Adjusting treatment effect estimates in randomized experiments with the Lasso

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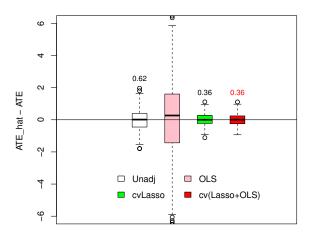
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Joint work with Adam Bloniarz, Hanzhong Liu, Cunhui Zhang, and Bin Yu, 2015

A simulation study based on experimental data with a large number (p) of covariates relative to sample size n

Boxplot with Standard Deviation on top



$$n=59*2$$
, $p=59$, $p_{A}=0.5$

Bowers et. al. 2015

The Challenge: Funky Outcomes and Many Covariates

The General Motivation: The civic effects of terrorism.

The Specific Study: The London Bombings of 2005 and voluntary activities (from the 2005 UK Home Office Survey).

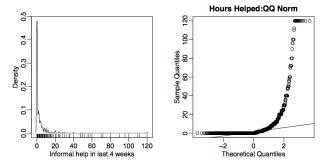


Figure 1: Hours of informal help for non-relatives in the last four weeks (UK Home Office Survey 2005). These observations from a window around the London Bombing of 2005 (-2 weeks $n_{\text{pre}} = 1195$, +8 weeks $n_{\text{post}} = 3341$).

Our plan

- Theoretical study of Lasso under Neyman-Rubin model to gain insights into when Lasso works as an adjustment method.
- Simulation and real data experiments to argument theoretical study about regularization parameter selection and compare Lasso and its variants.
- Estimation model is not being assumed. Useful framework to study other ML methods: SVM, deep learning, etc.

Related work

- Regression adjustment for fix p under Neyman-Rubin:
 - Freedman DA 2008;
 - Freedman DA 2008 (several treatments);
 - Lin W 2013;
- Regression adjustment for p > n under regression model:
 - Belloni A, Chernozhukov V, Hansen C 2013;
 - Belloni A, Chernozhukov V, Fernandez-Val I, Hansen C 2013;
 - Tian L, Alizadeh A, Gentles A, Tibshirani R 2014;
 - Rosenblum M, Liu H, En-Hsu Y 2014;

Other Related Work

- Imai and Ratkovic 2013: findit
- Ratkovic and Tingley 2015: sparsereg
- Grimmer, Messing, Westwood 2014
- Bowers, Panagopoulos, and Fredrickson 2013; Bowers 2014; Bowers, Fredrickson, Hansen 2015
- Athey and Imbens 2015

Neyman-Rubin model

SUTVA (Rubin, 1980) (Stable Unit Treatment Value Assumption)

- No interference
- Only a single version of each treatment level

Under SUTVA

- a_i, b_i: potential outcomes for unit i under treatment and control
- The parameter we are trying to estimate in this work is the Average Treatment Effect

$$ATE = \frac{1}{n} \sum_{i=1}^{n} a_i - \frac{1}{n} \sum_{i=1}^{n} b_i$$

Randomized experiment

- Randomness comes from treatment assignment
- T_i: random indicator of treatment for unit i
- A: set of treated units (random)

$$A = \{i, T_i = 1\}$$

B: set of control units (random)

$$B = \{i, T_i = 0\}$$

y_i: observed outcome

$$y_i = a_i T_i + b_i (1 - T_i), i = 1, ..., n$$

Notations

• n, n_A, n_B: number of treated and control units

$$p_{A} = n_{A}/n; \ p_{B} = n_{B}/n$$

Average on the population/treated/control

$$\bar{a} = \frac{1}{n} \sum_{i=1}^{n} a_i, \ \bar{a}_A = \frac{1}{n_A} \sum_{i \in A} a_i$$

$$\bar{b} = \frac{1}{n} \sum_{i=1}^{n} b_i, \ \bar{b}_B = \frac{1}{n_B} \sum_{i \in B} b_i$$

