

Supplementary Material for Bias-adjusted Kaplan-Meier Survival Curves for Marginal Treatment Effect in Observational Studies

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This paper has been submitted for consideration for publication in *JBS*

Theoretical Results

Here, we give the detailed and complete proof of the main results in Section 3 in the paper.

Let

$$\begin{aligned}
N_{1i}(t) &= I(Y_{1i} \leq t, \delta_{1i} = 1) & N_{1i}^{EL}(t) &= p_i D_i I(Y_{1i} \leq t, \delta_{1i} = 1) \\
Y_{1i}(t) &= I(Y_{1i} \geq t) & Y_{1i}^{EL}(t) &= p_i D_i I(Y_{1i} \geq t) \\
M_{1i}(t) &= N_{1i}(t) - \int_0^t Y_{1i}(s) d\Lambda_1(s) & M_{1i}^{EL}(t) &= N_{1i}^{EL}(t) - \int_0^t Y_{1i}^{EL}(s) d\Lambda_1(s) \\
\bar{N}_1(t) &= \sum_{i=1}^{n_1} I(Y_{1i} \leq t, \delta_{1i} = 1) & \bar{Y}_1(t) &= \sum_{i=1}^{n_1} I(Y_{1i} \geq t), \bar{Y}_1^{EL}(t) = \sum_{i=1}^n Y_{1i}^{EL}(t) \\
\bar{M}_1(t) &= \sum_{i=1}^{n_1} M_{1i}(t), \pi_1(t) = P(Y_1 \geq t) & \bar{M}_1^{EL}(t) &= \sum_{i=1}^n M_{1i}^{EL}(t) = \sum_{i=1}^n p_i D_i M_{1i}(t).
\end{aligned}$$

Similarly, if the subscript 1 is replaced with subscript 0 in all the aforementioned equations, it will denote the corresponding quantities of $S_0(t)$. For simplicity, we omit it here.

Let $\xi = (\xi_1, \xi_2)^T$, $\xi_1 = \frac{n_1 - \gamma^T a}{n}$, $\xi_2 = \frac{\gamma^T}{n}$ and $\psi(Z_i) = (1, \phi^T(Z_i))^T$. Then $p_i D_i = \frac{1}{n} \frac{D_i}{\xi^T \psi(Z_i)}$ and $p_i(1 - D_i) = \frac{1}{n} \frac{1 - D_i}{1 - \xi^T \psi(Z_i)}$. Let $\hat{\xi}$ be the solution of equations (2.1) and (2.2). Then we have

$$\frac{1}{n} \sum_{i=1}^n \left[\frac{D_i}{\hat{\xi}^T \psi(Z_i)} - \frac{1 - D_i}{1 - \hat{\xi}^T \psi(Z_i)} \right] \psi(Z_i) = 0. \quad (0.1)$$

LEMMA 1: $\hat{\xi}$ is the solution of equation (0.1), and ξ_0 is the value satisfies $\xi_0^T \psi(Z) = e(Z)$, which is the true value of the propensity score $P(D = 1|Z)$. Then under the conditions in Theorem 1, we have

$$\sqrt{n}(\hat{\xi} - \xi_0) = A^{-1} \frac{1}{\sqrt{n}} \sum_{i=1}^n \frac{(D_i - e(Z_i))\psi(Z_i)}{e(Z_i)(1 - e(Z_i))} + o_p(1),$$

where $A = E \left[\frac{\psi(Z_i)\psi^T(Z_i)}{e(Z_i)(1 - e(Z_i))} \right]$.

Proof. Since $\hat{\xi}$ is the solution of (0.1), using a Taylor series expansion at ξ_0 , we have

$$\frac{1}{n} \sum_{i=1}^n \left[\frac{D_i}{e(Z_i)} - \frac{1 - D_i}{1 - e(Z_i)} \right] \psi(Z_i) - E \left[\frac{\psi(Z_i)\psi^T(Z_i)}{e(Z_i)(1 - e(Z_i))} \right] (\hat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}) = 0$$

Then the results can be obtained immediately.

