

Supplemental Materials for “Generalized Ordinary Differential Equation Models”

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A Proof of Theorem

Theorem 1. Assume that there exists a $\lambda > 0$ such that $\delta = O(N^{-\lambda})$, then under Assumptions A1-A8, we have $\hat{\alpha} \xrightarrow{p} \alpha_0$.

Proof: Denote

$$\begin{aligned} & \tilde{M}_n(\alpha) \\ = & \frac{1}{N} L(\alpha; \mathbf{y}, \tilde{\mathbf{x}}, \mathbf{z}) = \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \ell(\alpha; y_{iks}, \tilde{\mathbf{x}}_i, \mathbf{z}_{ik}) \\ = & \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \left[\frac{y_{iks} \cdot b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}), \boldsymbol{\beta}) - b\{b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}), \boldsymbol{\beta})\}}{a(\phi)} + c(y_{iks}, \phi) \right], \end{aligned}$$

$$\begin{aligned} & M_n(\alpha) \\ = & \frac{1}{N} L(\alpha; \mathbf{y}, \mathbf{x}, \mathbf{z}) = \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \ell(\alpha; y_{iks}, \mathbf{x}_i, \mathbf{z}_{ik}) \\ = & \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \left[\frac{y_{iks} \cdot b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \boldsymbol{\theta}), \boldsymbol{\beta}) - b\{b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \boldsymbol{\theta}), \boldsymbol{\beta})\}}{a(\phi)} + c(y_{iks}, \phi) \right], \end{aligned}$$

and

$$M(\alpha) = \ell(\alpha; \mathbf{y}, \mathbf{x}, \mathbf{z}) = \frac{y \cdot b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}, \mathbf{x}(t, \boldsymbol{\theta}), \boldsymbol{\beta}) - b\{b'^{-1} \circ g^{-1} \circ g^*(\mathbf{z}, \mathbf{x}(t, \boldsymbol{\theta}), \boldsymbol{\beta})\}}{a(\phi)} + c(y, \phi).$$

First, we claim that $E_0[M(\alpha)]$ reaches its unique maximum at $\alpha = \alpha_0$. In fact, by the Jensen's inequality and the convexity of function $-\log x$, we have

$$E_0[M(\alpha)] - E_0[M(\alpha_0)] = E_0[\ell(\alpha; \mathbf{y}, \mathbf{x}, \mathbf{z}) - \ell(\alpha_0; \mathbf{y}, \mathbf{x}, \mathbf{z})] \leq 0,$$

i.e., $\alpha = \alpha_0$ is one maximum point of function $E_0[M(\alpha)]$. Moreover, from Assumption A6, it follows that it is the unique maximum point. Thus the above claim holds.

Next, with the Taylor expansion, for any $\alpha_1, \alpha_2 \in \Theta \times \mathcal{B} \times \Phi$, we can easily obtain

$$|M_n(\alpha_1) - M_n(\alpha_2)| \leq C(\mathbf{y}, \mathbf{x}, \mathbf{z}) \|\alpha_1 - \alpha_2\|, \quad (\text{A.1})$$

where $C(\mathbf{y}, \mathbf{x}, \mathbf{z})$ is bounded. By the weak law of large numbers, it follows that for every fixed α , $M_n(\alpha) - E_0[M(\alpha)] \xrightarrow{p} 0$. Then under Assumption A1, by Corollary 2 in Andrews (1987), we have that $\sup_{\alpha} |M_n(\alpha) - E_0[M(\alpha)]| \xrightarrow{p} 0$. Thus it follows that $M_n(\hat{\alpha}) - E_0[M(\hat{\alpha})] \xrightarrow{p} 0$ and $M_n(\alpha_0) - E_0[M(\alpha_0)] \xrightarrow{p} 0$.

Based on the formulation of ℓ in Section 3.1, for a variate x , $\frac{\partial}{\partial x} \ell(\boldsymbol{\alpha}; y_{iks}, x, \mathbf{z}_{ik})$ is continuous and bounded. In addition, the function $\mathbf{x}_i = \mathbf{x}(t_i, \boldsymbol{\theta})$ is continuous and bounded. Thus $\frac{\partial}{\partial x} \ell(\boldsymbol{\alpha}; y_{iks}, x, \mathbf{z}_{ik})|_{x=\mathbf{x}(t_i, \boldsymbol{\theta})}$ is $O_p(1)$. It is similar for $\tilde{\mathbf{x}}_i$. Then by Lemma 1 in [Xue et al. \(2010\)](#) and the first-order Taylor expansion, it follows

$$\begin{aligned}
\tilde{M}_n(\boldsymbol{\alpha}) &= \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \ell(\boldsymbol{\alpha}; y_{iks}, \tilde{\mathbf{x}}_i, \mathbf{z}_{ik}) \\
&= \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \ell(\boldsymbol{\alpha}; y_{iks}, \mathbf{x}_i + O(N^{-\lambda(p \wedge 4)}), \mathbf{z}_{ik}) \\
&= \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \ell(\boldsymbol{\alpha}; y_{iks}, \mathbf{x}_i, \mathbf{z}_{ik}) + O_p(N^{-\lambda(p \wedge 4)}) \\
&= M_n(\boldsymbol{\alpha}) + O_p(N^{-\lambda(p \wedge 4)}).
\end{aligned}$$

