

Appendix: Proof of Proposition 1

Without loss of generality, we consider $k = 1$. Let us call T^* to the random variable $T(e, M_1(e^*), W_1(e^{**}))$, being $(e, e^*, e^{**}) \in \{0, 1\}^3$. Then, the rate can be expressed as:

$$\gamma(t; e, M_1(e^*), W_1(e^{**})) = \lim_{dt \rightarrow 0} \frac{1}{dt} P(T^* \in [t, t + dt] \mid T^* \geq t).$$

It holds that:

$$P(T^* \in [t, t + dt] \mid T^* \geq t) = \mathbb{E}_{\{X|T^* > t\}}[P(T^* \in [t, t + dt] \mid X = x, T^* \geq t)]$$

and, similarly, being $F(m_1, w_1)$ the distribution function of $M_1(e^*), W_1(e^{**})$ given that $X = x$ and $T^* > t$:

$$\begin{aligned} P(T^* \in [t, t + dt] | X = x, T^* \geq t) &= \int_{\mathbb{R}^K} P(T^* \in [t, t + dt] \mid X = x, M_1 = m_1, W_1 = w_1, T^* \geq t) dF(m_1, w_1) \\ &= \int_{\mathbb{R}^K} P(T(e, m_1, w_1) \in [t, t + dt] \mid X = x, M_1 = m_1, W_1 = w_1, T(e, m_1, w_1) > t) dF(m_1, w_1) \\ &= \int_{\mathbb{R}^K} P(T(e, m_1, w_1) \in [t, t + dt] | X = x, E = e, M_1 = m_1, W_1 = w_1, T(e, m_1, w_1) > t) dF(m_1, w_1). \end{aligned}$$

Using the bounded convergency theorem and the additive risk hypothesis:

$$\begin{aligned} \gamma(t; e, M_1(e^*), W_1(e^{**})) &= \mathbb{E}_{\{X|T^* > t\}} \left[\int_{\mathbb{R}^K} (\lambda_0(t) + \lambda_1 e + \lambda_2^T x + \lambda_3^T(m_1, w_1^T)^T) dF(m_1, w_1) \right] \\ &= \lambda_0(t) + \lambda_1 e + \lambda_2^T \mathbb{E}(X | T^* > t) + \mathbb{E}_{\{X|T^* > t\}} \left[\int_{\mathbb{R}^K} \lambda_3^T(m_1, w_1^T)^T dF(m_1, w_1) \right]. \end{aligned}$$

In addition,

$$\begin{aligned} \int_{\mathbb{R}^K} \lambda_3^T(m_1, w_1^T)^T dF(m_1, w_1) &= \int_{\mathbb{R}^K} \lambda_3^T(m_1, w_1^T)^T f(M_1(e^*) = m_1, W_1(e^{**}) = w_1 \mid X = x, T^* > t) dm_1 dw_1 \\ &= \int_{\mathbb{R}^K} \lambda_3^T(m_1, w_1^T)^T \frac{P(T^* > t \mid M_1 = m_1, W_1 = w_1, X = x) f(M_1(e^*) = m_1, W_1(e^{**}) = w_1 \mid X = x)}{P(T^* > t | X = x)} dm_1 dw_1. \end{aligned}$$

Following the same arguments and taking into account that Aalen additive hazards models have been used for a time-to-event setting:

$$\begin{aligned} P(T^* > t \mid M_1 = m_1, W_1 = w_1, X = x) &= P(T(e, m_1, w_1) > t \mid M_1 = m_1, W_1 = w_1, X = x, E = e) \\ &= \exp\left\{-\int_0^t \lambda_0(u)du - \lambda_1 et - \lambda_2^T xt - \lambda_3^T (m_1, w_1^T)^T t\right\}. \end{aligned}$$

Also,

$$\begin{aligned} P(T^* > t \mid X = x) &= \int_{\mathbb{R}^K} P(T^* > t \mid M_1 = m_1, W_1 = w_1, X = x) dF(m_1, w_1) \\ &= \exp\left\{-\int_0^t \lambda_0(u)du - \lambda_1 et - \lambda_2^T xt\right\} \mathbb{E}(\exp\{-\lambda_3^T (m_1, w_1^T)^T t\}) \end{aligned}$$

Hence, defining $V = \lambda_3^T (m_1, w_1^T)^T$ and putting together the above results, it holds that:

$$\int_{\mathbb{R}^K} \lambda_3^T (m_1, w_1^T)^T dF(m_1, w_1) = \int_{\mathbb{R}^K} V dF(m_1, w_1) = \frac{\mathbb{E}(V \exp\{-tV\} \mid X = x)}{\mathbb{E}(\exp\{-tV\} \mid X = x)}.$$

The distribution of $(M_1(e^*), W_1(e^{**}) \mid X = x)$ is multivariate normal [?] with covariance matrix Σ , which does not depend on the exposures, and expected values:

$$\mathbb{E}(M_1(e^*) \mid X = x) = \alpha_{01} + \alpha_{11}e^* + \alpha_{21}x$$

$$\mathbb{E}(M_j(e^{**}) \mid X = x) = \alpha_{0j} + \alpha_{1j}e^{**} + \alpha_{2j}x, \quad j = 2, \dots, K.$$

Thus, the distribution of V is normal with:

$$\mathbb{E}(V \mid X = x) = \lambda_3^T \alpha_0 + \lambda_{31} \alpha_{11} e^* + \sum_{j=2}^K \lambda_{3j} \alpha_{1j} e^{**} + \lambda_3^T \alpha_2 x$$

$$\text{Var}(V \mid X = x) = \lambda_3^T \Sigma \lambda_3.$$

Thus, following Lange and Hansen 2011 [?],

$$\begin{aligned} \frac{\mathbb{E}(V \exp\{-tV\} | X = x)}{\mathbb{E}(\exp\{-tV\} | X = x)} &= \mathbb{E}(V | X = x) - t \text{Var}(V | X = x) \\ &= \lambda_3^T \alpha_0 + \lambda_{31} \alpha_{11} e^* + \sum_{j=2}^K \lambda_{3j} \alpha_{1j} e^{**} + \lambda_3^T \alpha_2 x - t \lambda_3^T \Sigma \lambda_3, \end{aligned}$$

and it holds that the counterfactual rate can be expressed as:

$$\gamma(t; e, M(e^*), W(e^{**})) = C(t) + \lambda_1 e + \lambda_{31} \alpha_{11} e^* + \sum_{j=2}^K \lambda_{3j} \alpha_{1j} e^{**},$$

being $C(t) = \lambda_0(t) + \lambda_2^T \mathbb{E}(X | T^* > t) + \lambda_3^T \alpha_0 + \lambda_3^T \alpha_2 \mathbb{E}(X | T^* > t) - t \lambda_3^T \Sigma \lambda_3$ a function of t that does not depend on the exposures.