LEMMA 2: When $S_1(t) > 0$, we have

$$\frac{\widehat{S}_1(t)}{S_1(t)} = 1 - \int_0^t \frac{\widehat{S}_1(s-)}{S_1(s)} \left\{ \frac{d\overline{N}_1^{EL}(s)}{\overline{Y}_1^{EL}(s)} - d\Lambda_1(s) \right\}, \quad (0.2)$$

where $\Lambda_1(\cdot)$ is the cumulative hazard function of T_1 .

Lemma 2 can be easily proved by the formulas for integration by parts and the differential of a reciprocal, similar to the proof of Theorem 3.2.3 in Fleming and Harrington (1991).

Proof of Theorem 1

Proof. Based on (0.2) in Lemma 2, we can obtain

$$\begin{aligned} \frac{S_1(t) - \widehat{S}_1(t)}{S_1(t)} &= \int_0^t \frac{\widehat{S}_1(s-)}{S_1(s)} \frac{I(\overline{Y}_1^{EL}(s) > 0)}{\overline{Y}_1^{EL}(s)} dM_1^{EL}(t) + \int_0^t \frac{\widehat{S}_1(s-)}{S_1(s)} I(\overline{Y}_1^{EL}(s) = 0) d\Lambda_1(s) \\ &= \sum_{i=1}^n \int_0^t \frac{\widehat{S}_1(s-)}{S_1(s)} \frac{I(\overline{Y}_1^{EL}(s) > 0)}{\overline{Y}_1^{EL}(s)} p_i D_i dM_{1i}(t) + \mathcal{B}_1(t), \end{aligned}$$

where $\mathcal{B}_1(t) = \int_0^t \frac{\widehat{S}_1(s-)}{S_1(s)} I(\overline{Y}_1^{EL}(s) = 0) d\Lambda_1(s)$.

Let $\tau = \inf\{s : \overline{Y}_1^{EL}(s) = 0\}$ and $\mathcal{B}(t) = \mathcal{B}_1(t)S_1(t)$. So $\forall t > \tau$, $\overline{Y}_1^{EL}(t) = \sum_{i=1}^n p_i D_i I(Y_{1i} \geq t) = 0$.

And we can obtain

$$\begin{aligned} \mathcal{B}(t) &= S_1(t) \int_\tau^t \frac{\widehat{S}_1(s-)}{S_1(s)} d\Lambda_1(s) I(\tau < t) \\ &= S_1(t) \widehat{S}_1(\tau) \frac{S_1(\tau) - S_1(t)}{S_1(\tau) S_1(t)} I(\tau < t) \\ &= \widehat{S}_1(\tau) \left(1 - \frac{S_1(t)}{S_1(\tau)} \right) I(\tau < t), \end{aligned}$$

therefore, as $n \rightarrow \infty$,

$$\begin{aligned} E[\mathcal{B}(t)] &= E\left\{ \widehat{S}_1(\tau) \left(1 - \frac{S_1(t)}{S_1(\tau)} \right) I(\tau < t) \right\} \\ &\leq E\left\{ I(\tau \leq t) (1 - S(t)) \right\} = (1 - S(t)) (1 - \pi_1(t))^{n_1} \rightarrow 0, \end{aligned}$$

which implies that as $n \rightarrow \infty$, $\mathcal{B}(t) \xrightarrow{P} 0$.

The derivatives of the consistency theorem can be divided into two steps. First, we show that for any fixed $u \in (0, \infty)$ such that as $n_1 \rightarrow \infty$, $\overline{Y}_1(u) \xrightarrow{P} \infty$, then $\sup_{0 \leq s \leq u} |\widehat{S}_1(s) -$

$|S_1(s)| \xrightarrow{P} 0$, as $n \rightarrow \infty$. Second, we show that if $t \in (0, \infty]$ is such that for any $u < t$, $\bar{Y}_1(u) \xrightarrow{P} \infty$, as $n \rightarrow \infty$. Then $\sup_{0 \leq s \leq t} |\hat{S}_1(s) - S_1(s)| \xrightarrow{P} 0$, as $n \rightarrow \infty$.