Simple estimator

- Understanding the assignment mechanism is crucial for causal inference
- If assignment is completely randomized, the ATE can be estimated without bias using the simple difference in means:

$$\widehat{ATE}_{unadj} = \bar{a}_A - \bar{b}_B$$

Regression adjustment

This leads us to consider estimators of the form

$$\widehat{ATE} = \left[\bar{a}_A - (\bar{x}_A - \bar{x})^T \beta^{(a)}\right] - \left[\bar{b}_B - (\bar{x}_B - \bar{x})^T \beta^{(b)}\right]$$

Regression adjustment with interaction (Lin W 2013¹)

$$y_i \sim T_i, x_i, T_i(x_i - \bar{x})$$

Minimizing

$$\sum_{i=1}^{n} \left\{ y_i - \tau_a T_i - \tau_b (1 - T_i) - T_i (x_i - \bar{x})^T \beta^{(a)} - (1 - T_i) (x_i - \bar{x})^T \beta^{(b)} \right\}^2$$

$$\widehat{ATE}_{OLS} = \left[\bar{a}_A - (\bar{x}_A - \bar{x})^T \hat{\beta}^{(a)}\right] - \left[\bar{b}_B - (\bar{x}_B - \bar{x})^T \hat{\beta}^{(b)}\right]$$

¹Lin W (2013). Agnostic notes on regression adjustments to experimental data: reexamining Freedman's critique. The Annals of Applied Statistics 7:295-318.

Regression adjustment

Benefits of regression adjustment:

Under regularity conditions, \widehat{ATE}_{OLS} is asymptotic normal with asymptotic variance **no larger than** that of the \widehat{ATE}_{unadi} (Lin W 2013)

- Question: what if p > n?
 - Observe many covariates
 - Main effects + interactions
 - Polynomial or splines

Regression adjustment using Lasso

- Sparsity: not all the covariates are relevant
- · Lasso (Tibshirani 1996), minimizing

$$\frac{1}{2n}\sum_{i=1}^{n}\left\{y_{i}-\tau_{a}T_{i}-\tau_{b}(1-T_{i})-T_{i}(x_{i}-\bar{x})^{T}\beta^{(a)}-(1-T_{i})(x_{i}-\bar{x})^{T}\beta^{(b)}\right\}^{2}$$
$$+\lambda_{a}||\beta^{(a)}||_{1}+\lambda_{b}||\beta^{(a)}||_{1}$$

Equivalent to (similarly for control group):

$$\hat{\beta}^{(a)} = \operatorname*{argmin}_{\beta} \frac{1}{2n_A} \sum_{i \in A} \left\{ a_i - \bar{a}_A - (x_i - \bar{x}_A)^T \beta \right\}^2 + \lambda_a ||\beta||_1$$

Regression adjustment using Lasso

Define Lasso adjusted ATE estimator

$$\widehat{\textit{ATE}}_{\texttt{Lasso}} = \left[\bar{a}_{\textit{A}} - (\bar{x}_{\textit{A}} - \bar{x})^{T} \hat{\beta}^{(a)}\right] - \left[\bar{b}_{\textit{B}} - (\bar{x}_{\textit{B}} - \bar{x})^{T} \hat{\beta}^{(b)}\right]$$

- We are interested in
 - Sufficient conditions for Lasso adjustment to work: Neyman-Rubin allows consideration of transformation before adjustment.

In particular, we hope to get

- asymptotic normality of the adjusted effect estimate, and
- estimate of the asymptotic variance

Assumptions

Assume there exist

- S^(a) ⊆ {1, 2, ..., p}: relevant covariates set for treatment, and
- $\beta^{(a)}$: projection coefficients.
- Then we can decompose potential outcome as follows:

$$a_i = \bar{a} + (\bar{x}_i - \bar{x})^T \beta^{(a)} + e_i^{(a)}$$

- Similar for control, then define $S = S^{(a)} \cup S^{(b)}$
- All the quantities above are fixed

Condition 1: Stability of treatment assignment probability

$$n_A/n \to p_A, \ n_B/n \to p_B, \ \text{as } n \to \infty,$$

for some $p_A, p_B \in (0, 1)$

Condition 2: The centered moment conditions

$$n^{-1} \sum_{i=1}^{n} (x_{ij} - \bar{x}_i)^4 \le L, \ \forall j$$

$$n^{-1} \sum_{i=1}^{n} (e_i^{(a)})^4 \le L; \ n^{-1} \sum_{i=1}^{n} (e_i^{(b)})^4 \le L$$

• Condition 3: The means $n^{-1} \sum_{i=1}^{n} (e_i^{(a)})^2$, $n^{-1} \sum_{i=1}^{n} (e_i^{(b)})^2$ and $n^{-1} \sum_{i=1}^{n} e_i^{(a)} e_i^{(b)}$ converge to finite limits.

