

AN ADDITIVE-MULTIPLICATIVE MEAN MODEL FOR MARKER DATA CONTINGENT ON RECURRENT EVENT WITH AN INFORMATIVE TERMINAL EVENT

Miao Han¹, Xinyuan Song², Liuquan Sun³ and Lei Liu⁴

¹*School of Statistics and Management, Shanghai University of Finance and Economics,
Shanghai, China*

²*Department of Statistics, The Chinese University of Hong Kong*

³*Institute of Applied Mathematics, Academy of Mathematics and Systems Science,
Chinese Academy of Sciences, Beijing, China*

⁴*Department of Preventive Medicine, Northwestern University, Chicago, USA*

Supplementary Material

This supplementary file contains the regularity conditions (C1)-(C6) and the proofs of Theorems 1-3 in Sections 3 and 4 of the paper.

S1 Proofs of Theorems

In order to study the asymptotic properties of the proposed estimators, we need the following regularity conditions:

- (C1) $\{m_i(\cdot), N_i(\cdot), T_i, \delta_i, X_i, W_i\}, i = 1, \dots, n$, are independent identically distributed.
- (C2) $N(\tau)$, X and W are bounded almost surely, and $P(T \geq \tau) > 0$.
- (C3) The link functions $g_\gamma(\cdot)$, $g_\beta(\cdot)$ and $g_\zeta(\cdot)$ are twice continuously differentiable with $g_\gamma(\cdot) > 0$
- (C4) The weight functions $W(t)$ and $Q(t)$ have bounded variation and converge to deterministic functions $w(t)$ and $q(t)$, respectively, in probability, uniformly in $t \in [0, \tau]$.
- (C5) Ω is nonsingular, where

$$\Omega = E \left[\int_0^\tau \{Z_i(t) - \bar{z}^D(t)\}^{\otimes 2} dN_i^D(t) \right]$$

and $\bar{z}^D(t)$ is the limit of $\bar{Z}^D(t; \eta_0)$.

- (C6) A is nonsingular, where

$$A = \begin{pmatrix} A_{11} & A_{12} \\ 0 & A_{22} \end{pmatrix},$$

$$A_{11} = E \left[\int_0^\tau w(t) \{Z_i - \bar{z}(t, Z_i)\} Y_i(t) \right. \\ \left. \times \begin{pmatrix} \{X_i \dot{g}_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \bar{x}(t, Z_i)\} dH(t, \log \Lambda_0(t) + \eta'_0 Z_i) \\ W_i \dot{g}_\zeta(\zeta'_0 W_i) dN_i(t) - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \bar{w}(t, Z_i) \end{pmatrix} \right],$$

$$A_{12} = E \left[\int_0^\tau w(t) \{Z_i - \bar{z}(t, Z_i)\} Y_i(t) \right. \\ \left. \times \{Z_i g_\beta(\beta'_0 X_i) \dot{g}_\gamma(\gamma'_0 Z_i) - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \bar{z}^\dagger(t, Z_i)\}' dH(t, \log \Lambda_0(t) + \eta'_0 Z_i) \right],$$

$$A_{22} = E \left[\int_0^\tau q(t) \{Z_i - \bar{z}^N(t, Z_i)\} Y_i(t) \{Z_i \dot{g}_\gamma(\gamma'_0 Z_i) - \bar{z}^*(t, Z_i) g_\gamma(\gamma'_0 Z_i)\}' d\mu_0(t; v_2) \right],$$

and $\bar{z}(t, Z_i)$, $\bar{x}(t, Z_i)$, $\bar{w}(t, Z_i)$, $\bar{z}^\dagger(t, Z_i)$ and $\bar{z}^N(t, Z_i)$ are the limits of $\bar{Z}_i(t; \beta_0, \gamma_0)$, $\bar{X}_i(t; \beta_0, \gamma_0)$, $\bar{W}_i(t; \theta_0, \gamma_0)$, $\bar{Z}_i^\dagger(t; \beta_0, \gamma_0)$, and $\bar{Z}_i^N(t; \gamma_0)$ conditional on Z_i , respectively.