Under the conditions in this theorem, for any $t \leq u$, $S_1(t) > 0$. Followed by Lemma 2, we have as $n \rightarrow \infty$,

$$P \left\{ \frac{S_1(t) - \hat{S}_1(t)}{S_1(t)} = U(t) \quad \text{on} \quad [0, u] \right\},$$

where

$$U(t) = \sum_{i=1}^n \int_0^t \frac{\hat{S}_1(s-)}{S_1(s)} \frac{I(\bar{Y}_1^{EL}(s) > 0)}{\bar{Y}_1^{EL}(s)} p_i D_i dM_{1i}(t). \quad (0.3)$$

Note that $\xi_0 \psi(Z_i) = e(Z_i)$, based on (2.3), we have

$$\begin{aligned} q_i = p_i D_i &= \frac{1}{n} \frac{D_i}{\hat{\xi} \psi(Z_i)} \\ &= \frac{1}{n} \frac{D_i}{e(Z_i)} - \frac{1}{n} \frac{D_i \psi^T(Z_i)}{e^2(Z_i)} (\hat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}), \end{aligned} \quad (0.4)$$

hence (0.3) becomes

$$\begin{aligned} U(t) &= \frac{1}{n} \sum_{i=1}^n \int_0^t \frac{\hat{S}_1(s-)}{S_1(s)} \frac{I(\bar{Y}_1^{EL}(s) > 0)}{\bar{Y}_1^{EL}(s)} \frac{D_i}{e(Z_i)} dM_{1i}(s) - \\ &\quad \frac{1}{n} \sum_{i=1}^n \int_0^t \frac{\hat{S}_1(s-)}{S_1(s)} \frac{I(\bar{Y}_1^{EL}(s) > 0)}{\bar{Y}_1^{EL}(s)} \frac{D_i \psi^T(Z_i)}{e^2(Z_i)} dM_{1i}(s) (\hat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}) \\ &=: U_1(t) - U_2(t) (\hat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}), \end{aligned}$$

where “=:” means “defined as” throughout our paper.

Combining equation (0.4) and Lemma 1, as $n \rightarrow \infty$, we can obtain

$$\frac{\hat{S}_1(s-)}{S_1(s)} \frac{I(\bar{Y}_1^{EL}(s) > 0)}{n \bar{Y}_1^{EL}(s)} \frac{D_i}{e(Z_i)} = \frac{1}{n} \frac{D_i}{e(Z_i)} \frac{\hat{S}_1^{(1)}(s-)}{S_1(s)} \frac{I(\bar{Y}_1(s) > 0)}{\pi_1(s)},$$

where $\hat{S}_1^{(1)}(s-) = \prod_{u < s} \left\{ 1 - \frac{\sum \frac{D_i}{e(Z_i)} \Delta N_{1i}(u)}{\sum \frac{D_i}{e(Z_i)} Y_{1i}(u)} \right\}$.

Let $U_1^*(t) = \frac{1}{n_1} \sum_{i=1}^{n_1} \int_0^t \frac{\hat{S}_1(s-)}{S_1(s)} \frac{I(\bar{Y}_1(s) > 0)}{\pi_1(s)} dM_{1i}(s)$. Followed the assumption about D and (Y_1, C) , we have $\sup_{0 \leq s \leq t} |U_1(t) - U_1^*(t)| \xrightarrow{P} 0$.

