Supplementary Material for Semiparametric Estimation in Secondary Analysis of Case-Control Studies

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S.1 Derivation of the Efficient Score

Having obtained both the score function and the two spaces Λ and Λ^{\perp} , we only need to project the score function onto Λ^{\perp} to obtain the efficient score \mathbf{S}_{eff} . To do this, we write $\mathbf{S}_{\theta} = \mathbf{S} - E(\mathbf{S} \mid D) = \mathbf{g}(\epsilon, \mathbf{X}) - E(\mathbf{g} \mid D) + \mathbf{S}_{\text{eff}}$, where $E_{\text{true}}(\mathbf{g}) = E_{\text{true}}(\epsilon \mathbf{g} \mid \mathbf{X}) = \mathbf{0}$. We alternatively write $\mathbf{S}_{\text{eff}} = \mathbf{S} - \mathbf{g}(\epsilon, \mathbf{X}) - E(\mathbf{S} - \mathbf{g} \mid D)$ and \mathbf{S}_{eff} satisfies $E\{\mathbf{S}_{\text{eff}} - E(\mathbf{S}_{\text{eff}} \mid D) \mid \epsilon, \mathbf{X}\} \sum_{d} (N_d/N) H(d, \mathbf{x}, Y) / p_D^{\text{true}}(d) = \epsilon \mathbf{a}(\mathbf{X})$ and $E(\mathbf{S}_{\text{eff}}) = 0$. However, $E(\mathbf{S}_{\text{eff}} \mid d) = 0$ automatically, hence we can ignore the second requirement and the first requirement simplifies to $E(\mathbf{S}_{\text{eff}} \mid \epsilon, \mathbf{X}) \sum_{d} (N_d/N) \{H(d, \mathbf{X}, Y) / p_D^{\text{true}}(d)\} = \epsilon \mathbf{a}(\mathbf{X})$. This gives

$$\epsilon \mathbf{a}(\mathbf{X}) \left\{ \sum_{d} \frac{N_d}{N} \frac{H(d, \mathbf{X}, Y)}{p_D^{\text{true}}(d)} \right\}^{-1} = E(\mathbf{S} - \mathbf{g} \mid \epsilon, \mathbf{X}) - E\left\{ E(\mathbf{S} - \mathbf{g} \mid D) \mid \epsilon, \mathbf{X} \right\}.$$

It follows that

$$f_{D|X,Y}(d,\mathbf{x},y) = \frac{N_d}{N} \frac{H(d,\mathbf{x},y)}{p_D^{\text{true}}(d)} \left\{ \sum_d \frac{N_d}{N} \frac{H(d,\mathbf{x},y)}{p_D^{\text{true}}(d)} \right\}^{-1}.$$

To simplify notation, in the following calculation we denote

$$\pi_{0} = p_{D}^{\text{true}}(0) = \int \eta_{1}(\mathbf{x})\eta_{2}(\epsilon, \mathbf{x})H(0, \mathbf{x}, y)d\mu(\mathbf{x})d\mu(y);$$

$$\pi_{1} = p_{D}^{\text{true}}(1) = \int \eta_{1}(\mathbf{x})\eta_{2}(\epsilon, \mathbf{x})H(1, \mathbf{x}, y)d\mu(\mathbf{x})d\mu(y);$$

$$b_{0} = E\{f_{D|\mathbf{X},Y}(1, \mathbf{X}, Y) \mid D = 0\};$$

$$b_{1} = E\{f_{D|\mathbf{X},Y}(0, \mathbf{X}, Y) \mid D = 1\};$$

$$\mathbf{c}_{0} = E(\mathbf{S} \mid D = 0) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid D = 0\};$$

$$\mathbf{c}_{1} = E(\mathbf{S} \mid D = 1) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid D = 1\};$$

$$\kappa(\mathbf{x}, y) = \left[\sum_{d=0}^{1} \{N_{d}H(d, \mathbf{x}, y)\}/(N\pi_{d})\right]^{-1};$$

$$\mathbf{u}_{0} = E\{\epsilon\mathbf{a}(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 0\};$$

$$\mathbf{u}_{1} = E\{\epsilon\mathbf{a}(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 1\};$$

$$\mathbf{v}_{0} = E(\mathbf{S} - \mathbf{g} \mid D = 0);$$

$$\mathbf{v}_{1} = E(\mathbf{S} - \mathbf{g} \mid D = 1).$$

Note that $\pi_0 + \pi_1 = 1$, $b_0 N_0 = b_1 N_1$, $\mathbf{c}_0 N_0 + \mathbf{c}_1 N_1 = \mathbf{0}$ and $\mathbf{v}_0 \pi_0 + \mathbf{v}_1 \pi_1 = \mathbf{0}$.

