta\ _2}{\sqrt{n}}$	Yes
(E2-a)	$\gamma < \gamma_u < \frac{1}{2}\gamma_\eta$	$\ \eta\ _\infty k \sqrt{\frac{\log p}{n}}$	\ll	$\ \eta\ _\infty k_u \sqrt{\frac{\log p}{n}}$	No
(E2-b)	$\gamma < \frac{1}{2}\gamma_\eta \leq \gamma_u$	$\ \eta\ _\infty k \sqrt{\frac{\log p}{n}}$	\ll	$\frac{\ \eta\ _2}{\sqrt{n}}$	No

- ▶ Cut-off for “dense” and “sparse” occurs at $\gamma_\eta = 2\gamma$.

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- ▶ Cut-off for “dense” and “sparse” occurs at $\gamma_\eta = 2\gamma$.
- ▶ If $\gamma_u \geq \frac{1}{2}\gamma_\eta$, then the optimal test is of order $\frac{\|\eta\|_2}{\sqrt{n}}$
- ▶ In absence of accurate sparsity information, the proposed inference procedure $\eta^\top \beta$ is **adaptive optimal** for **all** exact loadings η .

Take Home Message

- ▶ The best we can aim for: $L_\alpha^*(\Theta(k), \Theta(k_u))$
- ▶ Dense linear functionals are harder than sparse ones.
- ▶ Uniform Procedure over all loadings.

Reference and Acknowledgement

Tony Cai and Zijian Guo. *Confidence intervals for high-dimensional linear regression: Minimax rates and adaptivity*. AOS, 2017.

Cai, T., Cai, T.T., Guo, Z. (2018). Individualized Treatment Selection: An Optimal Hypothesis Testing Approach In High-dimensional Models. Submitted.

Acknowledgement to NSF and NIH for fundings.

Thank you!

CI for $\eta^\top \beta$ (Cai and G., '16)

Fundamental difference in terms of minimaxity and adaptivity,

1. Sparse loading $\eta : \beta_i$
2. Dense loading $\eta : \sum_{i=1}^p \beta_i$

CI for $\eta^\top \beta$ (Cai and G., '16)

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Plug-in Lasso Estimators

$$\begin{aligned}\beta_1 : \quad & \widehat{\beta}_1 - \beta_1 = \langle \mathbf{e}_1, \widehat{\beta} - \beta \rangle \\ \eta^\top \beta : \quad & \eta^\top \widehat{\beta} - \eta^\top \beta = \langle \eta, \widehat{\beta} - \beta \rangle\end{aligned}$$

CI for $\eta^\top \beta$ (Cai and G., '16)

Fundamental difference in terms of minimaxity and adaptivity,

1. Sparse loading $\eta : \beta_i$
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Plug-in Lasso Estimators

$$\begin{aligned}\beta_1 : \quad & \widehat{\beta}_1 - \beta_1 = \langle \mathbf{e}_1, \widehat{\beta} - \beta \rangle \\ \eta^\top \beta : \quad & \eta^\top \widehat{\beta} - \eta^\top \beta = \langle \eta, \widehat{\beta} - \beta \rangle\end{aligned}$$

- **Sparse η :** Correct the bias \Rightarrow Similar to β_1 .
- **Dense η :** NOT correct the bias \Rightarrow Inflated variance.

Balance bias and variance.

Simulation Setting

Simulation Setting with $\eta^\top \beta = 1.08$

- ▶ $p = 501, n = n_2 = n$
- ▶ $\beta_{1,0} = -0.1, \beta_{1,j} = 0.4(j - 1)$ for $1 \leq j \leq 10$
- ▶ $\beta_{2,0} = -0.5, \beta_{2,j} = 0.2(j - 1)$ for $1 \leq j \leq 5$
- ▶ $X_{new,j} \sim N(0, 1)$ for $1 \leq i \leq 10$ and
 $X_{new,j} \sim 0.2 * N(0, 1)$ for $i \geq 11$

- ▶ **Adaptive optimality**: If the sparsity is unknown, what is the optimal length of CI?

Size

The parameter space

$$\Theta(s) = \left\{ \boldsymbol{\theta} = \begin{pmatrix} \beta, \Sigma_1, \sigma_1 \\ \beta_2, \Sigma_2, \sigma_2 \end{pmatrix} : \|\beta\|_0 \leq s, 0 < \sigma_k \leq M_0, \lambda_{\min}(\Sigma_k) \geq c_0, \text{ for } k = 1, 2 \right\},$$

For a test ϕ , its size is

$$\alpha(s, \phi) = \sup_{\theta \in \mathcal{H}_0(s)} \mathbb{E}_{\theta} \phi. \quad (3)$$

with

$$\mathcal{H}_0(s) = \{ \boldsymbol{\theta} \in \Theta(s) : \eta^T (\beta - \beta_2) \leq 0 \}$$

Power

The local alternative parameter space

$$\mathcal{H}_1(s, \tau) = \{\boldsymbol{\theta} \in \Theta(s) : \mathbf{x}_{\text{new}}^\top (\boldsymbol{\beta} - \boldsymbol{\beta}_2) = \tau > 0\}.$$