The process $M_{1i}(t)$ is a martingale and the process $\frac{\hat{S}_1^{(1)}(s-)}{S_1(s)} \frac{I(\bar{Y}_1(s) > 0)}{\pi_1(s)}$ is a predictable and bounded process, by Corollary 3.4.1 in Fleming and Harrington (1991), $\forall \epsilon, \eta > 0$, for any

$0 \leq t < u$, we can obtain

$$\begin{aligned} P\left\{\sup_{0 \leq s \leq t} |U_1^*(s)|^2 \geq \epsilon\right\} &\leq \frac{\eta}{\epsilon} + P\left\{\int_0^t \frac{\widehat{S}_1^{(1)}(s-)^2 \sum_{i=1}^{n_1} I(Y_{1i} \geq s)}{S_1^2(s)} I(\overline{Y}_1(s) > 0) d\Lambda_1(s) \geq \eta\right\} \\ &\leq \frac{\eta}{\epsilon} + P\left\{\frac{\Lambda_1(t)}{n S_1^2(t) \pi_1(t)} \geq \eta\right\}. \end{aligned}$$

Therefore, as $n \rightarrow \infty$, $\sup_{0 \leq s < u} |U_1(s)| \xrightarrow{P} 0$. Under the conditions in this theorem, as $n \rightarrow \infty$, $U_2(s) = O_p(1)$. On the other side, we have showed that $\|\widehat{\xi} - \xi_0\| = O_p(n^{-\frac{1}{2}})$, therefore as $n \rightarrow \infty$, $\sup_{0 \leq s < u} |U_2(s)(\widehat{\xi} - \xi_0)| \xrightarrow{P} 0$. The first step of consistency theory is finished.

Next, we are going to show that if $u = \sup\{t : \pi_1(t) > 0\}$. Then $\sup_{0 \leq s \leq u} |\widehat{S}_1(s) - S_1(s)| \xrightarrow{P} 0$, as $n \rightarrow \infty$.

So it implies to find a t_0 such that as $n \rightarrow \infty$, $\sup_{t_0 \leq s \leq u} |\widehat{S}_1(s) - S_1(s)| \xrightarrow{P} 0$.

For $t_0 \leq s \leq u$,

$$S_1(u) \leq S_1(s) \leq S_1(t_0)$$

$$\widehat{S}_1(u) \leq \widehat{S}_1(s) \leq \widehat{S}_1(t_0),$$

hence, after simple algorithms, we have

$$\sup_{t_0 \leq s \leq u} |\widehat{S}_1(s) - S_1(s)| \leq |\widehat{S}_1(t_0) - S_1(t_0)| + 2|S_1(t_0) - S_1(u)| + |\widehat{S}_1(u) - S_1(u)|.$$

We have shown that $|\widehat{S}_1(t_0) - S_1(t_0)| \xrightarrow{P} 0$, as $n \rightarrow \infty$ in the first step, and the second term in last equation tends to 0 by the continuity of $S_1(\cdot)$ if we choose some t_0 close to u . Therefore, it suffices to show that as $n \rightarrow \infty$,

$$|\widehat{S}_1(u) - S_1(u)| \xrightarrow{P} 0.$$

We discuss the two cases: $S_1(u) = 0$ and $S_1(u) > 0$, separately.

When $S_1(u) = 0$, there exists a t_0 such that $|S_1(t_0) - S_1(u)| \xrightarrow{P} 0$, which implies

$|S_1(t_0)| \xrightarrow{P} 0$, note that $u > t_0$, then

$$\begin{aligned} |\widehat{S}_1(u) - S_1(u)| &\leq |\widehat{S}_1(u)| + |S_1(u)| \\ &\leq |\widehat{S}_1(t_0)| + |S_1(t_0)| \\ &\leq |\widehat{S}_1(t_0) - S_1(t_0)| + 2|S_1(t_0)|. \end{aligned}$$

Together with the results in the first step, the first term in last equation converges in probability to 0 as $n \rightarrow \infty$.