Then

$$\tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\boldsymbol{\alpha}_0)] \leq \tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\hat{\boldsymbol{\alpha}})] = M_n(\hat{\boldsymbol{\alpha}}) + O_p(N^{-\lambda(p \wedge 4)}) - E_0[M(\hat{\boldsymbol{\alpha}})].$$

and

$$\tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\boldsymbol{\alpha}_0)] \geq \tilde{M}_n(\boldsymbol{\alpha}_0) - E_0[M(\boldsymbol{\alpha}_0)] = M_n(\boldsymbol{\alpha}_0) + O_p(N^{-\lambda(p \wedge 4)}) - E_0[M(\boldsymbol{\alpha}_0)].$$

Hence $\tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\boldsymbol{\alpha}_0)] \xrightarrow{p} 0$, when $N \rightarrow \infty$ and $\lambda > 0$. Thus

$$|E_0[M(\hat{\boldsymbol{\alpha}})] - E_0[M(\boldsymbol{\alpha}_0)]| \leq |\tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\hat{\boldsymbol{\alpha}})]| + |\tilde{M}_n(\hat{\boldsymbol{\alpha}}) - E_0[M(\boldsymbol{\alpha}_0)]| \xrightarrow{p} 0. \quad (\text{A.2})$$

Now, we claim that $\hat{\boldsymbol{\alpha}}$ weakly converges to $\boldsymbol{\alpha}_0$. Otherwise, for sequence $\{\hat{\boldsymbol{\alpha}}\}$ in the compact subset $\Theta \times \mathcal{B} \times \Phi$, there exists a convergent subsequence, $\{\hat{\boldsymbol{\alpha}}_{n_k}\}$, such that $\hat{\boldsymbol{\alpha}}_{n_k} \xrightarrow{p} \boldsymbol{\alpha}_*$ and $\boldsymbol{\alpha}_* \neq \boldsymbol{\alpha}_0$. From the continuous mapping theorem, it follows that $E_0[M(\hat{\boldsymbol{\alpha}}_{n_k})] \xrightarrow{p} E_0[M(\boldsymbol{\alpha}_*)]$. From [\(A.2\)](#), we have $E_0[M(\hat{\boldsymbol{\alpha}}_{n_k})] \xrightarrow{p} E_0[M(\boldsymbol{\alpha}_0)]$. Then $E_0[M(\boldsymbol{\alpha}_*)] = E_0[M(\boldsymbol{\alpha}_0)]$. Since $\boldsymbol{\alpha}_0$ is the unique maximum point of $E_0[M(\boldsymbol{\alpha})]$, we have $\boldsymbol{\alpha}_* = \boldsymbol{\alpha}_0$, which is contradictory to $\boldsymbol{\alpha}_* \neq \boldsymbol{\alpha}_0$. Thus $\hat{\boldsymbol{\alpha}} \xrightarrow{p} \boldsymbol{\alpha}_0$. \square

Theorem 2. (i) For $\delta = O(N^{-\lambda})$ with $\lambda > 1/(p \wedge 4)$ where p is the order of the numerical method, under Assumptions A1-A10, we have that $\sqrt{N}(\hat{\alpha} - \alpha_0) \xrightarrow{d} \mathcal{N}(0, \mathbf{H}^{-1})$;

(ii) For $\delta = O(N^{-\lambda})$ with $0 < \lambda \leq 1/(p \wedge 4)$, under Assumptions A1-A10, we have that $\sqrt{N}(\hat{\alpha} - \tilde{\alpha}) \xrightarrow{d} \mathcal{N}(0, \tilde{\mathbf{H}}^{-1})$ with $\|\tilde{\alpha} - \alpha_0\| = O_p(\delta^{(p \wedge 4)/2}) = O_p(N^{-\lambda(p \wedge 4)/2})$ and $\|\tilde{\mathbf{H}} - \mathbf{H}\| = O_p(\delta^{(p \wedge 4)/2}) = O_p(N^{-\lambda(p \wedge 4)/2})$.