Proof of Theorem 1. Define

$$\Psi_i(t, Z; \eta, \Lambda) = I\{\log \Lambda(T_i) + \eta' Z_i \geq \log \Lambda(t) + \eta' Z \geq \log \Lambda(t) + \eta' Z_i\},$$

$$d\bar{N}(t, Z; \eta, \Lambda) = \frac{\sum_{j=1}^n dN_j(t) \Psi_j(t, Z; \eta, \Lambda)}{\sum_{j=1}^n g_\gamma(\gamma'_0 Z_j) \Psi_j(t, Z; \eta, \Lambda)},$$

and

$$d\bar{N}_0(t, Z; \eta, \Lambda) = \frac{E[dN_j(t) \Psi_j(t, Z; \eta, \Lambda) | Z]}{E[g_\gamma(\gamma'_0 Z_j) \Psi_j(t, Z; \eta, \Lambda) | Z]}.$$

Let $\Psi_i(t, Z) = \Psi_i(t, Z; \eta_0, \Lambda_0)$. It then follows from the functional delta method (van der Vaart and Wellner, 1996, Theorem 3.9.4, p.374) that

$$\begin{aligned} & n^{-1/2} \sum_{i=1}^n \int_0^\tau q(t) \{Z_i - \bar{z}^N(t, Z_i)\} Y_i(t) g_\gamma(\gamma'_0 Z_i) \{d\bar{N}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{N}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} \\ &= n^{-1/2} \sum_{i=1}^n \int_0^\tau \int q(t) \{z - \bar{z}^N(t, z)\} I(y \geq t) \frac{g_\gamma(\gamma'_0 z) \Psi_i(t, z)}{E[g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)]} dP(z, y) dN_i(t) \\ & \quad - n^{-1/2} \sum_{i=1}^n \int \left[\int_0^\tau q(t) \{z - \bar{z}^N(t, z)\} I(y \geq t) g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z) \right. \\ & \quad \left. \times \frac{E[\Psi_i(t, z) dN_i(t)]}{(E[g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)])^2} \right] g_\gamma(\gamma'_0 z) dP(z, y) + o_p(1), \end{aligned} \quad (S.1)$$

where $P(z, y)$ is the joint probability measure of (Z_i, T_i) . In addition, according to Fleming and Harrington (1991, p.299), we have

$$\hat{\eta} - \eta_0 = \Omega^{-1} n^{-1} \sum_{i=1}^n \int_0^\tau \{Z_i - \bar{z}^D(t)\} dM_i^D(t) + o_p(n^{-1/2}),$$

and

$$\hat{\Lambda}_0(t) - \Lambda_0(t) = n^{-1} \sum_{i=1}^n \int_0^t \frac{dM_i^D(u)}{s^{(0)}(u; \eta_0)} - \int_0^t \bar{z}^D(u)' d\Lambda_0(u) (\hat{\eta} - \eta_0) + o_p(n^{-1/2}),$$

where

$$M_i^D(t) = N_i^D(t) - \int_0^t Y_i(u) \exp(\eta'_0 Z_i) d\Lambda_0(u),$$

S1. PROOFS OF THEOREMS3

and $s^{(0)}(t; \eta_0)$ is the limit of $S^{(0)}(t; \eta_0)$. Let $dR_\eta(t, Z)$ and $dR_\Lambda(t, Z)$ be the derivative and the Hadamard derivative of $d\bar{N}_0(t, Z; \eta_0, \Lambda_0)$ with respect to η and Λ , respectively. Then by the functional delta method, we obtain

$$\begin{aligned} n^{-1/2} \sum_{i=1}^n \int_0^\tau q(t) \{Z_i - z^N(t, Z_i)\} Y_i(t) g_\gamma(\gamma'_0 Z_i) \{d\bar{N}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{N}_0(t, Z_i; \eta_0, \Lambda_0)\} \\ = n^{-1/2} \sum_{i=1}^n \int_0^\tau \left[B_1 \Omega^{-1} \{Z_i - \bar{z}^D(t)\} + \frac{C_1(t)}{s^{(0)}(t; \eta_0)} \right] dM_i^D(t) + o_p(1), \end{aligned} \quad (S.2)$$