Under a true model, $\pi_0, \pi_1, b_0, b_1, \mathbf{c}_0, \mathbf{c}_1$ are known quantities, while $\mathbf{u}_1, \mathbf{u}_1, \mathbf{v}_0, \mathbf{v}_1$ are not known because $\mathbf{g} = \mathbf{g}(\epsilon, \mathbf{x})$ and $\mathbf{a} = \mathbf{a}(\mathbf{x})$ are not specified. To further obtain $\mathbf{u}_1, \mathbf{u}_1, \mathbf{v}_0, \mathbf{v}_1$,

$$\epsilon \mathbf{a}(\mathbf{x})\kappa(\mathbf{x},y) = E(\mathbf{S} - \mathbf{g} \mid \epsilon, \mathbf{X} = \mathbf{x}) - \mathbf{v}_0 f_{D|X,Y}(0,\mathbf{x},y) - \mathbf{v}_1 f_{D|X,Y}(1,\mathbf{x},y)$$
$$= E(\mathbf{S} \mid \epsilon, \mathbf{X} = \mathbf{x}) - \mathbf{g} - \mathbf{v}_0 f_{D|X,Y}(0,\mathbf{x},y) - \mathbf{v}_1 f_{D|X,Y}(1,\mathbf{x},y).$$

Alternatively, we also have

$$\mathbf{g}(\epsilon, \mathbf{x}) = E(\mathbf{S} \mid \epsilon, \mathbf{X} = \mathbf{x}) - \epsilon \mathbf{a}(\mathbf{x}) \kappa(\mathbf{x}, y) - \mathbf{v}_0 f_{D|X,Y}(0, \mathbf{x}, y) - \mathbf{v}_1 f_{D|X,Y}(1, \mathbf{x}, y).$$
(S.1)

Since $\mathbf{v}_0 = E(\mathbf{S} - \mathbf{g} \mid D = 0)$, we obtain

$$\mathbf{v}_0 = E(\mathbf{S} \mid D = 0) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) - \epsilon \mathbf{a}(\mathbf{X}) \kappa(\mathbf{X}, Y) - \mathbf{v}_0 f_{D\mid X, Y}(0, \mathbf{X}, Y) - \mathbf{v}_1 f_{D\mid X, Y}(1, \mathbf{X}, Y) \mid D = 0\}$$

$$= \mathbf{c}_0 + \mathbf{u}_0 + \mathbf{v}_0 (1 - b_0) + \mathbf{v}_1 b_0.$$

Thus, we have $b_0 \mathbf{v}_0 - b_0 \mathbf{v}_1 - \mathbf{u}_0 = \mathbf{c}_0$. Similarly, from $\mathbf{v}_1 = E(\mathbf{S} - \mathbf{g} \mid D = 1)$, we obtain

$$\mathbf{v}_{1} = E(\mathbf{S} \mid D = 1) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) - \epsilon \mathbf{a}(\mathbf{X}) \kappa(\mathbf{X}, Y) - \mathbf{v}_{0} f_{D\mid X, Y}(0, \mathbf{X}, Y) - \mathbf{v}_{1} f_{D\mid X, Y}(1, \mathbf{X}, Y) \mid D = 1\}$$

$$= \mathbf{c}_{1} + \mathbf{u}_{1} + \mathbf{v}_{0} b_{1} + \mathbf{v}_{1} (1 - b_{1}).$$

Thus, we have $-b_1\mathbf{v}_0 + b_1\mathbf{v}_1 - \mathbf{u}_1 = \mathbf{c}_1$. Since

$$E\left\{\epsilon \mathbf{a}(\mathbf{X})\kappa(\mathbf{X},Y)\right\} = \mathbf{0},$$

we have $\mathbf{u}_0 N_0 + \mathbf{u}_1 N_1 = \mathbf{0}$. Since $E_{\text{true}}(\mathbf{S} - \mathbf{g}) = \mathbf{0}$, we have

$$\mathbf{0} = \sum_{d} \int (\mathbf{S} - \mathbf{g}) f_{\mathbf{X},Y|D}(\mathbf{x}, y, d) \{ f_{X,Y,D}^{\text{true}}(\mathbf{x}, y, d) / f_{\mathbf{X},Y|D}(\mathbf{x}, y, d) \} d\mu(\mathbf{x}) d\mu(y)$$

$$= \sum_{d} \int (\mathbf{S} - \mathbf{g}) f_{\mathbf{X},Y|D}(\mathbf{x}, y, d) \{ f_{X,Y,D}^{\text{true}}(\mathbf{x}, y, d) / f_{\mathbf{X},Y|D}^{\text{true}}(\mathbf{x}, y, d) \} d\mu(\mathbf{x}) d\mu(y)$$

$$= \sum_{d} \int (\mathbf{S} - \mathbf{g}) f_{\mathbf{X},Y|D}(\mathbf{x}, y, d) p_{D}^{\text{true}}(d) d\mu(\mathbf{x}) d\mu(y)$$