Proof: For the proof of Part (i), it suffices to verify conditions of Theorem 3.1 in [Newey and McFadden \(1994\)](#) on asymptotic normality for a M-estimator. Denote

$$\begin{aligned} \tilde{G}_n(\alpha) &= \begin{pmatrix} \frac{\partial \tilde{M}_n(\alpha)}{\partial \theta} \\ \frac{\partial \tilde{M}_n(\alpha)}{\partial \beta} \\ \frac{\partial \tilde{M}_n(\alpha)}{\partial \phi} \end{pmatrix} = \begin{pmatrix} \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \tilde{\mathbf{x}}(t_i, \theta), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}(t_i, \theta), \beta)}{\partial \tilde{\mathbf{x}}} \Big|_{\tilde{\mathbf{x}}=\tilde{\mathbf{x}}(t_i, \theta)} \frac{\partial \tilde{\mathbf{x}}(t_i, \theta)}{\partial \theta} \\ \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \tilde{\mathbf{x}}(t_i, \theta), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}(t_i, \theta), \beta)}{\partial \beta} \\ \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \tilde{\mathbf{x}}(t_i, \theta), \mathbf{z}_{ik})}{\partial \phi} \end{pmatrix}, \\ G_n(\alpha) &= \begin{pmatrix} \frac{\partial M_n(\alpha)}{\partial \theta} \\ \frac{\partial M_n(\alpha)}{\partial \beta} \\ \frac{\partial M_n(\alpha)}{\partial \phi} \end{pmatrix} = \begin{pmatrix} \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \mathbf{x}(t_i, \theta), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \theta), \beta)}{\partial \mathbf{x}} \Big|_{\mathbf{x}=\mathbf{x}(t_i, \theta)} \frac{\partial \mathbf{x}(t_i, \theta)}{\partial \theta} \\ \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \mathbf{x}(t_i, \theta), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \theta), \beta)}{\partial \beta} \\ \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\alpha; y_{iks}, \mathbf{x}(t_i, \theta), \mathbf{z}_{ik})}{\partial \phi} \end{pmatrix}, \\ G(\alpha) &= \begin{pmatrix} \frac{\partial E_0[M(\alpha)]}{\partial \theta} \\ \frac{\partial E_0[M(\alpha)]}{\partial \beta} \\ \frac{\partial E_0[M(\alpha)]}{\partial \phi} \end{pmatrix} = \begin{pmatrix} E_0 \left[\frac{\partial \ell(\alpha; y, \mathbf{x}(t, \theta), \mathbf{z})}{\partial g^*} \frac{\partial g^*(\mathbf{z}, \mathbf{x}, \beta)}{\partial \mathbf{x}} \Big|_{\mathbf{x}=\mathbf{x}(t, \theta)} \frac{\partial \mathbf{x}(t, \theta)}{\partial \theta} \right] \\ E_0 \left[\frac{\partial \ell(\alpha; y, \mathbf{x}(t, \theta), \mathbf{z})}{\partial g^*} \frac{\partial g^*(\mathbf{z}, \mathbf{x}(t, \theta), \beta)}{\partial \beta} \right] \\ E_0 \left[\frac{\partial \ell(\alpha; y, \mathbf{x}(t, \theta), \mathbf{z})}{\partial \phi} \right] \end{pmatrix}, \end{aligned}$$

where $\frac{\partial \ell}{\partial g^*} = \frac{\partial}{\partial g^*} \ell(\cdot, g^*, \cdot) \Big|_{g^*=g^*(\mathbf{z}, \mathbf{x}, \beta)}$ with $g^*(\mathbf{z}, \mathbf{x}, \beta)$ defined in (2.5). Since $\hat{\alpha}$ and α_0 are maximum points of $\tilde{M}_n(\alpha)$ and $E_0[M(\alpha)]$, respectively, $\tilde{G}_n(\hat{\alpha}) = 0$ and $G(\alpha_0) = 0$.