where

$$C_1(t) = E \left[\int_t^\tau q(u) \{Z_i - \bar{z}^N(u, Z_i)\} Y_i(u) g_\gamma(\gamma'_0 Z_i) dR_\Lambda(u, Z_i) \right],$$

and

$$B_1 = E \left[\int_0^\tau q(t) \{Z_i - \bar{z}^N(t, Z_i)\} Y_i(t) g_\gamma(\gamma'_0 Z_i) \left\{ dR_\eta(t, Z_i) - \left(\int_0^t \bar{z}^D(u)' d\Lambda_0(u) \right) dR_\Lambda(t, Z_i) \right\} \right].$$

Note that

$$\begin{aligned} n^{-1/2} U_\gamma(\gamma_0) &= \sum_{i=1}^n \int_0^\tau Q(t) \{Z_i - \bar{Z}_i^N(t; \gamma_0)\} Y_i(t) [dN_i(t) - g_\gamma(\gamma'_0 Z_i) d\bar{N}_0(t, Z_i; \eta_0, \Lambda_0) \\ &\quad - g_\gamma(\gamma'_0 Z_i) \{d\bar{N}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{N}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} \\ &\quad - g_\gamma(\gamma'_0 Z_i) \{d\bar{N}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{N}_0(t, Z_i; \eta_0, \Lambda_0)\}] + o_p(1). \end{aligned} \quad (S.3)$$

Since $\sup_{i,t} |\bar{Z}_i^N(t; \gamma_0) - z^N(t, Z_i)| \rightarrow 0$ in probability, it follows from (S.1), (S.2) and (S.3) that

$$n^{-1/2} U_\gamma(\gamma_0) = n^{-1/2} \sum_{i=1}^n \varphi_i + o_p(1), \quad (S.4)$$

where

$$\begin{aligned} \varphi_i &= \int_0^\tau q(t) \{Z_i - \bar{z}^N(t, Z_i)\} Y_i(t) [dN_i(t) - g_\gamma(\gamma'_0 Z_i) d\bar{N}_0(t, Z_i; \eta_0, \Lambda_0)] \\ &\quad - \int_0^\tau \int q(t) \{z - \bar{z}^N(t, z)\} I(y \geq t) \frac{g_\gamma(\gamma'_0 z) \Psi_i(t, z)}{E[g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)]} dP(z, y) dN_i(t) \\ &\quad + \int \left[\int_0^\tau q(t) \{z - \bar{z}^N(t, z)\} I(y \geq t) g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z) \right. \\ &\quad \times \left. \frac{E[\Psi_i(t, z) dN_i(t)]}{(E[g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)])^2} \right] g_\gamma(\gamma'_0 z) dP(z, y) \\ &\quad - \int_0^\tau \left[B_1 \Omega^{-1} \{Z_i - \bar{z}^D(t)\} + \frac{C_1(t)}{s^{(0)}(t; \eta_0)} \right] dM_i^D(t). \end{aligned}$$

Define $dM_i(t) = \{m_i(t) - g_\zeta(\zeta'_0 W_i)\} dN_i(t)$,

$$d\bar{M}(t, Z; \eta, \Lambda) = \frac{\sum_{j=1}^n dM_j(t) \Psi_j(t, Z; \eta, \Lambda)}{\sum_{j=1}^n g_\beta(\beta'_0 X_j) g_\gamma(\gamma'_0 Z_j) \Psi_j(t, Z; \eta, \Lambda)},$$

and

$$d\bar{M}_0(t, Z; \eta, \Lambda) = \frac{E[dM_j(t)\Psi_j(t, Z; \eta, \Lambda)|Z]}{E[g_\beta(\beta'_0 X_j)g_\gamma(\gamma'_0 Z_j)\Psi_j(t, Z; \eta, \Lambda)|Z]}.$$