$$= \pi_{0} \mathbf{v}_{0} + \pi_{1} \mathbf{v}_{1}.$$

Combining the above relations, we have obtained $N_0\mathbf{u}_0 + N_1\mathbf{u}_1 = \mathbf{0}$, $\pi_0\mathbf{v}_0 + \pi_1\mathbf{v}_1 = \mathbf{0}$, $b_0\mathbf{v}_0 - b_0\mathbf{v}_1 - \mathbf{u}_0 = \mathbf{c}_0$ and $-b_1\mathbf{v}_0 + b_1\mathbf{v}_1 - \mathbf{u}_1 = \mathbf{c}_1$. The last two equations are equivalent so one is redundant. Using these relations, we can rewrite $\mathbf{u}_1, \mathbf{v}_0, \mathbf{v}_1$ as a function of \mathbf{u}_0 :

$$\mathbf{u}_1 = -(N_0/N_1)\mathbf{u}_0, \quad \mathbf{v}_0 = (\pi_1/b_0)(\mathbf{u}_0 + \mathbf{c}_0), \quad \mathbf{v}_1 = -(\pi_0/b_0)(\mathbf{u}_0 + \mathbf{c}_0).$$
 (S.2)

We cannot obtain a more explicit expression for \mathbf{u}_0 at this stage, but we can further obtain $\mathbf{a}(\mathbf{x})$ as a function of \mathbf{u}_0 . Using (S.1) and since $E_{\text{true}}(\epsilon \mathbf{g} \mid \mathbf{X}) = \mathbf{0}$, we have

$$E_{\text{true}} \left\{ \epsilon E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid \mathbf{X} \right\} - E_{\text{true}} \left\{ \epsilon^2 \kappa(\mathbf{X}, Y) \mid \mathbf{X} \right\} \mathbf{a}(\mathbf{X}) \\ - \mathbf{v}_0 E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X} \right\} - \mathbf{v}_1 E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(1, \mathbf{X}, Y) \mid \mathbf{X} \right\} = \mathbf{0}.$$

Hence

$$\mathbf{a}(\mathbf{X}) = \begin{bmatrix} E_{\text{true}} \left\{ \epsilon^{2} \kappa(\mathbf{X}, Y) \mid \mathbf{X} \right\} \end{bmatrix}^{-1} \\ \begin{bmatrix} E_{\text{true}} \left\{ \epsilon E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid \mathbf{X} \right\} - \mathbf{v}_{0} E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X} \right\} \\ - \mathbf{v}_{1} E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(1, \mathbf{X}, Y) \mid \mathbf{X} \right\} \end{bmatrix} \\ = \begin{bmatrix} E_{\text{true}} \left\{ \epsilon^{2} \kappa(\mathbf{X}, Y) \mid \mathbf{X} \right\} \end{bmatrix}^{-1} \begin{bmatrix} E_{\text{true}} \left\{ \epsilon E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid \mathbf{X} \right\} \\ - (\pi_{1}/b_{0})(\mathbf{u}_{0} + \mathbf{c}_{0}) E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X} \right\} \\ + (\pi_{0}/b_{0})(\mathbf{u}_{0} + \mathbf{c}_{0}) E_{\text{true}} \left\{ \epsilon f_{D|X,Y}(1, \mathbf{X}, Y) \mid \mathbf{X} \right\} \end{bmatrix}.$$

To further simplify notation, denote

$$t_{1}(\mathbf{X}) = \left[E_{\text{true}}\left\{\epsilon^{2}\kappa(\mathbf{X},Y) \mid \mathbf{X}\right\}\right]^{-1};$$

$$\mathbf{t}_{2}(\mathbf{X}) = E_{\text{true}}\left\{\epsilon E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid \mathbf{X}\right\} - (\pi_{1}/b_{0})\mathbf{c}_{0}E_{\text{true}}\left\{\epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X}\right\}$$

$$+(\pi_{0}/b_{0})\mathbf{c}_{0}E_{\text{true}}\left\{\epsilon f_{D|X,Y}(1, \mathbf{X}, Y) \mid \mathbf{X}\right\}$$

$$= E_{\text{true}}\left\{\epsilon E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid \mathbf{X}\right\} - (\mathbf{c}_{0}/b_{0})E_{\text{true}}\left\{\epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X}\right\};$$

$$t_{3}(\mathbf{X}) = -(\pi_{1}/b_{0})E_{\text{true}}\left\{\epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X}\right\} + (\pi_{0}/b_{0})E_{\text{true}}\left\{\epsilon f_{D|X,Y}(1, \mathbf{X}, Y) \mid \mathbf{X}\right\}$$

$$= -b_{0}^{-1}E_{\text{true}}\left\{\epsilon f_{D|X,Y}(0, \mathbf{X}, Y) \mid \mathbf{X}\right\}.$$
(S.3)