First, we verify the result that $\sqrt{N}[\tilde{G}_n(\alpha_0) - G(\alpha_0)] \xrightarrow{d} \mathcal{N}(0, \mathbf{H})$ with \mathbf{H} defined by (4.18). For fixed t , by the Landau-Kolmogorov inequality between different derivatives of a function, we have $\left\| \frac{\partial \tilde{\mathbf{x}}(t, \theta)}{\partial \theta} - \frac{\partial \mathbf{x}(t, \theta)}{\partial \theta} \right\|_{\infty} \leq C \left\| \frac{\partial^2 \tilde{\mathbf{x}}(t, \theta)}{\partial \theta \partial \theta^T} - \frac{\partial^2 \mathbf{x}(t, \theta)}{\partial \theta \partial \theta^T} \right\|_{\infty}^{1/2} \left\| \tilde{\mathbf{x}}(t, \theta) - \mathbf{x}(t, \theta) \right\|_{\infty}^{1/2} \leq C' \left\| \tilde{\mathbf{x}}(t, \theta) - \mathbf{x}(t, \theta) \right\|_{\infty}^{1/2}$ for two constants C and C' , where the second inequality holds because of the uniform boundedness of both $\frac{\partial^2 \mathbf{x}(t, \theta)}{\partial \theta \partial \theta^T}$ and $\frac{\partial^2 \tilde{\mathbf{x}}(t, \theta)}{\partial \theta \partial \theta^T}$ under Assumptions A7-A8, and $\|\Upsilon(\theta)\|_{\infty} = \sup_{\theta} |\Upsilon(\theta)|$ is the supremum norm of a function Υ . Based on $\sup_{t \in [t_0, T]} \left\| \tilde{\mathbf{x}}(t, \theta) - \mathbf{x}(t, \theta) \right\|_{\infty} = O(N^{-\lambda(p \wedge 4)})$ from Lemma 1 in [Xue et al. \(2010\)](#), it follows that $\left\| \frac{\partial \tilde{\mathbf{x}}(t, \theta)}{\partial \theta} - \frac{\partial \mathbf{x}(t, \theta)}{\partial \theta} \right\| = O(N^{-\lambda(p \wedge 4)/2})$.

Then we have

$$\begin{aligned}
& \sqrt{N}[\tilde{G}_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] \\
&= \left(\begin{aligned} & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}, \boldsymbol{\beta}_0)}{\partial \tilde{\mathbf{x}}} \Big|_{\tilde{\mathbf{x}}=\tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0)} \frac{\partial \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0)}{\partial \boldsymbol{\theta}_0} \\ & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0), \boldsymbol{\beta}_0)}{\partial \boldsymbol{\beta}_0} \\ & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \tilde{\mathbf{x}}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial \phi_0} \end{aligned} \right) \\
&= \left(\begin{aligned} & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \left[\frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} + O_p(N^{-\lambda(p \wedge 4)}) \right] \left[\frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}, \boldsymbol{\beta}_0)}{\partial \mathbf{x}} \Big|_{\mathbf{x}=\mathbf{x}(t_i, \boldsymbol{\theta}_0)} + O_p(N^{-\lambda(p \wedge 4)}) \right] \\ & \quad \times \left[\frac{\partial \mathbf{x}(t_i, \boldsymbol{\theta}_0)}{\partial \boldsymbol{\theta}_0} + O_p(N^{-\lambda(p \wedge 4)/2}) \right] \\ & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \left[\frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} + O_p(N^{-\lambda(p \wedge 4)}) \right] \left[\frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \boldsymbol{\beta}_0)}{\partial \boldsymbol{\beta}_0} + O_p(N^{-\lambda(p \wedge 4)}) \right] \\ & \quad \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \left[\frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial \phi_0} + O_p(N^{-\lambda(p \wedge 4)}) \right] \end{aligned} \right) \\
&= \left(\begin{aligned} & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}, \boldsymbol{\beta}_0)}{\partial \mathbf{x}} \Big|_{\mathbf{x}=\mathbf{x}(t_i, \boldsymbol{\theta}_0)} \frac{\partial \mathbf{x}(t_i, \boldsymbol{\theta}_0)}{\partial \boldsymbol{\theta}_0} \\ & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial g^*} \frac{\partial g^*(\mathbf{z}_{ik}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \boldsymbol{\beta}_0)}{\partial \boldsymbol{\beta}_0} \\ & \frac{1}{\sqrt{N}} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} \frac{\partial \ell(\boldsymbol{\alpha}_0; y_{iks}, \mathbf{x}(t_i, \boldsymbol{\theta}_0), \mathbf{z}_{ik})}{\partial \phi_0} \end{aligned} \right) + O_p(N^{-\lambda(p \wedge 4)/2+1/2}) \\
&= \sqrt{N}[G_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] + O_p(N^{-\lambda(p \wedge 4)/2+1/2}).
\end{aligned}$$