Similarly, it follows from the functional delta method that

$$\begin{aligned} & n^{-1/2} \sum_{i=1}^n \int_0^\tau w(t) \{Z_i - \bar{z}(t, Z_i)\} Y_i(t) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \{d\bar{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} \\ &= n^{-1/2} \sum_{i=1}^n \int_0^\tau \int w(t) \{z - \bar{z}(t, z)\} I(y \geq t) \frac{g_\beta(\beta'_0 x) g_\gamma(\gamma'_0 z) \Psi_i(t, z)}{E[g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)]} dF(z, x, y) dM_i(t) \\ &\quad - n^{-1/2} \sum_{i=1}^n \int \left[\int_0^\tau w(t) \{z - \bar{z}(t, z)\} I(y \geq t) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z) \right. \\ &\quad \left. \times \frac{E[\Psi_i(t, z) dM_i(t)]}{(E[g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(t, z)])^2} \right] g_\beta(\beta'_0 x) g_\gamma(\gamma'_0 z) dF(z, x, y) + o_p(1), \end{aligned} \quad (S.5)$$

where $F(z, x, y)$ is the joint probability measure of (Z_i, X_i, T_i) . Let $dR_\eta^*(t, Z)$ and $dR_\Lambda^*(t, Z)$ be the derivative and the Hadamard derivative of $d\bar{M}_0(t, Z; \eta_0, \Lambda_0)$ with respect to η and Λ , respectively. Then in a similar manner, we get

$$\begin{aligned} & n^{-1/2} \sum_{i=1}^n \int_0^\tau w(t) \{Z_i - \bar{z}(t, Z_i)\} Y_i(t) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \{d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \eta_0, \Lambda_0)\} \\ &= n^{-1/2} \sum_{i=1}^n \int_0^\tau \left[B_2 \Omega^{-1} \{Z_i - \bar{z}^D(t)\} + \frac{C_2(t)}{s^{(0)}(t; \eta_0)} \right] dM_i^D(t) + o_p(1), \end{aligned} \quad (S.6)$$

where

$$C_2(t) = E \left[\int_t^\tau q(u) \{Z_i - \bar{z}(u, Z_i)\} Y_i(u) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) dR_\Lambda^*(u, Z_i) \right],$$

and

$$\begin{aligned} B_2 &= E \left[\int_0^\tau q(t) \{Z_i - \bar{z}(t, Z_i)\} Y_i(t) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \right. \\ &\quad \left. \times \left\{ dR_\eta^*(t, Z_i) - \left(\int_0^t \bar{z}^D(u)' d\Lambda_0(u) \right) dR_\Lambda^*(t, Z_i) \right\} \right]. \end{aligned}$$

Note that

$$\begin{aligned} U_\theta(\theta_0, \gamma_0) &= \sum_{i=1}^n \int_0^\tau W(t) \{Z_i - \bar{Z}_i(t; \beta_0, \gamma_0)\} Y_i(t) \left[dM_i(t) - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) d\bar{M}_0(t, Z_i; \eta_0, \Lambda_0) \right. \\ &\quad - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \{d\bar{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} \\ &\quad \left. - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \{d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \eta_0, \Lambda_0)\} \right] + o_p(1), \end{aligned} \quad (S.7)$$

and $\sup_{i,t} |\bar{Z}_i(t; \beta_0, \gamma_0) - \bar{z}(t, Z_i)| \rightarrow 0$ in probability. Thus, it follow from (S.5), (S.6) and (S.7) that

$$n^{-1/2} U_\theta(\theta_0, \gamma_0) = n^{-1/2} \sum_{i=1}^n \xi_i + o_p(1), \quad (S.8)$$