Then

$$\mathbf{a}(\mathbf{x}) = t_1(\mathbf{x})\{\mathbf{t}_2(\mathbf{x}) + t_3(\mathbf{x})\mathbf{u}_0\},\tag{S.4}$$

hence the definition of \mathbf{u}_0 yields

$$\mathbf{u}_0 = E\left[\epsilon \left[t_1(\mathbf{X}) \{\mathbf{t}_2(\mathbf{X}) + t_3(\mathbf{X})\mathbf{u}_0\}\right] \kappa(\mathbf{X}, Y) \mid D = 0\right]$$

$$= E\left[\epsilon t_1(\mathbf{X})\mathbf{t}_2(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 0\right] + E\left[\epsilon t_1(\mathbf{X})t_3(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 0\right] \mathbf{u}_0.$$

This yields

$$\mathbf{u}_0 = (1 - E\left[\epsilon t_1(\mathbf{X})t_3(\mathbf{X})\kappa(\mathbf{X},Y) \mid D = 0\right])^{-1} E\left[\epsilon t_1(\mathbf{X})\mathbf{t}_2(\mathbf{X})\kappa(\mathbf{X},Y) \mid D = 0\right].$$
 (S.5)

Combining the above results, we have obtained the analytic form of $\mathbf{S}_{\text{eff}} = \mathbf{S} - \mathbf{g} - E(\mathbf{S} - \mathbf{g} \mid D = d)$, where \mathbf{g} is given in (S.1), $\mathbf{a}(\mathbf{x})$ is given in (S.4), \mathbf{v}_0 , \mathbf{v}_1 are given in (S.2) \mathbf{u}_0 is given in (S.5) and the functions t_1 , t_2 , t_3 are given in (S.3).

In forming the estimating equation $\sum_{i=1}^{N} \mathbf{S}_{\text{eff}} = 0$, we will have $\sum_{i=1}^{N} [\mathbf{S}(\mathbf{X}_i, Y_i, D_i) - \mathbf{g}\{Y_i - m(\mathbf{X}_i, \boldsymbol{\beta}), \mathbf{X}_i\}] - N_0 E(\mathbf{S} - \mathbf{g} \mid D = 0) - N_1 E(\mathbf{S} - \mathbf{g} \mid D = 1) = \mathbf{0}$. Using (S.1), we obtain

$$E(\mathbf{S} - \mathbf{g} \mid D = 0) = E(\mathbf{S} \mid D = 0) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid D = 0\} + E\{\epsilon \mathbf{a}(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 0\} + \mathbf{v}_0 E\{f_{D\mid X, Y}(0, \mathbf{X}, Y) \mid D = 0\} + \mathbf{v}_1 E\{f_{D\mid X, Y}(1, \mathbf{X}, Y) \mid D = 0\}$$

$$= \mathbf{c}_0 + \mathbf{u}_0 + \mathbf{v}_0(1 - b_0) + \mathbf{v}_1 b_0$$

and

$$E(\mathbf{S} - \mathbf{g} \mid D = 1) = E(\mathbf{S} \mid D = 1) - E\{E(\mathbf{S} \mid \epsilon, \mathbf{X}) \mid D = 1\} + E\{\epsilon \mathbf{a}(\mathbf{X})\kappa(\mathbf{X}, Y) \mid D = 1\} + \mathbf{v}_0 E\{f_{D\mid X, Y}(0, \mathbf{X}, Y) \mid D = 1\} + \mathbf{v}_1 E\{f_{D\mid X, Y}(1, \mathbf{X}, Y) \mid D = 1\}$$

$$= \mathbf{c}_1 + \mathbf{u}_1 + \mathbf{v}_0 b_1 + \mathbf{v}_1 (1 - b_1),$$

hence

$$N_0 E(\mathbf{S} - \mathbf{g} \mid D = 0) + N_1 E(\mathbf{S} - \mathbf{g} \mid D = 1)$$

$$N_0 \{ \mathbf{c}_0 + \mathbf{u}_0 + \mathbf{v}_0 (1 - b_0) + \mathbf{v}_1 b_0 \} + N_1 \{ \mathbf{c}_1 + \mathbf{u}_1 + \mathbf{v}_0 b_1 + \mathbf{v}_1 (1 - b_1) \}$$

$$(N_0 \mathbf{c}_0 + N_1 \mathbf{c}_1) + (N_0 \mathbf{u}_0 + N_1 \mathbf{u}_1) + (N_0 \mathbf{v}_0 + N_1 \mathbf{v}_1) + (\mathbf{v}_1 - \mathbf{v}_0) (N_0 \mathbf{b}_0 - N_1 \mathbf{b}_1).$$

Thus, the estimating equation simplifies to $\sum_{i=1}^{N} [\mathbf{S}(\mathbf{X}_i, Y_i, D_i) - \mathbf{g}\{Y_i - m(\mathbf{X}_i, \boldsymbol{\beta}), \mathbf{X}_i\}] - N_0 \mathbf{v}_0 - N_1 \mathbf{v}_1 = \mathbf{0}.$

S.2 Verification of Assumptions 1-2 in the Linear Model as in Remark 2

Let $\mathcal{D}^c = [-K, K]$ for a sufficiently large K. We have assumed that the conditional distribution of ϵ satisfies the property that $\operatorname{pr}(|\epsilon| > K | \mathbf{X} = \mathbf{x}) \to 0$ as $K \to \infty$ uniformly in \mathbf{x} , and hence $\operatorname{pr}(\epsilon \in \mathcal{D})$ can be made arbitrarily small uniformly in \mathbf{x} .

