

## Homework 9

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1

$$h(x) = \frac{2x}{(1+x^2)}$$

Let  $H(x)$  be cumulative hazard function,  $S(x)$  be Survival function, and  $f(x)$  be density function.

$$\begin{aligned} H(x) &= \int_0^x h(t) dt \\ &= \int_0^x \frac{2}{1+t^2} dt = \left[ \log(1+t^2) \right]_0^x \\ &= \log(1+x^2) \end{aligned}$$

$$S(x) = \exp\{-H(x)\} = \exp\{-\log(1+x^2)\} = \frac{1}{1+x^2}$$

$$\begin{aligned} h(x) = \frac{f(x)}{S(x)} &\Rightarrow f(x) = h(x) \times S(x) = \frac{2x}{(1+x^2)} \times \frac{1}{(1+x^2)} \\ &= \frac{2x}{(1+x^2)^2} \end{aligned}$$

2

$t_i$	$n_i$	$d_i$	$c_i$	$\hat{\lambda}_i = \frac{d_i}{n_i}$	$\hat{S}(t)$
1	10	1	0	1/10	$1 \times (1 - 1/10) = 9/10$
2	9	2	0	2/9	$9/10 \times (1 - 2/9) = 7/10$
4	7	0	1	0/7	$7/10 \times (1 - 0/7) = 7/10$
5	6	0	1	0/6	$7/10 \times (1 - 0/6) = 7/10$
6	5	1	0	1/5	$7/10 \times (1 - 1/5) = 14/25$
7	4	0	1	0/4	$14/25 \times (1 - 0/4) = 14/25$
8	3	0	1	0/3	$14/25 \times (1 - 0/3) = 14/25$
9	2	0	1	0/2	$14/25 \times (1 - 0/2) = 14/25$
10	1	0	1	0/1	$14/25 \times (1 - 0/1) = 14/25$

 $t_i$ : time to event $n_i$ : number of observations at  $t_i$  $d_i$ : number of events at  $t_i$  $c_i$ : number of censored observations at  $t_i$  $\hat{\lambda}_i$ : conditional probability of event at  $t_i$  given that the individual is still alive at  $t_i$  $\hat{S}(t)$ : Kaplan-Meier estimator of survival function

(a) Kaplan-Meier estimate of the survival function

$$\hat{S}(t) = \prod_{i=1}^k (1 - \hat{\lambda}_i) = \prod_{i=1}^k (1 - d_i/n_i), \quad t_k \leq t < t_{k+1}$$

The values for each  $t_k \leq t < t_{k+1}$  ( $k=1, 2, \dots, 10$ ) are stated in the above table.

(b) Nelson-Aalen estimate of the cumulative hazard function

$$\hat{H}(t) = \begin{cases} 0 & 0 \leq t < t_1 \\ \sum_{t_i \leq t} d_i/n_i & t \geq t_1 \end{cases} \Rightarrow$$

$t_i$	$d_i/n_i$	$\hat{H}(t)$
1	1/10	1/10 = 0.1
2	2/9	1/10 + 2/9 = 0.32
4	0/7	0.32
5	0/6	0.32
6	1/5	0.32 + 1/5 = 0.52
7	0/4	0.52
8	0/3	0.52
9	0/2	0.52
10	0/1	0.52

(c) Fleming-Harrington estimate of the survival function

$t_i$	$\exp[-\hat{H}(t_i)]$
1	0.905
2	0.726
4	0.726
5	0.726
6	0.595
7	0.595
8	0.595
9	0.595
10	0.595

### 3

tongue data contains the following columns:

- **type** Tumor DNA profile (1 = Aneuploid Tumor, 2 = Diploid Tumor)
- **time** Time to death or on-study time, weeks
- **delta** Death indicator (0 = alive, 1 = dead)