When $S_1(u) > 0$, since

$$|\widehat{S}_1(u) - S_1(u)| \leq |\widehat{S}_1(u) - \widehat{S}_1(t_0)| + |\widehat{S}_1(t_0) - S_1(t_0)| + |S_1(t_0) - S_1(u)|,$$

so it suffices to show that as $n \rightarrow \infty$, $|\widehat{S}_1(u) - \widehat{S}_1(t_0)| \xrightarrow{P} 0$.

For any $\epsilon > 0$,

$$\begin{aligned} P\left\{|\widehat{S}_1(t) - \widehat{S}_1(t_0)| > \epsilon\right\} &= P\left\{\left|\int_{t_0}^t I\{\overline{Y}^{EL}(t) > 0\} \widehat{S}_1(s-) \frac{d\overline{N}_1^{EL}(s)}{\overline{Y}_1^{EL}(s)}\right| > \epsilon\right\} \\ &\leq P\left\{\left|\int_{t_0}^t I\{\overline{Y}(t) > 0\} \left\{\frac{d\overline{N}_1^{EL}(s)}{\overline{Y}_1^{EL}(s)} - d\Lambda_1(s)\right\}\right| > \epsilon/2\right\} + (0.5) \\ &\quad P\left\{\left|\int_{t_0}^t d\Lambda_1(s)\right| > \epsilon/2\right\}. \end{aligned}$$

By the continuity of $S_1(t)$, we can find a t_0 such that

$$\left|\int_{t_0}^t d\Lambda_1(s)\right| = \left|-\int_{t_0}^t \frac{dS_1(t)}{S_1(t)}\right| \leq |S_1(t) - S_1(t_0)| < \epsilon/4.$$

Similar to the proof of uniformly consistency of the process $U_1(t)$ in the first step, we can easily show that as $n \rightarrow \infty$

$$P\left\{\left|\int_{t_0}^t I\{\overline{Y}(t) > 0\} \left\{\frac{d\overline{N}_1^{EL}(s)}{\overline{Y}_1^{EL}(s)} - d\Lambda_1(s)\right\}\right| > \epsilon/2\right\} < \epsilon/2.$$

Here we omit the detailed proof. Hence as $n \rightarrow \infty$, $|\widehat{S}_1(t) - \widehat{S}_1(t_0)| \xrightarrow{P} 0$ when $S_1(t) > 0$.

We have finished the proof of the consistency theorem.

Proof of Theorem 2

Proof. By the derivations of the consistency theorem and the consistency of $\widehat{S}_1(t)$, we can obtain:

$$\begin{aligned} \frac{S_1(t) - \widehat{S}_1(t)}{S_1(t)} &= \frac{1}{n} \sum_{i=1}^n \int_0^t \frac{1}{\pi_1(s)} \frac{D_i}{e(Z_i)} dM_{1i}(s) \\ &- B_1 \cdot (\widehat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}), \end{aligned}$$

where $B_1 = E \left[\int_0^t \frac{1}{\pi_1(s)} \frac{D_i \psi^T(Z_i)}{e^2(Z_i)} dM_{1i}(s) \right]$.

Together with Lemma 1, we have

$$\begin{aligned} \frac{S_1(t) - \widehat{S}_1(t)}{S_1(t)} &= \frac{1}{n} \sum_{i=1}^n \int_0^t \frac{1}{\pi_1(s)} \frac{D_i}{e(Z_i)} dM_{1i}(s) \\ &- B_1 A^{-1} \frac{1}{n} \sum_{i=1}^n \frac{(D_i - e(Z_i)) \psi(Z_i)}{e(Z_i)(1 - e(Z_i))} + o_p(n^{-\frac{1}{2}}). \end{aligned} \quad (0.6)$$

By the central limit theory of independent identically random variables, we can obtain for any $t \in \mathcal{I}_1$, $\sqrt{n}S_1(t)^{-1}(\widehat{S}_1(t) - S_1(t))$ converges to a normal distribution $N(0, V_1)$, where

$$V_1 = E \left[\frac{D}{e(Z)} \int_0^t \frac{1}{\pi_1(s)} dM_1(s) \right]^2 - B_1 A^{-1} B_1^T.$$

Note that the two terms in equation (0.6) are correlated. We have finished the proof of this theorem.