Without loss of generality, assume the first component of $\alpha_1 + \beta_1 \alpha_2$ is not zero. We consider three situations, 1. $\widetilde{\beta}_c - \beta_c \neq 0$, 2. $\widetilde{\beta}_c - \beta_c = 0$ and the first component of $\widetilde{\beta}_1 - \beta_1$ is not zero and 3) $\widetilde{\beta}_c - \beta_c = 0$ and the first component of $\widetilde{\beta}_1 - \beta_1$ is zero. Note that ϵ has conditional mean zero. Thus for most common density functions with bounded variance function, this requirement is satisfied.

Case 1 and Case 2: If the first component of $\alpha_1 + \beta_1 \alpha_2$ is positive, let $\mathbf{c}_1 = (\infty, \mathbf{0}^{\mathrm{T}})^{\mathrm{T}}$, otherwise, let $\mathbf{c}_1 = (-\infty, \mathbf{0}^{\mathrm{T}})^{\mathrm{T}}$. Let $c_2 = 0$, $\mathcal{D}^c = [-K, K]$ for a sufficiently large K. Then

$$\sup_{\epsilon \in \mathcal{D}^c} \lim_{\mathbf{x} \to \mathbf{c}_1} |(1 + \exp[\alpha_c + u\{\mathbf{x}, m(\mathbf{x}, \boldsymbol{\beta}) + \epsilon, \boldsymbol{\alpha}_1, \alpha_2\}])^{-1} - c_2|$$

$$= \sup_{-K \le \epsilon \le K} \lim_{\mathbf{x} \to \mathbf{c}_1} |[1 + \exp\{\alpha_c + \mathbf{x}^{\mathrm{T}}(\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2) + \beta_c \alpha_2 + \epsilon \alpha_2\}]^{-1}|$$

$$= 0.$$

Finally, since $\mathcal{D} = (-\infty, -K) \cup (K, \infty)$, we have that every element $u \in \mathcal{D}$ satisfies u > 1 as long as K > 1. We have thus verified Assumption 1.

We have $c(\boldsymbol{\beta}, \widetilde{\boldsymbol{\beta}}) = \mathbf{c}_1^{\mathrm{T}}(\widetilde{\boldsymbol{\beta}}_1 - \boldsymbol{\beta}_1) + (\widetilde{\boldsymbol{\beta}}_c - \boldsymbol{\beta}_c)$, which is not zero if $\widetilde{\boldsymbol{\beta}}_c - \boldsymbol{\beta}_c \neq 0$ (case 1), and is also not zero if the first component of $\widetilde{\boldsymbol{\beta}}_1 - \boldsymbol{\beta}_1$ is not zero (case 2). Thus, Assumption 2 also holds.

Case 3: Since $\tilde{\boldsymbol{\beta}} \neq \boldsymbol{\beta}$, without loss of generality, we assume the second component of $\tilde{\boldsymbol{\beta}}_1 - \boldsymbol{\beta}_1$ is not zero in this case. We select \mathbf{c}_1 as follows. If the first component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is positive and the second component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is non-negative, let $\mathbf{c}_1 = (\infty, \infty, \mathbf{0}^T)^T$. If the first component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is negative, and the second component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is negative and the second component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is negative and the second component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is non-negative, let $\mathbf{c}_1 = (-\infty, \infty, \mathbf{0}^T)^T$. If the first component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is negative and the second component of $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1 \alpha_2$ is negative, let $\mathbf{c}_1 = (-\infty, -\infty, \mathbf{0}^T)^T$. The selection of c_2, \mathcal{D}, K remains the same as in Cases 1 and 2. We an see the same arguments lead to the verification of Assumption 1. In addition, $c(\boldsymbol{\beta}, \boldsymbol{\tilde{\beta}}) = \mathbf{c}_1^T (\boldsymbol{\tilde{\beta}}_1 - \boldsymbol{\beta}_1)$, which is either ∞ or $-\infty$, and is thus not zero. Thus, Assumption 2 also holds.