The power of ϕ over $\mathcal{H}_1(s, \tau)$ is defined as

$$\omega(s, \tau, \phi) = \inf_{\boldsymbol{\theta} \in \mathcal{H}_1(s, \tau)} \mathbb{E}_{\boldsymbol{\theta}} \phi. \quad (4)$$

Optimality: identify the **smallest** τ

- ▶ The size is controlled over $\mathcal{H}_0(s)$;
- ▶ The corresponding power over $\mathcal{H}_1(s, \tau)$ is large

Minimax Detection Boundary

Minimax detection boundary is defined as

$$\tau_{\min}(k, x_{\text{new}}) = \arg \min_{\tau} \left\{ \tau : \sup_{\phi: \alpha(s, \phi) \leq \alpha} \omega(s, \tau, \phi) \geq 1 - \eta \right\}.$$

A test ϕ is minimax optimal if

$$\alpha(s, \phi) \leq \alpha \quad \text{and} \quad \omega(s, \phi, \tau) \geq 1 - \eta \quad \text{for } \tau \asymp \tau_{\min}(k, x_{\text{new}})$$

Minimax assumes s is known.

Unknown Sparsity Level

Capture the optimality for unknown sparsity level?

We consider two sparsity levels, $k \leq k_u$.

- ▶ k denotes the true sparsity level;
- ▶ k_u denotes an upper bound for the sparsity level.

The size is uniformly controlled over $\mathcal{H}_0(k_u)$,

$$\alpha(k_u, \phi) = \sup_{\theta \in \mathcal{H}_0(k_u)} \mathbb{E}_{\theta} \phi \leq \alpha. \quad (5)$$

Adaptive Detection Boundary

The adaptive detection boundary $\tau_{\text{adap}}(k_u, k, x_{\text{new}})$

$$\tau_{\text{adap}}(k_u, k, x_{\text{new}}) = \arg \min_{\tau} \left\{ \tau : \sup_{\phi: \alpha(k_u, \phi) \leq \alpha} \omega(\mathbf{k}, \tau, \phi) \geq 1 - \eta \right\}.$$

A test ϕ is adaptive optimal if

$$\alpha(k_u, \phi) \leq \alpha \quad \text{and} \quad \omega(k, \tau, \phi) \geq 1 - \eta \quad \text{for } \tau \asymp \tau_{\text{adap}}(k_u, k, x_{\text{new}})$$

An adaptive optimal test would be the best that we can aim for if there is lack of accurate information on sparsity.

Adaptive Hypothesis Testing

- ▶ If $\tau_{\min}(k, x_{\text{new}}) \asymp \tau_{\text{adap}}(k_u, k, x_{\text{new}})$, the testing problem is adaptive.
- ▶ If $\tau_{\min}(k, x_{\text{new}}) \ll \tau_{\text{adap}}(k_u, k, x_{\text{new}})$, the testing problem is NOT adaptive.

Numerical Comparison

Other methods

1. HITS
2. Plug-in scaled Lasso: $x_{\text{new}}^T(\widehat{\beta} - \widehat{\beta}_2)$
3. Plug-in debiased Lasso: $x_{\text{new}}^T(\widetilde{\beta} - \widetilde{\beta}_2)$

Numerical Comparison

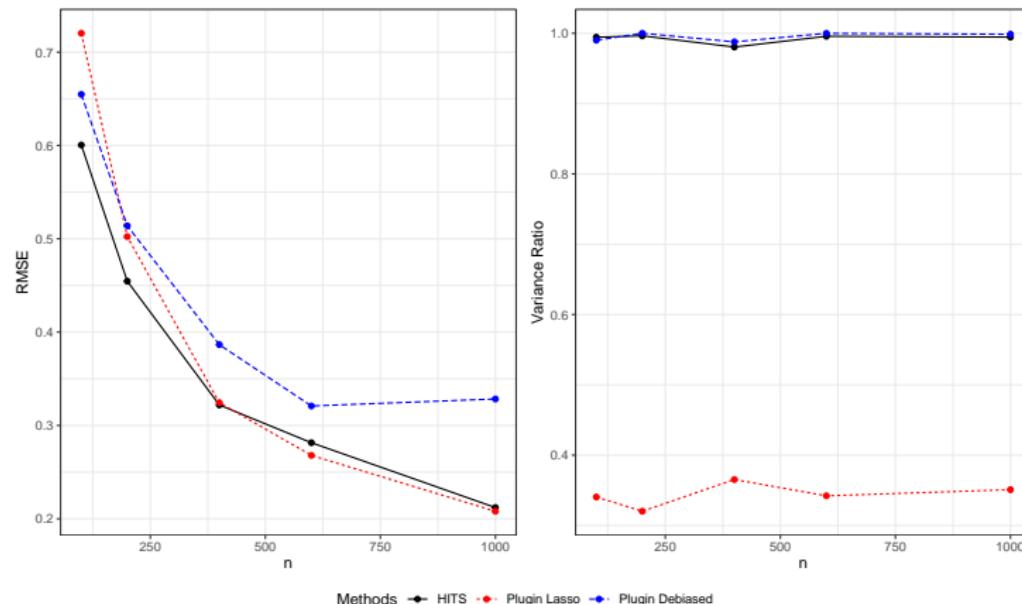
Other methods

1. HITS
2. Plug-in scaled Lasso: $x_{\text{new}}^T(\hat{\beta} - \hat{\beta}_2)$
3. Plug-in debiased Lasso: $x_{\text{new}}^T(\tilde{\beta} - \tilde{\beta}_2)$