where

$$\begin{aligned}
\xi_i &= \int_0^\tau w(t)\{Z_i - \bar{z}(t, Z_i)\}Y_i(t)[dM_i(t) - g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)d\bar{M}_0(t, Z_i; \eta_0, \Lambda_0)] \\
&\quad - \int_0^\tau \int w(t)\{z - \bar{z}(t, z)\}I(y \geq t) \frac{g_\beta(\beta'_0 x)g_\gamma(\gamma'_0 z)\Psi_i(t, z)}{E[g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z)]} dF(z, x, y)dM_i(t) \\
&\quad + \int \left[\int_0^\tau w(t)\{z - \bar{z}(t, z)\}I(y \geq t)g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z) \right. \\
&\quad \times \left. \frac{E[\Psi_i(t, z)dM_i(t)]}{(E[g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z)])^2} \right] g_\beta(\beta'_0 x)g_\gamma(\gamma'_0 z)dF(z, x, y) \\
&\quad - \int_0^\tau \left[B_2 \Omega^{-1}\{Z_i - \bar{z}^D(t)\} + \frac{C_2(t)}{s^{(0)}(t; \eta_0)} \right] dM_i^D(t).
\end{aligned}$$

Note that $-n^{-1}\partial U_\theta(\theta_0; \gamma_0)/\partial\theta$, $-n^{-1}\partial U_\theta(\theta_0; \gamma_0)/\partial\gamma$ and $-n^{-1}\partial U_\gamma(\gamma_0)/\partial\gamma$ converge in probability to A_{11} , A_{12} and A_{22} , respectively. It then follows from (S.4), (S.8) and the Taylor expansion that

$$\begin{aligned}
n^{1/2} \begin{pmatrix} \hat{\theta} - \theta_0 \\ \hat{\gamma} - \gamma_0 \end{pmatrix} &= A^{-1}n^{-1/2} \begin{pmatrix} U_\theta(\theta_0; \gamma_0) \\ U_\gamma(\gamma_0) \end{pmatrix} + o_p(1), \\
&= A^{-1}n^{-1/2} \sum_{i=1}^n \begin{pmatrix} \xi_i \\ \varphi_i \end{pmatrix} + o_p(1),
\end{aligned}$$

which implies that $n^{1/2}(\hat{\theta} - \theta_0)$ and $n^{1/2}(\hat{\gamma} - \gamma_0)$ have asymptotically a joint normal distribution with mean zero and covariance matrix $A^{-1}\Sigma(A')^{-1}$, where $\Sigma = E[(\xi'_i, \varphi'_i)' \otimes 2]$.

Proof of Theorem 2. In view of the consistency of $\hat{\theta}$, $\hat{\gamma}$, $\hat{\eta}$ and $\hat{\Lambda}_0(t)$, using the Theorem 3.6.13 of van der Vaart and Wellner (1996), we obtain that conditional on the observed data,

$$\begin{aligned}
\Phi_1^* &= \sum_{i=1}^n G_i \int_0^\tau w(t)\{Z_i - \bar{z}(t, Z_i)\}Y_i(t)[dM_i(t) \\
&\quad - g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)d\bar{M}_0(t, Z_i; \eta_0, \Lambda_0)] + o_p(n^{1/2}). \tag{S.9}
\end{aligned}$$

Likewise, it can be shown that conditional on the observed data,

$$\begin{aligned}
\Phi_2^* &= - \sum_{i=1}^n G_i \int_0^\tau \int w(t)\{z - \bar{z}(t, z)\}I(y \geq t) \frac{g_\beta(\beta'_0 x)g_\gamma(\gamma'_0 z)\Psi_i(t, z)}{E[g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z)]} dF(z, x, y)dM_i(t) \\
&\quad + \sum_{i=1}^n G_i \int \left[\int_0^\tau w(t)\{z - \bar{z}(t, z)\}I(y \geq t)g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z) \right. \\
&\quad \times \left. \frac{E[\Psi_i(t, z)dM_i(t)]}{(E[g_\beta(\beta'_0 X_i)g_\gamma(\gamma'_0 Z_i)\Psi_i(t, z)])^2} \right] g_\beta(\beta'_0 x)g_\gamma(\gamma'_0 z)dF(z, x, y) + o_p(n^{1/2}). \tag{S.10}
\end{aligned}$$