Two quantities needed for high-dim case

Sparsity measures (number of nonzero coefficients)

$$s = |\{j : \beta_i^{(a)} \neq 0 \text{ or } \beta_i^{(b)} \neq 0\}|$$

Maximum covariance

$$\delta_n = \max_{\omega = a, b} \left\{ \max_{j} \left| \frac{1}{n} \sum_{i=1}^{n} (x_{ij} - \bar{x}_j) \left(e_i^{(\omega)} - \bar{e}^{(\omega)} \right) \right| \right\}$$

Further assumptions for consistency of Lasso

Condition 4: Decay and scaling

$$\delta_n = o\left(\frac{1}{s\sqrt{\log p}}\right); \quad (s\log p)/\sqrt{n} = o(1)$$

 $||h_{S}||_{1} < C_{S}||\hat{\Sigma}h||_{\infty}, \ \forall h \in \mathcal{C} = \{h : ||h_{S^{c}}||_{1} < \xi ||h_{S}||_{1}\}$

· Condition 5: Cone invertibility factor

$$\hat{\Sigma} = n^{-1} \sum_{i=1}^{n} (x_i - \bar{x})(x_i - \bar{x})^T$$

Condition 6: Tuning parameter

$$\lambda_{a} \in \left(\frac{1}{\eta}, M\right] \times \left(\frac{11\sqrt{L}}{3p'_{A}}\sqrt{\frac{\log p}{n}} + \delta_{n}\right)$$
$$\lambda_{b} \in \left(\frac{1}{\eta}, M\right] \times \left(\frac{11\sqrt{L}}{3p'_{B}}\sqrt{\frac{\log p}{n}} + \delta_{n}\right)$$

Asymptotic Normality

Theorem 1

Assume conditions 1 - 6 hold. Then

$$\sqrt{n}\left(\widehat{ATE}_{Lasso} - ATE\right) \overset{d}{
ightarrow} \mathcal{N}\left(0, \sigma^2\right),$$

$$\sigma^2 = \lim_{n \to \infty} \left[\frac{1 - p_A}{p_A} \sigma_{e^{(a)}}^2 + \frac{p_A}{1 - p_A} \sigma_{e^{(b)}}^2 + 2\sigma_{e^{(a)}e^{(b)}} \right]$$

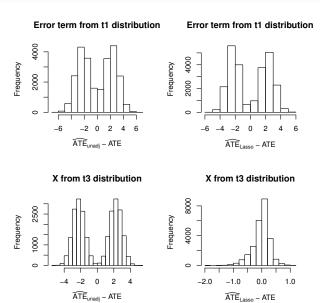
which is no greater than the asymptotic variance of the $\sqrt{n}\left(\widehat{ATE}_{\mathsf{unadj}} - ATE\right)$. The difference is $\frac{1}{p_A(1-p_A)}\Delta$.

$$\Delta = -\lim_{n \to \infty} \|X\beta_E\|_2^2 \le 0, \ \beta_E = (1 - p_A)\beta^{(a)} + p_A\beta^{(b)}$$

Our conditions are telling

- In simulation studies, we see that some moment conditions are necessary. That is, we find that the distribution of the Lasso adjusted estimator can be non-normal when these conditions do not hold.
- Nevertheless, in our simulation studies, Lasso adjusted estimator still has a smaller MSE than the unadjusted estimator

When moment conditions fail



Conservative variance estimate

As stated in Theorem 1, asymptotic variance

$$\sigma^2 = \lim_{n \to \infty} \left[\frac{1 - p_A}{p_A} \sigma_{e^{(a)}}^2 + \frac{p_A}{1 - p_A} \sigma_{e^{(b)}}^2 + 2\sigma_{e^{(a)}e^{(b)}} \right]$$

