

**MATH-449 - Biostatistics**  
**EPFL, Spring 2025**  
**Problem Set 5**

Consider the following filtrations from the lectures:

$$\begin{aligned}\mathcal{F}_t &= \sigma(L_0, N(u), Z(u); 0 \leq u \leq t) && \text{(the information we have)} \\ \mathcal{F}_t^c &= \sigma(L_0, N^c(u); 0 \leq u \leq t) && \text{(what we want to make inference about)} \\ \mathcal{G}_t &= \sigma(L_0, N^c(u), C(u); 0 \leq u \leq t) && \text{(auxiliary filtration to define independent censoring)}\end{aligned}$$

where  $C(t) = I(t \leq T^*)$  and  $T^*$  is the censoring time,  $N^c(t) = I(t \geq T)$  and  $T$  is the event time,  $N(t) = I(t \geq \tilde{T}, D = 1)$  with  $\tilde{T} = \min(T, T^*)$  and  $D = I(T < T^*)$ , and  $Z(t) = I(t \leq \tilde{T})$ . Thus,  $\mathcal{F}_t \subseteq \mathcal{G}_t \supseteq \mathcal{F}_t^c$ .  $L_0$  is the (possibly empty) set of covariates known at time  $t = 0$ .

1. From the lectures we recall that the censoring is independent if the compensator  $\Lambda^c$  of  $N^c$  with respect to  $\mathcal{F}^c$  is also the compensator of  $N^c$  with respect to  $\mathcal{G}$ . This can be rephrased as

$$E[N^c(t)|\mathcal{G}_t] = E[N^c(t)|\mathcal{F}_t^c]. \quad (1)$$

Thus, (1) hold if and only if we have independent censoring. We will often focus on the intensity  $\lambda^c$  instead of the cumulative intensity  $\Lambda^c(t) = \int_0^t \lambda^c(s)ds$ .

In the lectures, you learned that the independent censoring assumption in this context is that the intensity of the observed counting process  $N$  with respect to the observed information  $\mathcal{F}$  is

$$\lambda(t) = Z(t)\alpha(t), \quad (2)$$

where  $\alpha(t)$  is the hazard function: <sup>1</sup>

$$\alpha(t) = \alpha(t, L_0) = \lim_{h \rightarrow 0+} \frac{1}{h} P(t \leq T < t+h | t \leq T, L_0).$$

It turns out that (2) defines the intensity of  $N$  with respect to  $\mathcal{F}$  if and only if <sup>2</sup>

$$\lim_{h \rightarrow 0+} \frac{1}{h} P(t \leq T < t+h | t \leq T, L_0) = \lim_{h \rightarrow 0+} \frac{1}{h} P(t \leq T < t+h | t \leq \tilde{T}, L_0). \quad (3)$$

- a) Show that independent censoring holds if  $T \perp\!\!\!\perp T^* | L_0$ , i.e. if we have random censoring when conditioning on  $L_0$ .
2. Consider the counting process  $N^c$  and suppose that the intensity  $\lambda^{\mathcal{G}}$  of  $N^c$  with respect to  $\mathcal{G}$  is predictable with respect to  $\mathcal{F}^c$ . Show that independent censoring is satisfied. <sup>3</sup>
  3. Suppose the intensity  $\lambda^{\mathcal{G}}(t)$  with respect to  $\mathcal{G}$  is  $I(T \geq t)(2 - I(T^* \geq t))$ .
    - a) Sketch  $\lambda^{\mathcal{G}}$  for the scenario  $T^* < T$ .
    - b) Which of the following is true: The short-term risk of death for a censored individual is
      - (i) higher than
      - (ii) the same as
      - (iii) lower than
 the short-term risk of death among the subjects that are alive and not censored. Can you think of an example where this is the case?

<sup>1</sup>Of course, we may remove  $L_0$  from the conditioning set if  $L_0 = \emptyset$ .

<sup>2</sup>See Fleming and Harrington (1991) for a proof.

<sup>3</sup>Hint: Use the innovation theorem and the fact that, if  $X$  is predictable with respect to  $\mathcal{F}$ , then  $X(t)$  is measurable with respect to  $\mathcal{F}(t-)$ .

- c) Show that independent censoring is not satisfied in this situation.
4. Calculate the Nelson-Aalen estimator  $\hat{H}(t)$ ,  $t \geq 0$ , for the data set below by hand.