Let $d\hat{M}_i(t) = \{m_i(t) - g_\zeta(\zeta' W_i)\}dN_i(t)$, and

$$d\hat{M}(t, Z; \eta, \Lambda) = \frac{\sum_{j=1}^n d\hat{M}_j(t)\Psi_j(t, Z; \eta, \Lambda)}{\sum_{j=1}^n g_\beta(\hat{\beta}' X_j)g_\gamma(\hat{\gamma}' Z_j)\Psi_j(t, Z; \eta, \Lambda)}.$$

Thus,

$$\Phi_3^* = \sum_{i=1}^n \int_0^\tau W(t) \{Z_i - \bar{Z}_i(t; \hat{\beta}, \hat{\gamma})\} Y_i(t) g_\beta(\hat{\beta}' X_i) g_\gamma(\hat{\gamma}' Z_i) \{d\hat{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\hat{M}(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*)\}.$$

It can be checked that

$$\begin{aligned} & d\hat{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\hat{M}(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*) = \{d\hat{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} \\ & - \{d\hat{M}(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*) - d\bar{M}_0(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*)\} + \{d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) - d\bar{M}_0(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*)\}. \end{aligned}$$

Similarly to (S.1), we obtain

$$\begin{aligned} & n^{-1/2} \sum_{i=1}^n \int_0^\tau W(t) \{Z_i - \bar{Z}_i(t; \hat{\beta}, \hat{\gamma})\} Y_i(t) g_\beta(\hat{\beta}' X_i) g_\gamma(\hat{\gamma}' Z_i) \left[\{d\hat{M}(t, Z_i; \hat{\eta}, \hat{\Lambda}_0) \right. \\ & \left. - d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)\} - \{d\hat{M}(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*) - d\bar{M}_0(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*)\} \right] = o_p(1). \end{aligned}$$

From an argument similar to that in the proof of (S.6), we have that conditional on the observed data,

$$\begin{aligned} & n^{-1/2} \sum_{i=1}^n \int_0^\tau W(t) \{Z_i - \bar{Z}_i(t; \hat{\beta}, \hat{\gamma})\} Y_i(t) g_\beta(\hat{\beta}' X_i) g_\gamma(\hat{\gamma}' Z_i) [d\bar{M}_0(t, Z_i; \hat{\eta}^*, \hat{\Lambda}_0^*) - d\bar{M}_0(t, Z_i; \hat{\eta}, \hat{\Lambda}_0)] \\ & = n^{-1/2} G_i \int_0^\tau \left[B_2 \Omega^{-1} \{Z_i - \bar{z}^D(t)\} + \frac{C_2(t)}{s^{(0)}(t; \eta_0)} \right] dM_i^D(t) + o_p(1). \end{aligned} \quad (\text{S.11})$$

It follows from (S.9), (S.10) and (S.11) that conditional on the observed data,

$$\hat{U}_1 = n^{-1/2} (\Phi_1^* + \Phi_2^* + \Phi_3^*) = n^{-1/2} \sum_{i=1}^n G_i \xi_i + o_p(1).$$

In a similar manner,

$$\hat{U}_2 = n^{-1/2} (\Phi_4^* + \Phi_5^* + \Phi_6^*) = n^{-1/2} \sum_{i=1}^n G_i \varphi_i + o_p(1).$$

Thus, by using theorem 3.6.13 of van der Vaart and Wellner (1996), $E_G[\hat{U}^{\otimes 2}]$ converges in probability to Σ .