Proof of Theorem 4

Proof. Followed by the proof of Theorem 2, for a fix time point t , we have

$$\begin{aligned} \widehat{\Delta}(t) - \Delta(t) &= \widehat{S}_1(t) - \widehat{S}_0(t) - (S_1(t) - S_0(t)) \\ &= S_1(t) \frac{1}{n} \sum_{i=1}^n \frac{D_i}{e(Z_i)} \int_0^t \frac{1}{\pi_1(s)} dM_{1i}(s) - S_1(t) B_1 (\widehat{\xi} - \xi_0) - \\ &\quad S_0(t) \frac{1}{n} \sum_{i=1}^n \frac{1 - D_i}{1 - e(Z_i)} \int_0^t \frac{1}{\pi_0(s)} dM_{0i}(s) + S_0(t) B_0 (\widehat{\xi} - \xi_0) + o_p(n^{-\frac{1}{2}}). \end{aligned}$$

Then the large sample properties can be easily obtained.

Additional Simulation Results

The simulation setups are the same as simulation study 2 except that the sample size is set to $n = 100$ and $n = 300$.

Results of simulation study 2 with $n = 100$

method	para	true	est.hat	bias	se	sd	RMSE
KM	$S_1(t)$	0.504	0.572	0.135	0.078	0.074	0.156
KM	$S_0(t)$	0.265	0.221	0.165	0.065	0.062	0.177
KM	$\Delta(t)$	0.239	0.351	0.467	0.103	0.097	0.478
IPW incorrect	$S_1(t)$	0.504	0.529	0.05	0.076	0.075	0.091
IPW incorrect	$S_0(t)$	0.265	0.254	0.041	0.073	0.07	0.084
IPW incorrect	$\Delta(t)$	0.239	0.275	0.149	0.103	0.103	0.181
IPW correct	$S_1(t)$	0.504	0.514	0.02	0.073	0.074	0.075
IPW correct	$S_0(t)$	0.265	0.265	0.002	0.074	0.072	0.074
IPW correct	$\Delta(t)$	0.239	0.248	0.039	0.099	0.104	0.107
EL	$S_1(t)$	0.504	0.508	0.008	0.071	0.068	0.071
EL	$S_0(t)$	0.265	0.272	0.027	0.076	0.067	0.081
EL	$\Delta(t)$	0.239	0.236	0.012	0.097	0.091	0.098

Results of simulation study 2 with $n = 300$

method	para	true	est.hat	bias	se	sd	RMSE	
KM	$S_1(t)$	0.504	0.569	0.129	0.044	0.044	0.136	67.5
KM	$S_0(t)$	0.265	0.224	0.155	0.039	0.037	0.16	76.7
KM	$\Delta(t)$	0.239	0.345	0.443	0.059	0.057	0.447	53.7
IPW incorrect	$S_1(t)$	0.504	0.524	0.04	0.043	0.044	0.059	92.2
IPW incorrect	$S_0(t)$	0.265	0.256	0.033	0.043	0.042	0.054	92.8
IPW incorrect	$\Delta(t)$	0.239	0.268	0.121	0.058	0.061	0.134	92.5
IPW correct	$S_1(t)$	0.504	0.509	0.01	0.042	0.043	0.043	95.1
IPW correct	$S_0(t)$	0.265	0.267	0.007	0.043	0.043	0.044	94.1
IPW correct	$\Delta(t)$	0.239	0.242	0.013	0.056	0.061	0.057	96.6
EL	$S_1(t)$	0.504	0.506	0.004	0.041	0.041	0.041	94.5
EL	$S_0(t)$	0.265	0.269	0.016	0.043	0.041	0.046	93.1
EL	$\Delta(t)$	0.239	0.237	0.01	0.054	0.055	0.055	95.2

References

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