$i$	$\tilde{T}_i$	$D_i$
1	2	1
2	2.5	0
3	5	1
4	5.5	0
5	9	1
6	12	1

Draw the result, and use "(,)", "[,]" to indicate the continuity properties of  $\hat{H}$  at the jump times (here "[,]" at a point indicates continuous from the right at that point, "(,)" at a point indicates not continuous from the right at that point, "[,)" at a point indicates continuous from the left at that point, and "(,)" at a point indicates not continuous from the left at that point, ).

5. Suppose we follow  $n$  individuals over a study period. We will now consider an estimator of the survival probability  $P(T > t)$  as a function of  $t$ . To formulate the estimator, we introduce the variables  $\{\tilde{T}_i, D_i\}_{i=1}^n$ , where  $D_i = 1$  if subject  $i$  dies in the study period (so that  $\tilde{T}_i = T_i$ ) and  $D_i = 0$  if subject  $i$  is censored at  $\tilde{T}_i$  (so that  $T_i > \tilde{T}_i$ ). The estimator, which is called the Kaplan-Meier estimator, then takes the form<sup>4</sup>

$$\hat{S}(t) = \prod_{j: T_j \leq t, D_j = 1} \left(1 - \frac{1}{Z(T_j)}\right),$$

so that the product is over the observed failure times, and  $Z(t) = \sum_{i=1}^n Z_i(t)$  is the number of individuals at risk (i.e. alive and not censored) just before  $t$ . Here,  $Z_i(t)$  is 1 if subject  $i$  is at risk just before  $t$ , and 0 otherwise.<sup>5</sup>

- a) Suppose there is no censoring, i.e. that all individuals are followed up over the entire study period. Show that then  $\hat{S}(t) = 1 - \hat{F}(t)$ , where  $\hat{F}$  is the *empirical distribution function*

$$\hat{F}(t) = \frac{1}{n} \sum_{i=1}^n I(T_i \leq t).$$

- b) A student (not enrolled in MATH-449 - Biostatistics) gets inspired by the relationship between the Kaplan-Meier estimator and the empirical distribution function. He reasons that, if he modifies the sample by just removing the subjects that are censored during the follow-up period, he can estimate the survival function by 1 minus the empirical distribution function of the modified sample. His proposed estimator is

$$\hat{S}^*(t) = 1 - \hat{F}^*(t),$$

where  $\hat{F}^*(t) = \frac{1}{n^*} \sum_{i=1}^n I(T_i \leq t, D_i = 1)$ , and  $n^* = \sum_{i=1}^n I(D_i = 1)$ .

Argue that the estimator  $\hat{S}^*$  will fail to estimate  $S$  in the presence of censoring, even if we have independent censoring.

<sup>4</sup>As in the lectures, we only consider the case without ties; the estimator looks slightly different if some event times are tied.

<sup>5</sup>In the lectures we will see that the Kaplan-Meier estimator is a consistent estimator under the independent censoring assumption. By consistent we mean that, for any  $\epsilon > 0$ ,  $\lim_{n \rightarrow \infty} P\left(\sup_{s \leq \tau} |\hat{S}(s) - S(s)| \geq \epsilon\right) = 0$ , where  $\tau$  is the end of the study period.

6. Prove the following result:

**Theorem 1 (identification under independent censoring)** *Under independent censoring, the intensity of the right-censored counting process  $N_i$  can be written as*

$$\lambda_i(t)dt = Z_i(t)\alpha_i(t)dt$$

where  $Z_i(t) = I(t \leq \tilde{T}_i)$  and  $\alpha_i$  is the hazard of the "complete" counting process

$$\lambda_i^c(t)dt = Z_i^c(t)\alpha_i(t)dt$$

where  $Z_i^c(t) = I(t \leq T_i)$ .

As a hint: use the Innovation theorem:

**Theorem 2 (Innovation theorem)** *An intensity  $\lambda_i^{\mathcal{F}''}(t)$  with respect to a filtration  $\{\mathcal{F}_t''\}$  such that  $\{\mathcal{F}_t'\} \supseteq \{\mathcal{F}_t''\}$ , satisfies*

$$\lambda_i^{\mathcal{F}''}(t) = \mathbb{E}(\lambda_i^{\mathcal{F}'}(t) \mid \mathcal{F}_{t-}'').$$