

## HW#6 Survival analysis II

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### Summary of section 2.1 and section 2.2 in

Emura T & Chen YH (2014), Gene selection for survival data under dependent censoring, a copula-based approach, *Statistical Methods in Medical Research*, DOI: 10.1177/0962280214533378

Section 2.1 is talk about the univariate selection for censored survival data. In univariate selection, a Cox regression based on univariate models

$$h(t | x_{ij}) = h_{0j}(t)e^{\beta_j x_{ij}}, \quad j = 1, \dots, p$$

and there is an assumption

**Assumption I:** The survival time  $T$  and censoring time  $U$  are conditionally independent given a gene  $x_{ij}$  for all  $j = 1, \dots, p$ .

Even when the previous model is incorrect, the univariate estimate  $\hat{\beta}_j$  still possesses a valid meaning under Assumption I. This result proved that Assumption I is more important than the previous model in applying univariate selection.

In section 2.2, there is a more reasonable assumption is the conditional independent in which  $T$  and  $U$  are conditionally independent given all components of  $\mathbf{x}$ . And Figure 1 shows an example of how Assumption I fails to hold. Then the result of analysis shows that the conditional independence yields dependency between  $T$  and  $U$  given only  $x_j$ , and thus Assumption I does not hold.

So in general,  $T$  and  $U$  may be dependent for any given  $x_j$  with an unknown dependence structure.

## Detail derivation of Eq. (2)

The Laplace transform of a random vector  $\mathbf{x}$  is defined as  $\varphi(\mathbf{u}) = E\{\exp(-\mathbf{u}'\mathbf{x})\}$ .

Hence,

$$\varphi_{\beta_{(-j)}, \gamma_{(-j)}}(u, v) = E\{\exp(-ue^{\beta_{(-j)}\mathbf{x}_{(-j)}})\exp(-ve^{\gamma_{(-j)}\mathbf{x}_{(-j)}}) | x_j\},$$

$$\varphi_{\beta_{(-j)}}(u) = E\{\exp(-ue^{\beta_{(-j)}\mathbf{x}_{(-j)}}) | x_j\} = \varphi_{\beta_{(-j)}, \gamma_{(-j)}}(u, 0),$$

$$\varphi_{\gamma_{(-j)}}(u) = E\{\exp(-ue^{\gamma_{(-j)}\mathbf{x}_{(-j)}}) | x_j\} = \varphi_{\beta_{(-j)}, \gamma_{(-j)}}(0, u)$$

are the Laplace transform for some random vectors.

Then we have

$$\begin{aligned} & P(T > t, U > u | x_j) \\ &= E\{P(T > t, U > u | \mathbf{x}) | x_j\} \\ &= E\{P(T > t | \mathbf{x})P(U > u | \mathbf{x}) | x_j\} \\ &= E[\exp\{-\Lambda_T(t)e^{\beta\mathbf{x}}\}\exp\{-\Lambda_U(u)e^{\gamma\mathbf{x}}\} | x_j] \\ &= E[\exp\{-\Lambda_T(t)e^{\beta_j x_j} e^{\beta_{(-j)}\mathbf{x}_{(-j)}}\}\exp\{-\Lambda_U(u)e^{\gamma_j x_j} e^{\gamma_{(-j)}\mathbf{x}_{(-j)}}\} | x_j] \\ &= \varphi_{\beta_{(-j)}, \gamma_{(-j)}}\{\Lambda_T(t)e^{\beta_j x_j}, \Lambda_U(u)e^{\gamma_j x_j}\}, \end{aligned}$$

and then

$$\begin{aligned} P(T > t | x_j) &= E\{P(T > t | \mathbf{x}) | x_j\} \\ &= E[\exp\{-\Lambda_T(t)e^{\beta\mathbf{x}}\} | x_j] \\ &= E[\exp\{-\Lambda_T(t)e^{\beta_j x_j} e^{\beta_{(-j)}\mathbf{x}_{(-j)}}\} | x_j] \\ &= \varphi_{\beta_{(-j)}}\{\Lambda_T(t)e^{\beta_j x_j}\}. \end{aligned}$$

Therefore,

$$\Lambda_T(t)e^{\beta_j x_j} = \varphi_{\beta_{(-j)}}^{-1}\{P(T > t | x_j)\}$$

and similarly,

$$\Lambda_U(u)e^{\gamma_j x_j} = \varphi_{\gamma_{(-j)}}^{-1}\{P(U > u | x_j)\}.$$

Hence we can obtain

$$P(T > t, U > u | x_j) = \varphi_{\beta_{(-j)}, \gamma_{(-j)}}[\varphi_{\beta_{(-j)}}^{-1}\{P(T > t | x_j)\}, \varphi_{\gamma_{(-j)}}^{-1}\{P(U > u | x_j)\}].$$

### Detail derivation of Eq. (3)

For the special case of  $\boldsymbol{\beta} = \boldsymbol{\gamma}$ ,

$$\begin{aligned} & P(T > t, U > u | x_j) \\ &= E\{ P(T > t, U > u | \mathbf{x}) | x_j \} \\ &= E\{ P(T > t | \mathbf{x})P(U > u | \mathbf{x}) | x_j \} \\ &= E[\exp\{-\Lambda_T(t)e^{\boldsymbol{\beta}'\mathbf{x}}\}\exp\{-\Lambda_U(u)e^{\boldsymbol{\beta}'\mathbf{x}}\} | x_j] \\ &= E(\exp[-\{\Lambda_T(t) + \Lambda_U(u)\}e^{\boldsymbol{\beta}'\mathbf{x}}] | x_j) \\ &= E(\exp[-\{\Lambda_T(t) + \Lambda_U(u)\}e^{\beta_j x_j} e^{\boldsymbol{\beta}'_{(-j)}\mathbf{x}_{(-j)}}] | x_j) \\ &= \varphi_{\boldsymbol{\beta}_{(-j)}}[\{\Lambda_T(t) + \Lambda_U(u)\}e^{\beta_j x_j}]. \end{aligned}$$

From the previous result we have

$$\Lambda_T(t)e^{\beta_j x_j} = \varphi_{\boldsymbol{\beta}_{(-j)}}^{-1}\{P(T > t | x_j)\}$$

and

$$\Lambda_U(u)e^{\beta_j x_j} = \varphi_{\boldsymbol{\beta}_{(-j)}}^{-1}\{P(U > u | x_j)\}.$$

Hence we can obtain the Archimedean copula representation

$$P(T > t, U > u | x_j) = \varphi_{\boldsymbol{\beta}_{(-j)}}[\varphi_{\boldsymbol{\beta}_{(-j)}}^{-1}\{P(T > t | x_j)\} + \varphi_{\boldsymbol{\beta}_{(-j)}}^{-1}\{P(U > u | x_j)\}].$$