Let

$$\hat{\sigma}_{e^{(a)}}^{2} = \frac{1}{n_{A} - df^{(a)}} \sum_{i \in A} \left\{ a_{i} - \bar{a}_{A} - (x_{i} - \bar{x}_{A})^{T} \hat{\beta}^{(a)} \right\}^{2}$$

$$df^{(a)} = \hat{s}^{(a)} + 1 = ||\hat{\beta}^{(a)}||_0 + 1$$

Define

$$\hat{\sigma}^2 = \frac{n}{n_A} \hat{\sigma}_{e^{(a)}}^2 + \frac{n}{n_B} \hat{\sigma}_{e^{(b)}}^2$$

• We show that: $\hat{\sigma}^2$ is asymptotically conservative estimate of σ^2

PAC-man data: results

- OLS is computed by using only 59 main effect
- PAC has no significant average treatment effect
- regression-based methods provide 20% shorter intervals
- cv(Lasso) selects 24, 8 covariates for treatment and control respectively
- cv(Lasso+OLS) selects 4, 5 covariates for treatment and control respectively;

PAC-man data: simulation

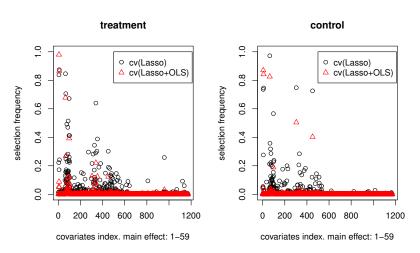
Matched on 59 main effects (ATE=-0.29), conduct 500 randomized experiments

Table 1: Results for the PAC-based simulations

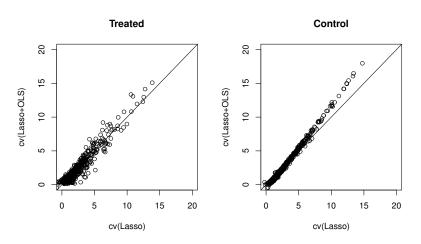
	Bias	SD	\sqrt{MSE}	Coverage	Length
Unadj	0.001	0.20	0.20	99%	1.06
OLS	0.002	0.18	0.18	99%	0.95
cv(Lasso)	0.001	0.17	0.17	99%	0.94
cv(Lasso+OLS)	0.000	0.17	0.17	99%	0.95

model size				
	treatment	control		
Unadj	0	0		
OLS	59	59		
cv(Lasso)	25	15		
cv(Lasso+OLS)	6	4		

Selection stability comparison: cv(Lasso) and cv(Lasso+OLS)



Adjustment value comparison: cv(Lasso) and cv(Lasso+OLS)



Bound (I): Massart concentration inequality

Proposition 1 (Massart concentration inequality for sampling without replacement)

Let $\{z_i, i=1,...,n\}$ be a finite population of real numbers. Let $A \subset \{i,...,n\}$ be a subset of deterministic size $|A| = n_A$ that is selected randomly without replacement. Define $p_A = n_A/n$, $\sigma^2 = n^{-1} \sum_{i=1}^n (z_i - \bar{z})^2$. Then, for any t > 0,

$$P(\bar{z}_A - \bar{z} \geq t) \leq \exp\left\{-\frac{p_A n_A t^2}{(1+\tau)^2 \sigma^2}\right\},$$

with $\tau = \min \{1/70, (3p_A)^2/70, (3-3p_A)^2/70\}.$

Summary of results

- Theoretical analysis of Lasso under Neyman-Rubin model, pointing to importance of moment conditions for covariates and error terms, and also cone invertibility for Lasso to work.
- Recommendation of cv(Lasso+OLS): much fewer covariates selected when compared with cv(Lasso) and with similar coverage and confidence interval length.
- Estimating Heterogeneous treatment effects?