S.3 Verification of Nonidentifiability in the Special Case in Remark 2

Here, we provide the details of the proof of the nonidentifiability result in Remark 2, which happens when when $m(\mathbf{x}, \boldsymbol{\beta}) = \mathbf{x}^{\mathrm{T}}\boldsymbol{\beta}_1 + \beta_c$ and $u(\mathbf{x}, y, \boldsymbol{\alpha}_1, \boldsymbol{\alpha}_c) = \mathbf{x}^{\mathrm{T}}\boldsymbol{\alpha}_1 + y\alpha_2 + \alpha_c$, and $\boldsymbol{\alpha}_1 + \boldsymbol{\beta}_1\alpha_2 = \mathbf{0}$. We first exclude a special case when $\alpha_2 = 0$. This special case implies $\boldsymbol{\alpha}_1 = \mathbf{0}$ and $\alpha_2 = 0$, hence the case-control sampling is in fact random sampling. Thus, in the following, we assume $\alpha_2 \neq 0$, We point out that

$$m_1 \equiv \int \epsilon \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + \epsilon \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + \epsilon \alpha_2)} \eta_2(\epsilon) d\mu(\epsilon) \neq 0.$$

This is because we can use the mean value theorem for integration to obtain

$$\int \epsilon \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + \epsilon \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + \epsilon \alpha_2)} \eta_2(\epsilon) d\mu(\epsilon)$$

$$= \int_{-\infty}^{0} \epsilon \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + \epsilon \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + \epsilon \alpha_2)} \eta_2(\epsilon) d\mu(\epsilon) + \int_{0}^{\infty} \epsilon \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + \epsilon \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + \epsilon \alpha_2)} \eta_2(\epsilon) d\mu(\epsilon)$$

$$= \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 - k_1 \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 - k_1 \alpha_2)} \int_{-\infty}^{0} \epsilon \eta_2(\epsilon) d\mu(\epsilon) + \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + k_2 \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + k_2 \alpha_2)} \int_{0}^{\infty} \epsilon \eta_2(\epsilon) d\mu(\epsilon)$$

$$= \left\{ \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 + k_2 \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 + k_2 \alpha_2)} - \frac{1 + \exp(\widetilde{\alpha}_c + \beta_c \alpha_2 - k_1 \alpha_2)}{1 + \exp(\alpha_c + \beta_c \alpha_2 - k_1 \alpha_2)} \right\} \int_{0}^{\infty} \epsilon \eta_2(\epsilon) d\mu(\epsilon)$$

$$= \frac{\{\exp(\widetilde{\alpha}_c) - \exp(\alpha_c)\} \exp(\beta_c \alpha_2) \{\exp(k_2 \alpha_2) - \exp(-k_1 \alpha_2)\}}{\{1 + \exp(\alpha_c + \beta_c \alpha_2 + k_2 \alpha_2)\} \{1 + \exp(\alpha_c + \beta_c \alpha_2 - k_1 \alpha_2)\}} \int_{0}^{\infty} \epsilon \eta_2(\epsilon) d\mu(\epsilon) \neq 0,$$

where k_1, k_2 are positive constants.

Following the notation in the proof of Proposition 1, we define $\eta_2(\epsilon, \mathbf{x}) = \eta_2(\epsilon)$,

$$\widetilde{\eta}_{1}(\mathbf{x}) = \frac{\widetilde{\pi}_{0}}{\pi_{0}} \eta_{1}(x) \int \frac{1 + \exp(\widetilde{\alpha}_{c} + \beta_{c}\alpha_{2} + \epsilon\alpha_{2})}{1 + \exp(\alpha_{c} + \beta_{c}\alpha_{2} + \epsilon\alpha_{2})} \eta_{2}(\epsilon) d\mu(\epsilon);$$

$$\widetilde{\beta}_{c} = \beta_{c} + \left\{ \int \frac{1 + \exp(\widetilde{\alpha}_{c} + \beta_{c}\alpha_{2} + \epsilon\alpha_{2})}{1 + \exp(\alpha_{c} + \beta_{c}\alpha_{2} + \epsilon\alpha_{2})} \eta_{2}(\epsilon) d\mu(\epsilon) \right\}^{-1} m_{1};$$

$$\widetilde{\eta}_{2}(\epsilon) = c_{0} \frac{1 + \exp(\widetilde{\alpha}_{c} + \widetilde{\beta}_{c}\alpha_{2} + \epsilon\alpha_{2})}{1 + \exp(\alpha_{c} + \widetilde{\beta}_{c}\alpha_{2} + \epsilon\alpha_{2})} \eta_{2}(\epsilon + \widetilde{\beta}_{c} - \beta_{c}),$$

where

$$c_0^{-1} = \int \frac{1 + \exp(\widetilde{\alpha}_c + \widetilde{\beta}_c \alpha_2 + \epsilon \alpha_2)}{1 + \exp(\alpha_c + \widetilde{\beta}_c \alpha_2 + \epsilon \alpha_2)} \eta_2(\epsilon + \widetilde{\beta}_c - \beta_c) d\mu(\epsilon).$$