When $\lambda > 1/(p \wedge 4)$, $O_p(N^{-\lambda(p \wedge 4)/2+1/2}) = o_p(1)$. So for the above expression, we have $\sqrt{N}[\tilde{G}_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] = \sqrt{N}[G_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] + o_p(1)$. Based on the special structure of ξ_{ik} and ϕ in (2.4), it is easy to follow that $E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}, \mathbf{z})}{\partial(\boldsymbol{\theta}^T, \boldsymbol{\beta}^T)} \frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}, \mathbf{z})}{\partial \phi} \right] \Big|_{\boldsymbol{\alpha}=\boldsymbol{\alpha}_0} = \mathbf{0}$, that is, the dispersion parameter ϕ is orthogonal to parameters $\boldsymbol{\theta}$ and $\boldsymbol{\beta}$ in the sense of [Cox and Reid \(1987\)](#). From the standard central limit theorem, we have $\sqrt{N}[G_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] \xrightarrow{d} \mathcal{N}(0, \mathbf{J})$, where

$$\mathbf{J} = \begin{pmatrix} E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}, \mathbf{z})}{\partial(\boldsymbol{\theta}^T, \boldsymbol{\beta}^T)^T} \frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}, \mathbf{z})}{\partial(\boldsymbol{\theta}^T, \boldsymbol{\beta}^T)} \right] \Big|_{\boldsymbol{\alpha}=\boldsymbol{\alpha}_0} & \mathbf{0} \\ \mathbf{0}^T & E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}, \mathbf{z})}{\partial \phi} \right]^2 \Big|_{\boldsymbol{\alpha}=\boldsymbol{\alpha}_0} \end{pmatrix}$$

Thus it follows that $\sqrt{N}[\tilde{G}_n(\boldsymbol{\alpha}_0) - G(\boldsymbol{\alpha}_0)] \xrightarrow{d} \mathcal{N}(0, \mathbf{J})$.

Next, by similar arguments to those in the proof of Theorem 1, it follows that $\sup_{\boldsymbol{\alpha} \in \Delta} \left\| \frac{\partial^2 M_n(\boldsymbol{\alpha})}{\partial \boldsymbol{\alpha} \partial \boldsymbol{\alpha}^T} + \mathbf{H}(\boldsymbol{\alpha}) \right\| \xrightarrow{P} 0$ with a neighborhood Δ of $\boldsymbol{\alpha}_0$. By Lemma 1 in [Xue et al. \(2010\)](#), it follows that $\sup_{\boldsymbol{\alpha} \in \Delta} \left\| \frac{\partial^2 M_n(\boldsymbol{\alpha})}{\partial \boldsymbol{\alpha} \partial \boldsymbol{\alpha}^T} - \frac{\partial^2 M_n(\boldsymbol{\alpha})}{\partial \boldsymbol{\alpha} \partial \boldsymbol{\alpha}^T} \right\| \xrightarrow{P} 0$. Then we have $\sup_{\boldsymbol{\alpha} \in \Delta} \left\| \frac{\partial^2 M_n(\boldsymbol{\alpha})}{\partial \boldsymbol{\alpha} \partial \boldsymbol{\alpha}^T} + \mathbf{H}(\boldsymbol{\alpha}) \right\| \xrightarrow{P} 0$. By Theorem 1, Assumptions A9 and A10, and $\mathbf{H} = \mathbf{J}$, all conditions of Theorem 3.1 in [Newey and McFadden \(1994\)](#) are satisfied, thus Theorem 2(i) holds with a variance-covariance matrix $(-\mathbf{H})^{-1} \mathbf{J} (-\mathbf{H})^{-1} = \mathbf{J}^{-1} = \mathbf{H}^{-1}$.

Now, we consider the proof of Theorem 2(ii). Define

$$\begin{aligned}
\tilde{M}(\boldsymbol{\alpha}) &= \ell(\boldsymbol{\alpha}; y, \tilde{\mathbf{x}}, \mathbf{z}) \\
&= \frac{y \cdot b^{-1} \circ g^{-1} \circ g^*(\mathbf{z}, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \boldsymbol{\beta}) - b\{b^{-1} \circ g^{-1} \circ g^*(\mathbf{z}, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \boldsymbol{\beta})\}}{a(\phi)} + c(y, \phi),
\end{aligned}$$

and

$$\tilde{G}(\boldsymbol{\alpha}) = \begin{pmatrix} \frac{\partial \tilde{M}(\boldsymbol{\alpha})}{\partial \boldsymbol{\theta}} \\ \frac{\partial \tilde{M}(\boldsymbol{\alpha})}{\partial \boldsymbol{\beta}} \\ \frac{\partial \tilde{M}(\boldsymbol{\alpha})}{\partial \phi} \end{pmatrix} = \begin{pmatrix} E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} \frac{\partial g^*(z, \tilde{\mathbf{x}}, \boldsymbol{\beta})}{\partial \tilde{\mathbf{x}}} \Big|_{\tilde{\mathbf{x}} = \tilde{\mathbf{x}}(t, \boldsymbol{\theta})} \frac{\partial \tilde{\mathbf{x}}(t, \boldsymbol{\theta})}{\partial \boldsymbol{\theta}} \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} \frac{\partial g^*(z, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \boldsymbol{\beta})}{\partial \boldsymbol{\beta}} \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \tilde{\mathbf{x}}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial \phi} \right] \end{pmatrix}.$$