PAC

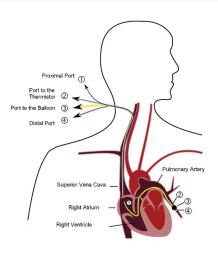


Figure 1: Pulmonary Artery Catheter (PAC), from Wikipedia

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Motivation example: PAC-man

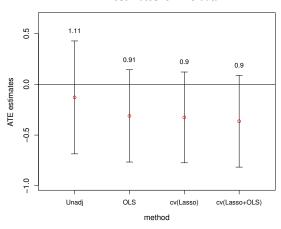
- Pulmonary Artery Catheter (PAC): monitoring device commonly inserted into critically ill patients
- Detecting complications, but invasive to patients and significant expenditure
- Question: does PAC have effect on patient survival?
- Observational study (Connors et al, 1996): PAC had an adverse effect on patient survival and led to increased cost of care

Motivation example: PAC-man data (Harvey et al, 2005)

- Randomized controlled trial on PAC, called PAC-Man: 65 UK intensive care units, 2001-2004
- 1013 patients: 506 treated and 507 control
- Outcome variable: quality-adjusted life years
- Observe many covariates including age, sex, some indicators, ...

PAC-man data: results

ATE estimates for PAC data



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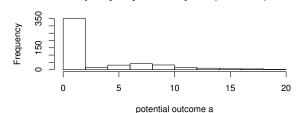
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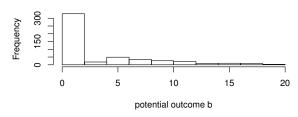
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quality-adjusted life years (treatment)



quality-adjusted life years (control)



Selected covariates for PAC data

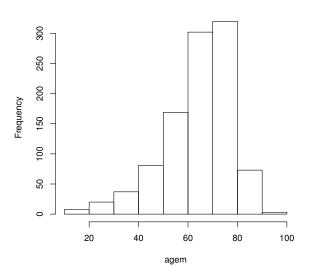
Table 2: Selected covariates for adjustment

Method	T/C	Selected covariates
cv(Lasso+OLS)	Т	age, p_death, age-age, age:p_death
cv(Lasso+OLS)	С	age, p_death, age-age, age:p_death, p_death:mech_vent
cv(Lasso)	Т	pac_rate, age, p_death, age-age, p_death-p_death,
		region:im_score, region:systemnew, pac_rate:age,
		pac_rate:p_death, pac_rate:systemnew, im_score:interactnew,
		age:glasgow, age:systemnew, interactnew:systemnew,
		pac_rate:creatinine, age:mech_vent, age:respiratory,
		age:p_death, age:creatinine, interactnew:systemnew,
		interactnew:mech_vent, interactnew:male, systemnew:male,
		p_death:mech_vent, glasgow:organ_failure
cv(Lasso)	С	age, p_death, age-age, unitsize:p_death, pac_rate:systemnew, age:p_death, interactnew:mech_vent, p_death:mech_vent

T: treated; C: control. Covariate meanings: age (patient's age); p_death (baseline probability of death); mech_vent (mechanical ventilation at admission); region (geographic region); pac_rate (PAC rate in unit); creatinine, respiratory, glasgow, interactnew, organ_failure, systemnew, im_score (various physiological indicators).

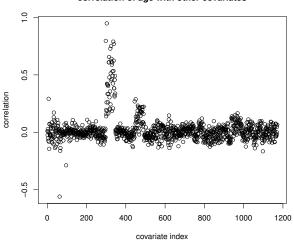
Histogram of the covariate: age

histogram of the covariate: age



Correlation

correlation of age with other covariates



Proof Sketch

Recall that

$$\widehat{\textit{ATE}}_{\text{Lasso}} = \left[\bar{a}_{\textit{A}} - \left(\bar{x}_{\textit{A}} - \bar{x}\right)^T \hat{\beta}^{(a)}\right] - \left[\bar{b}_{\textit{B}} - \left(\bar{x}_{\textit{B}} - \bar{x}\right)^T \hat{\beta}^{(b)}\right]$$

· Since,

$$\begin{split} &\sqrt{n}\left[\bar{a}_{A}-\left(\bar{x}_{A}-\bar{x}\right)^{T}\hat{\beta}^{(a)}\right]\\ &=\sqrt{n}\left[\bar{a}_{A}-\left(\bar{x}_{A}-\bar{x}\right)^{T}\beta^{(a)}\right]+\sqrt{n}(\bar{x}_{A}-\bar{x})^{T}(\beta^{(a)}-\hat{\beta}^{(a)}) \end{split}$$