Obviously $\int \widetilde{\eta}_2(\epsilon) d\mu(\epsilon) = 1$. We can easily verify that

$$\begin{split} \int \widetilde{\eta}_{1}(\mathbf{x}) d\mu(\mathbf{x}) &= \frac{\widetilde{\pi}_{0}}{\pi_{0}} \int \eta_{1}(x) \frac{1 + \exp(\widetilde{\alpha}_{c} + \beta_{c}\alpha_{2} + \epsilon \alpha_{2})}{1 + \exp(\alpha_{c} + \beta_{c}\alpha_{2} + \epsilon \alpha_{2})} \eta_{2}(\epsilon) d\mu(\epsilon) d\mu(\mathbf{x}) \\ &= \frac{\widetilde{\pi}_{0}}{\pi_{0}} \{ \pi_{0} + \exp(\widetilde{\alpha}_{c} - \alpha_{c}) \pi_{1} \} \\ &= \widetilde{\pi}_{0} + \frac{\widetilde{\pi}_{0} \exp(\widetilde{\alpha}_{c}) \pi_{1}}{\pi_{0} \exp(\alpha_{c})} = \widetilde{\pi}_{0} + \widetilde{\pi}_{1} = 1, \end{split}$$

using the intermediate results in the proof of Proposition 1. We can also obtain

$$\int \epsilon \widetilde{\eta}_{2}(\epsilon) d\mu(\epsilon) = c_{0} \int \epsilon \frac{1 + \exp(\widetilde{\alpha}_{c} + \widetilde{\beta}_{c}\alpha_{2} + \epsilon\alpha_{2})}{1 + \exp(\alpha_{c} + \widetilde{\beta}_{c}\alpha_{2} + \epsilon\alpha_{2})} \eta_{2}(\epsilon + \widetilde{\beta}_{c} - \beta_{c}) d\mu(\epsilon)
= c_{0} \int (t + \beta_{c} - \widetilde{\beta}_{c}) \frac{1 + \exp(\widetilde{\alpha}_{c} + \beta_{c}\alpha_{2} + t\alpha_{2})}{1 + \exp(\alpha_{c} + \beta_{c}\alpha_{2} + t\alpha_{2})} \eta_{2}(t) d\mu(\epsilon) = 0.$$

Now we can easily verify that the two parameter sets $\{\alpha_c, \boldsymbol{\alpha}_1, \alpha_2, \boldsymbol{\beta}_1, \beta_c, \eta_1(\mathbf{x}), \eta_2(\epsilon)\}$ and $\{\widetilde{\alpha}_c, \boldsymbol{\alpha}_1, \alpha_2, \boldsymbol{\beta}_1, \widetilde{\beta}_c, \widetilde{\eta}_1(\mathbf{x}), \widetilde{\eta}_2(\epsilon)\}$ satisfy (A.1), hence the problem is not identifiable.

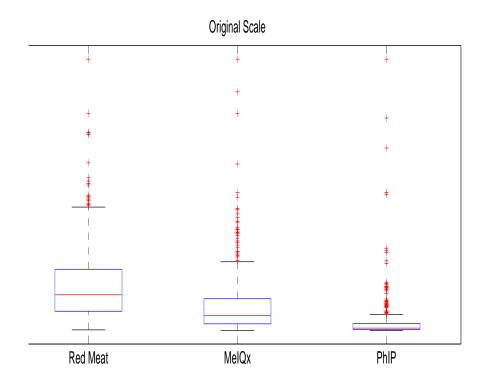


Figure S.1: Boxplots of the variables in the <u>original</u> data scale among the controls. Each variable has been normalized to have maximum value 1.0. This and Figure S.2 indicate a need for data transformation.

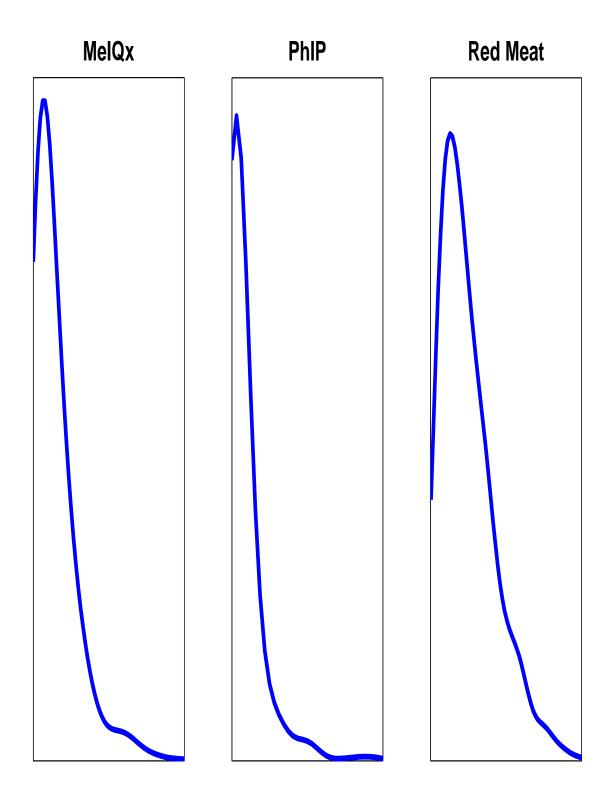


Figure S.2: Kernel density estimates in the <u>original</u> data scale, among the controls. This and Figure S.1 indicate a need for data transformation.