Computation comparison

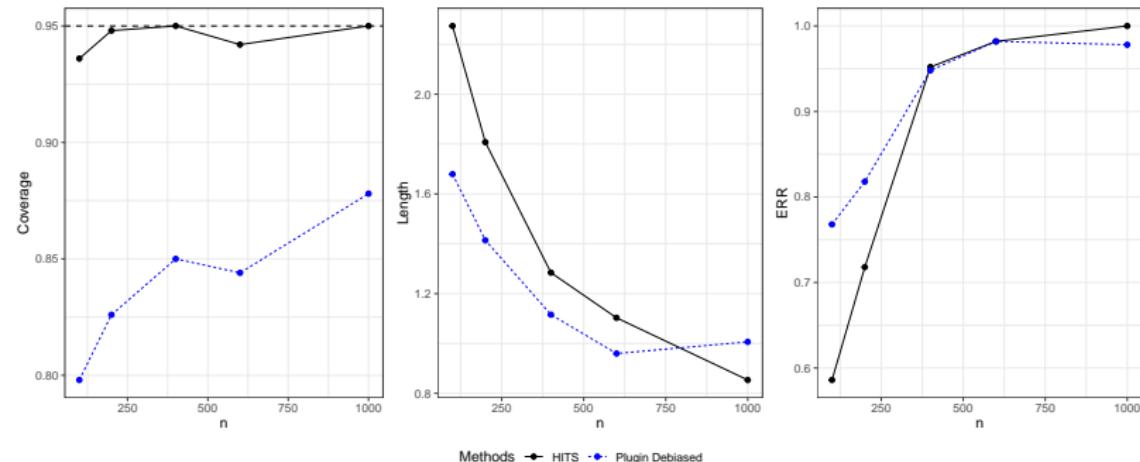
1. HITS: 4 Lasso
2. Plug-in scaled Lasso: 2 Lasso
3. Plug-in debiased Lasso: 1,004 Lasso ($2p + 2$)

RMSE



- ▶ Plug-in Lasso: hard to do inference
- ▶ HITS has smaller RMSE than Plug-in Debiased

ITE and CI



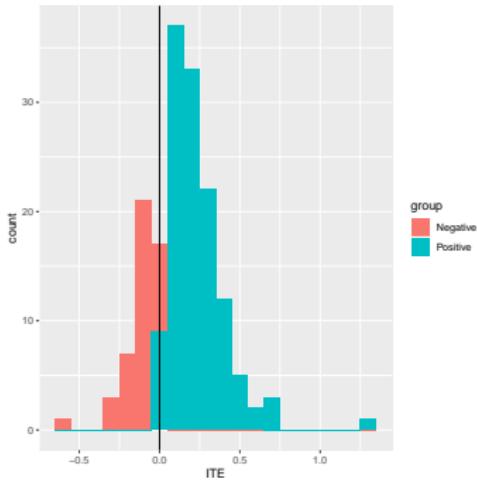
- ▶ Better coverage
- ▶ Computationally more efficient
- ▶ Comparable length and ERR

Real Data Analysis

Rheumatoid Arthritis (RA)

- ▶ Treatment 1: methotraxate+ anti-TNF (92 patients)
- ▶ Treatment 2: methotraxate (91 patients)
- ▶ Outcome – $\log(\text{CRP})$
Higher value of $Y \rightarrow$ Better treatment response.
- ▶ 171 Predictors, including Clinical measurement, EHR and SNP

Real Data Analysis

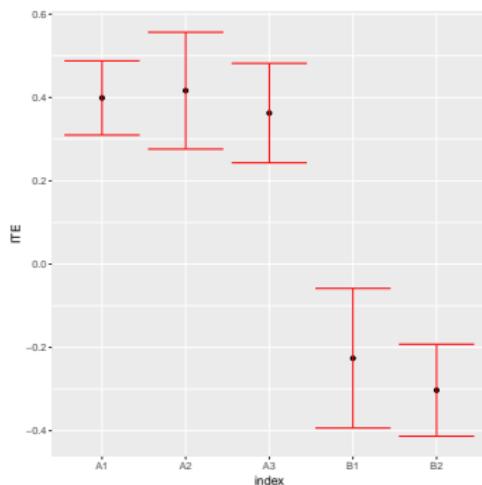


- ▶ About 72% benefit from the combination therapy.

Real Data Analysis

Patients	rs12506688	SLE mention	rs2843401	rs8043085	...
A	=0	≥ 1	= 0	> 0	...
B	>0	No	>0	=0	...

(SLE= Systemic Lupus Erythematosus)



The treatment effect is **heterogeneous** across patients.