We have²

$$\sqrt{n}(\widehat{ATE}_{Lasso} - ATE)$$

$$= \sqrt{n} \left\{ (\bar{a}_A - (\bar{x}_A - \bar{x})^T \beta^{(a)}) - (\bar{b}_B - (\bar{x}_B - \bar{x})^T \beta^{(b)}) - ATE \right\} (1)$$

$$+ \sqrt{n}(\bar{x}_A - \bar{x})^T (\beta^{(a)} - \hat{\beta}^{(a)}) - \sqrt{n}(\bar{x}_B - \bar{x})^T (\beta^{(b)} - \hat{\beta}^{(b)}) (2)$$

²Freedman DA (2008). On regression adjustments in experiments with several treatments. The Annals of Applied Statistics 2(1):176-196.

Proof Sketch

· Enough to show:

$$\sqrt{n}(\bar{x}_A - \bar{x})^T (\beta^{(a)} - \hat{\beta}^{(a)}) \rightarrow_{p} 0$$
 (3)

$$\sqrt{n}(\bar{x}_B - \bar{x})^T (\beta^{(b)} - \hat{\beta}^{(b)}) \rightarrow_p 0 \tag{4}$$

By Hölder inequality,

$$|\sqrt{n}(\bar{x}_{A} - \bar{x})^{T}(\beta^{(a)} - \hat{\beta}^{(a)})| \leq ||\sqrt{n}(\bar{x}_{A} - \bar{x})||_{\infty}||\beta^{(a)} - \hat{\beta}^{(a)}||_{1}$$

Need to control

$$(I): ||\bar{x}_A - \bar{x}||_{\infty} \text{ and } (II): ||\beta^{(a)} - \hat{\beta}^{(a)}||_{1}$$

Bound (I): Cont

Lemma 2

Under the fourth moment condition on the covariates, if we let $c_n = \frac{(1+\tau)L^{1/4}}{p_A} \sqrt{\frac{2\log p}{n}}$, then as $n \to \infty$,

$$P(\|\bar{x}_{A} - \bar{x}\|_{\infty} > c_{n}) \to 0 \tag{5}$$

Thus,
$$||\bar{x}_A - \bar{x}||_{\infty} = O_p\left(\sqrt{\frac{\log p}{n}}\right)$$
.

Bound (II): Consistency of Lasso

 Proceed with the similar procedure with Lasso in linear regression, we can show

$$||\beta^{(a)} - \hat{\beta}^{(a)}||_1 = O_p\left(\frac{s\sqrt{\log p}}{\sqrt{n}} + s\delta_n\right) = o_p\left(\frac{1}{\sqrt{\log p}}\right)$$

Both needs to show the following event happens w.h.p.

$$\mathcal{L} := ||\frac{2}{n_A} \sum_{i \in A} (e_i - \bar{e}_A) (x_i - \bar{x}_A)^T ||_{\infty} \le \frac{\lambda_a}{2}$$
 (6)

What are the differences?

Bound (II): Consistency of Lasso

Difference is here

- Linear regression
 - A is fixed at {1,2,...,n}
 - e_i's are i.i.d zero mean Gaussian or Subgaussian random variable
 - Concentration inequality for i.i.d Gaussian or Subgaussian
- Neyman-Rubin model
 - e_i's are fixed number
 - A is a random set with fixed size
 - Concentration (Massart) inequality for sampling with replacement

Bound (I) + Bound (II)

We have shown

$$||\beta^{(a)} - \hat{\beta}^{(a)}||_{1} = O_{p}\left(\frac{s\sqrt{\log p}}{\sqrt{n}} + s\delta_{n}\right) = o_{p}\left(\frac{1}{\sqrt{\log p}}\right)$$
$$||\bar{x}_{A} - \bar{x}||_{\infty} = O_{p}\left(\sqrt{\frac{\log p}{n}}\right)$$

Therefore,

$$||\sqrt{n}(\bar{x}_A - \bar{x})||_{\infty}||\beta^{(a)} - \hat{\beta}^{(a)}||_{1} = \sqrt{n}O_{\rho}\left(\sqrt{\frac{\log \rho}{n}}\right)o_{\rho}\left(\frac{1}{\sqrt{\log \rho}}\right)$$
$$= o_{\rho}(1)$$