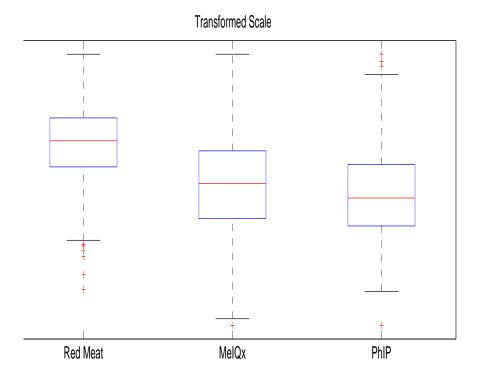


Figure S.3: Boxplots of the variables in the <u>transformed</u> data scale, among the controls. Each variable has been normalized to have maximum value 1.0. Contrast with the boxplots in the original data scale in Figure S.1.

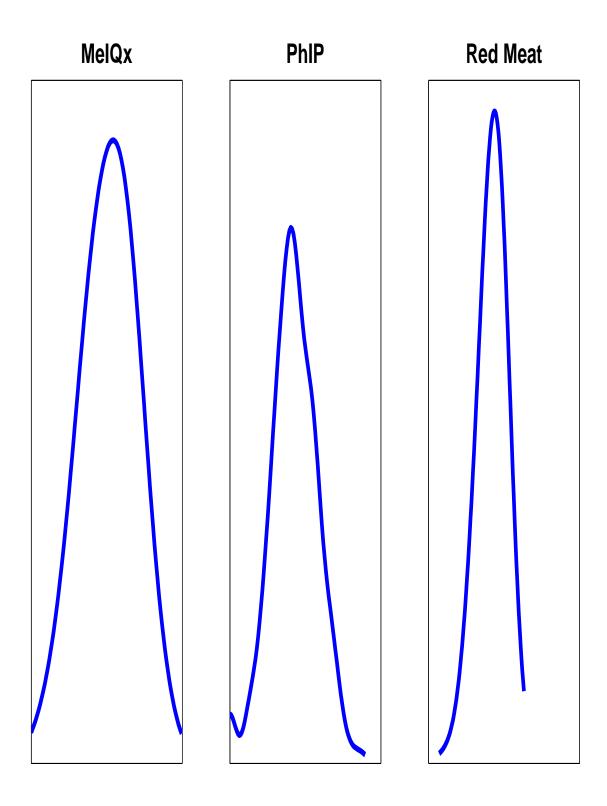


Figure S.4: Kernel density estimates in the $\underline{\text{transformed}}$ data scale, among the controls. Contrast with the boxplots in the original data scale in Figure S.2

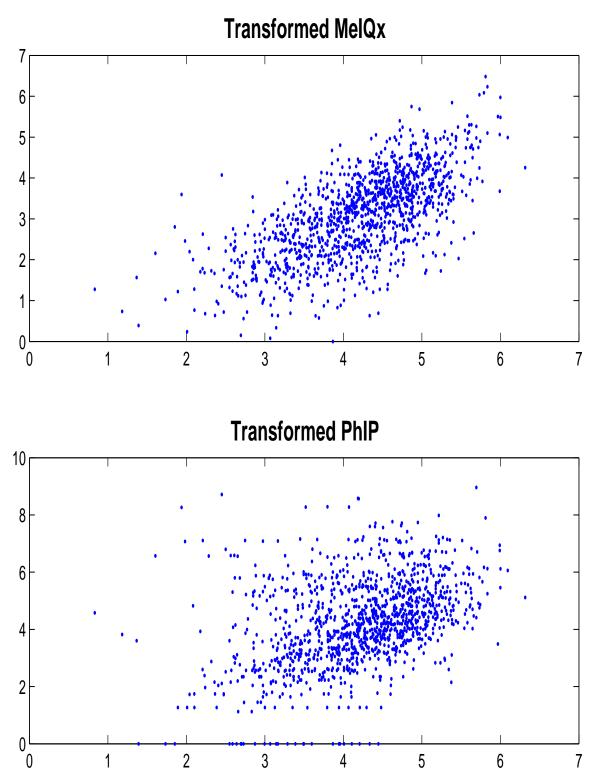


Figure S.5: Scatterplots of $\underline{\text{transformed}}$ MeIQX and PhIP against transformed Red Meat, among the controls.