Proof of Theorem 3. Note that

$$\begin{aligned} \mathcal{F}(z, t) &= n^{-1/2} \sum_{i=1}^n \int_0^t I(Z_i \leq z) Y_i(u) \left[\{m_i(u) - g_\zeta(\zeta_0' W_i)\} dN_i(t) \right. \\ & \left. - g_\beta(\beta_0' X_i) g_\gamma(\gamma_0' Z_i) \{d\bar{m}_i(u; \beta_0, \gamma_0) - d\bar{g}_i(u; \theta_0, \gamma_0)\} \right] \\ & - \Gamma_1(z, t)' n^{1/2} (\hat{\theta} - \theta_0) - \Gamma_2(z, t)' n^{1/2} (\hat{\gamma} - \gamma_0) + o_p(1), \end{aligned} \quad (\text{S.12})$$

where $\Gamma_1(z, t)$ and $\Gamma_2(z, t)$ are the limit of $\hat{\Gamma}_1(z, t)$ and $\hat{\Gamma}_2(z, t)$, respectively. By following similar arguments as in the proof of Theorem 1, the first term on the right-hand side of (S.12) equals

$$n^{-1/2} \sum_{i=1}^n \phi_i(z, t) + o_p(1), \quad (\text{S.13})$$

where

$$\begin{aligned} \phi_i(z, t) = & \int_0^t I(Z_i \leq z) Y_i(u) [dM_i(u) - g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) d\bar{M}_0(u, Z_i; \eta_0, \Lambda_0)] \\ & - \int_0^t \int I(Z_i \leq z) I(y \geq u) \frac{g_\beta(\beta'_0 x) g_\gamma(\gamma'_0 s) \Psi_i(u, s)}{E[g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(u, s)]} dF(s, x, y) dM_i(u) \\ & + \int \left[\int_0^t I(Z_i \leq z) I(y \geq u) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(u, s) \right. \\ & \times \left. \frac{E[\Psi_i(u, s) dM_i(u)]}{(E[g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \Psi_i(u, s)])^2} \right] g_\beta(\beta'_0 x) g_\gamma(\gamma'_0 s) dF(s, x, y) \\ & - B_3(t, z) \Omega^{-1} \int_0^\tau \{Z_i - \bar{z}^D(u)\} dM_i^D(u) - \int_0^t \frac{C_3(t, u, z)}{s^{(0)}(u; \eta_0)} dM_i^D(u), \end{aligned}$$

$$\begin{aligned} B_3(t, z) = & E \left[\int_0^t I(Z_i \leq z) Y_i(u) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) \left\{ dR_\eta^*(u, Z_i) \right. \right. \\ & \left. \left. - \left(\int_0^u \bar{z}^D(s)' d\Lambda_0(s) \right) dR_\Lambda^*(u, Z_i) \right\} \right], \end{aligned}$$

and

$$C_3(t, u, z) = E \left[\int_u^t I(Z_i \leq z) Y_i(s) g_\beta(\beta'_0 X_i) g_\gamma(\gamma'_0 Z_i) dR_\Lambda^*(s, Z_i) \right].$$

Thus, it follows from (S.12), (S.13) and Theorem 1 that

$$\mathcal{F}(z, t) = n^{-1/2} \sum_{i=1}^n \left[\phi_i(z, t) - \Gamma(z, t)' A^{-1}(\xi'_i, \varphi'_i)' \right] + o_p(1), \quad (\text{S.14})$$

where $\Gamma(z, t) = (\Gamma_1(z, t)', \Gamma_2(z, t)')'$. By the multivariate central limit theorem, $\hat{\mathcal{F}}(z, t)$ converges in finite-dimensional distribution to a zero-mean Gaussian process. The first term of (S.14) is tight because any function of bounded variation can be written as the difference of two increasing functions. Note that $\Gamma(z, t)$ is a deterministic function, and ξ_i and φ_i do not involve t . Thus the second term is also tight. Hence $\mathcal{F}(z, t)$ is tight and converges weakly to a zero-mean Gaussian process. By the arguments analogous to those in the proof of Theorem 2, we obtain that $\mathcal{F}(z, t)$ can be approximated by the zero-mean Gaussian process $\hat{\mathcal{F}}(z, t)$ given by (7).

REFERENCES

- Fleming T. R. and Harrington D. P. (1991). *Counting Processes and Survival Analysis*. New York: Wiley.
- van der Vaart, A. W. and Wellner, J. A. (1996). *Weak Convergence and Empirical Processes*. New York: Springer.