Since $E_0[\tilde{M}(\boldsymbol{\alpha})]$ reaches its maximum at $\boldsymbol{\alpha} = \tilde{\boldsymbol{\alpha}}$, then the first-order derivative of $E_0[\tilde{M}(\boldsymbol{\alpha})]$ at $\tilde{\boldsymbol{\alpha}}$ equals to 0, i.e., $\tilde{G}(\tilde{\boldsymbol{\alpha}}) = 0$. Then similar to the proof of Case (i) above, we have

$$\begin{aligned} \tilde{G}(\boldsymbol{\alpha}) &= \begin{pmatrix} E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} + O_p(N^{-\lambda(p \wedge 4)}) \right] \left[\frac{\partial g^*(z, \mathbf{x}, \boldsymbol{\beta})}{\partial \mathbf{x}} \Big|_{\mathbf{x} = \mathbf{x}(t, \boldsymbol{\theta})} + O_p(N^{-\lambda(p \wedge 4)}) \right] \\ \quad \times \left[\frac{\partial \mathbf{x}(t, \boldsymbol{\theta})}{\partial \boldsymbol{\theta}} + O_p(N^{-\lambda(p \wedge 4)/2}) \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} + O_p(N^{-\lambda(p \wedge 4)}) \right] \left[\frac{\partial g^*(z, \mathbf{x}(t, \boldsymbol{\theta}), \boldsymbol{\beta})}{\partial \boldsymbol{\beta}} + O_p(N^{-\lambda(p \wedge 4)}) \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial \phi} + O_p(N^{-\lambda(p \wedge 4)}) \right] \end{pmatrix} \\ &= \begin{pmatrix} E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} \frac{\partial g^*(z, \mathbf{x}(t, \boldsymbol{\theta}), \boldsymbol{\beta})}{\partial \mathbf{x}} \Big|_{\mathbf{x} = \mathbf{x}(t, \boldsymbol{\theta})} \frac{\partial \mathbf{x}(t, \boldsymbol{\theta})}{\partial \boldsymbol{\theta}} \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial g^*} \frac{\partial g^*(z, \mathbf{x}(t, \boldsymbol{\theta}), \boldsymbol{\beta})}{\partial \boldsymbol{\beta}} \right] \\ E_0 \left[\frac{\partial \ell(\boldsymbol{\alpha}; y, \mathbf{x}(t, \boldsymbol{\theta}), \mathbf{z})}{\partial \phi} \right] \end{pmatrix} + O_p(N^{-\lambda(p \wedge 4)/2}) \\ &= G(\boldsymbol{\alpha}) + O_p(N^{-\lambda(p \wedge 4)/2}). \end{aligned}$$

It follows that $\tilde{G}(\tilde{\boldsymbol{\alpha}}) = G(\tilde{\boldsymbol{\alpha}}) + O_p(N^{-\lambda(p \wedge 4)/2})$, then $G(\tilde{\boldsymbol{\alpha}}) = O_p(N^{-\lambda(p \wedge 4)/2})$ from $\tilde{G}(\tilde{\boldsymbol{\alpha}}) = 0$. The Taylor series expansion yields that there exist constants $0 < c_1, c_2 < \infty$ such that

$$c_1 \|\tilde{\boldsymbol{\alpha}} - \boldsymbol{\alpha}_0\| \leq |G(\tilde{\boldsymbol{\alpha}}) - G(\boldsymbol{\alpha}_0)| \leq c_2 \|\tilde{\boldsymbol{\alpha}} - \boldsymbol{\alpha}_0\|.$$

Thus $\|\tilde{\boldsymbol{\alpha}} - \boldsymbol{\alpha}_0\| = O_p(N^{-\lambda(p \wedge 4)/2})$ from $G(\boldsymbol{\alpha}_0) = 0$. It follows that $\tilde{\boldsymbol{\alpha}} \xrightarrow{p} \boldsymbol{\alpha}_0$. By Assumption A9, we have that $\tilde{\boldsymbol{\alpha}}$ is also an interior point of $\Theta \times \mathcal{B} \times \Phi$ for sufficiently large N . Similarly we can show that $\|\tilde{\mathbf{H}} - \mathbf{H}\| = O_p(N^{-\lambda(p \wedge 4)/2})$. By Assumption A10, we have that $\tilde{\mathbf{H}}$ is also nonsingular for sufficiently large N . Then it is straightforward to derive the asymptotic normality of Theorem 2(ii) by verifying the conditions of Theorem 3.3 in [Newey and McFadden \(1994\)](#) on asymptotic normality for MLE. \square

Theorem 3. For the weighted MLE $\hat{\alpha}^*$ in (3.14), under the same assumptions as those in Theorem 2 as well as Assumption A11, we have that,

(i) for $\lambda > 1/(p \wedge 4)$, $\sqrt{N/v_0}(\hat{\alpha}^* - \hat{\alpha})$ has the same conditional limiting distribution as $\sqrt{N}(\hat{\alpha} - \alpha_0)$ has unconditionally, i.e., $\left(\sqrt{N/v_0}(\hat{\alpha}^* - \hat{\alpha}) \mid \{t_i, z_{ik}, y_{iks}\}\right) \xrightarrow{d} \sqrt{N}(\hat{\alpha} - \alpha_0)$. Thus $\sqrt{N}(\hat{\alpha}^* - \alpha_0) \xrightarrow{d} N(0, (1 + v_0)\mathbf{H}^{-1})$.

(ii) for $0 < \lambda \leq 1/(p \wedge 4)$, $\left(\sqrt{N/v_0}(\hat{\alpha}^* - \hat{\alpha}) \mid \{t_i, z_{ik}, y_{iks}\}\right) \xrightarrow{d} \sqrt{N}(\hat{\alpha} - \tilde{\alpha})$. Thus $\sqrt{N}(\hat{\alpha}^* - \tilde{\alpha}) \xrightarrow{d} N(0, (1 + v_0)\tilde{\mathbf{H}}^{-1})$.

Proof: Replace all likelihood functions $\ell(\cdot)$ in the proofs of Theorems 1-2 with the weight likelihood functions $w\ell(\cdot)$ and denote

$$\begin{aligned}\tilde{M}_n^*(\alpha) &= \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} w_{iks} \ell(\alpha; y_{iks}, \tilde{\mathbf{x}}_i, \mathbf{z}_{ik}), \\ M_n^*(\alpha) &= \frac{1}{N} \sum_{i=1}^n \sum_{k=1}^{K_i} \sum_{s=1}^{S_{ik}} w_{iks} \ell(\alpha; y_{iks}, \mathbf{x}_i, \mathbf{z}_{ik}), \\ M^*(\alpha) &= w\ell(\alpha; y, \mathbf{x}, \mathbf{z}),\end{aligned}$$

$$\text{and } \tilde{G}_n^*(\alpha) = \frac{\partial \tilde{M}_n^*(\alpha)}{\partial \alpha}, G_n^*(\alpha) = \frac{\partial M_n^*(\alpha)}{\partial \alpha}, G^*(\alpha) = \frac{\partial E_0[M^*(\alpha)]}{\partial \alpha}.$$

First, we claim that under Assumptions A1-A9 and A11, we have $\hat{\alpha}^* - \alpha_0 \rightarrow 0$, almost surely under P_{α_0} , i.e., Theorem 1 still holds for the weighted MLE $\hat{\alpha}^*$. In fact, by the Jensen's inequality, the convexity of function $-\log x$, $E(w) = 1$ and the independence between w and (t, y, z) , we have

$$\begin{aligned}& E_0[M^*(\alpha)] - E_0[M^*(\alpha_0)] \\ &= E_0[w\ell(\alpha; y, \mathbf{x}, \mathbf{z}) - w\ell(\alpha_0; y, \mathbf{x}, \mathbf{z})] \\ &= E(w)E_0[\ell(\alpha; y, \mathbf{x}, \mathbf{z}) - \ell(\alpha_0; y, \mathbf{x}, \mathbf{z})] \\ &\leq 0,\end{aligned}$$

i.e., $\alpha = \alpha_0$ is still the maximum point of function $E_0[M^*(\alpha)]$. In addition, from Assumption A6, it follows that it is the unique maximum point. Under Assumption A11, similar to (A.1), for any $\alpha_1, \alpha_2 \in \Theta \times \mathcal{B} \times \Phi$, we have

$$|M_n^*(\alpha_1) - M_n^*(\alpha_2)| \leq Q|M_n(\alpha_1) - M_n(\alpha_2)| \leq C(\mathbf{y}, \mathbf{x}, \mathbf{z})Q\|\alpha_1 - \alpha_2\|.$$

The remaining steps to prove this claim are similar to those in the proof of Theorem 1.

Next, we claim that $\sqrt{N}(\hat{\alpha}^* - \alpha_0) \xrightarrow{d} \mathcal{N}(0, (1 + v_0)\mathbf{H}^{-1})$ or $\sqrt{N}(\tilde{\alpha}^* - \tilde{\alpha}) \xrightarrow{d} \mathcal{N}(0, (1 + v_0)\tilde{\mathbf{H}}^{-1})$ for $\lambda > 1/(p \wedge 4)$ and $0 < \lambda \leq 1/(p \wedge 4)$, respectively. We verify it as follows.

Since $\hat{\alpha}^*$ and α_0 are the maximum points of $\tilde{M}_n^*(\alpha)$ and $E_0[M^*(\alpha)]$, respectively, $\tilde{G}_n^*(\hat{\alpha}^*) = 0$ and $G^*(\alpha_0) = 0$. By similar arguments to those in the proof of Theorem 2, we have $\sqrt{N}[\tilde{G}_n^*(\alpha_0) - G^*(\alpha_0)] = \sqrt{N}[G_n^*(\alpha_0) - G^*(\alpha_0)] + o_p(1)$. From the standard central limit theorem and $E(w^2) = 1 + v_0$, we have $\sqrt{N}[G_n^*(\alpha_0) - G^*(\alpha_0)] \xrightarrow{d} \mathcal{N}(0, (1 + v_0)\mathbf{J})$. Thus it follows that $\sqrt{N}[\tilde{G}_n^*(\alpha_0) - G^*(\alpha_0)] \xrightarrow{d} \mathcal{N}(0, (1 + v_0)\mathbf{J})$. In addition, by the weak law of large number and $E(w) = 1$, it follows that

$$\frac{\partial^2 M_n^*(\alpha)}{\partial \alpha \partial \alpha^T} \xrightarrow{p} E_0 \left[\frac{\partial^2 M^*(\alpha)}{\partial \alpha \partial \alpha^T} \right] = E(w)E_0 \left[\frac{\partial^2 M(\alpha)}{\partial \alpha \partial \alpha^T} \right] = -\mathbf{H}.$$

The remaining steps to prove this claim are similar to those in the proof of Theorem 2. Combining the second claim and Theorem 2, we have that

$$\left(\sqrt{N/v_0}(\hat{\alpha}^* - \hat{\alpha}) \mid \{t_i, z_{ik}, y_{iks}\}\right) \xrightarrow{d} \sqrt{N}(\hat{\alpha} - \alpha_0)$$

for $\lambda > 1/(p \wedge 4)$ and

$$\left(\sqrt{N/v_0}(\hat{\alpha}^* - \hat{\alpha}) \mid \{t_i, z_{ik}, y_{iks}\}\right) \xrightarrow{d} \sqrt{N}(\hat{\alpha} - \tilde{\alpha})$$

for $0 < \lambda \leq 1/(p \wedge 4)$, respectively. □

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B Additional Simulation Result

Table B.1: Evaluation of the GODE method by calculating average relative errors (ARE) in parameter estimates based on 500 simulation runs with ρ_E fixed at zero.

distribution	# of obs. per dilu. factor	variance	ARE(%) of GODE				ARE(%) of ESB					
			ρ_E	β_E	δ_{E^*}	c_V	β	ρ_E	β_E	δ_{E^*}	c_V	
binomial	5	-	fixed	8.53%	32.5%	27.7%	3.97%	fixed	91.9%	72.8%	36.5%	3.85%
	10	-	fixed	6.49%	18.9%	18.9%	2.85%	fixed	96.3%	72.1%	30.1%	2.83%
	20	-	fixed	4.27%	12.4%	12.5%	2.02%	fixed	87.5%	38.6%	19.2%	2.33%
Poisson	5	-	fixed	13.0%	45.6%	42.1%	7.05%	fixed	68.1%	85.3%	38.1%	6.95%
	10	-	fixed	8.67%	31.5%	28.8%	4.91%	fixed	72.8%	66.7%	29.4%	4.94%
	20	-	fixed	6.38%	21.2%	19.8%	3.45%	fixed	69.2%	51.2%	19.5%	3.49%
Gamma	5	3.0	fixed	10.3%	48.3%	58.1%	84.3%	fixed	86.0%	86.5%	58.6%	84.6%
	10	1.5	fixed	6.46%	48.8%	44.3%	76.9%	fixed	107%	83.8%	29.8%	77.1%
	20	0.75	fixed	11.8%	35.9%	25.0%	69.5%	fixed	113%	89.4%	21.2%	69.7%
normal	5	3.0	fixed	47.7%	72.4%	116%	54.8%	fixed	66.3%	74.2%	46.1%	71.8%
	10	1.5	fixed	20.1%	53.9%	60.2%	13.9%	fixed	66.7%	61.8%	31.8%	15.4%
	20	0.75	fixed	9.11%	30.5%	27.8%	6.41%	fixed	62.5%	52.0%	22.2%	6.33%