Here we consider individuals with **delta** = 0 as right censored ones.

```
# data import
data("tongue")

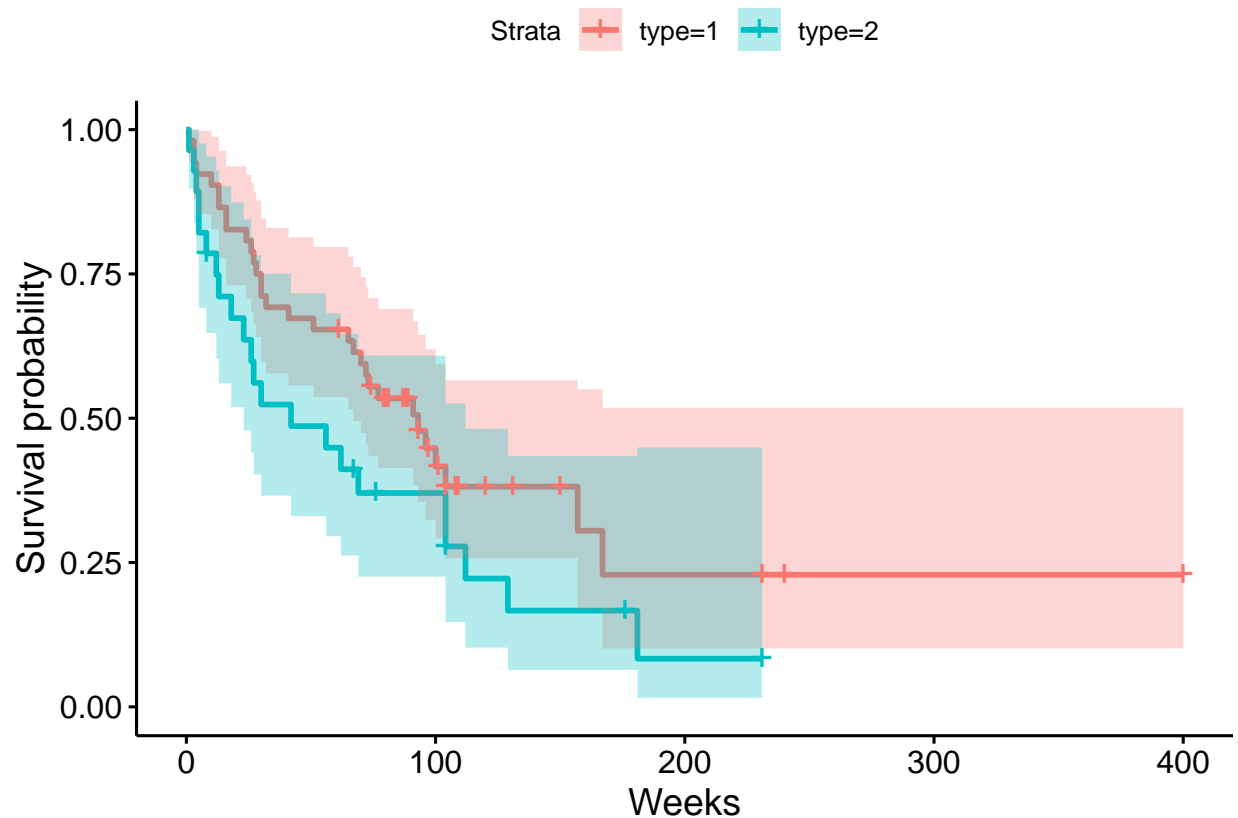
str(tongue)
```

```
## 'data.frame': 80 obs. of 3 variables:
## $ type : int 1 1 1 1 1 1 1 1 1 1 ...
## $ time : int 1 3 3 4 10 13 13 16 16 24 ...
## $ delta: int 1 1 1 1 1 1 1 1 1 1 ...
```

```
# check censored
Surv(tongue$time, tongue$delta, type = "right")
```

```
## [1] 1 3 3 4 10 13 13 16 16 24 26 27 28 30 30
## [16] 32 41 51 65 67 70 72 73 77 91 93 96 100 104 157
## [31] 167 61+ 74+ 79+ 80+ 81+ 87+ 87+ 88+ 89+ 93+ 97+ 101+ 104+ 108+
## [46] 109+ 120+ 131+ 150+ 231+ 240+ 400+ 1 3 4 5 5 8 12 13
## [61] 18 23 26 27 30 42 56 62 69 104 104 112 129 181 8+
## [76] 67+ 76+ 104+ 176+ 231+
```

```
# plot Kaplan-Meier curve of survival function
ggsurvplot(survfit(Surv(time, delta) ~ type, data = tongue, conf.type = "log"), conf.int = TRUE, xlab =
```



The Kaplan-Meier curve of survival function and its pointwise 95% CI using the log transformation is shown above.

```
# estimated 1-year (52 weeks) survival rate and 95% CI
KM <- survfit(Surv(time, delta) ~ type, data = tongue, conf.type = "log")

summary(KM, times = 52)
```

```
## Call: survfit(formula = Surv(time, delta) ~ type, data = tongue, conf.type = "log")
##
##               type=1
##      time      n.risk  n.event  survival  std.err lower 95% CI
##    52.000     34.000    18.000    0.654    0.066    0.537
## upper 95% CI
##    0.797
##
##               type=2
##      time      n.risk  n.event  survival  std.err lower 95% CI
##    52.000     13.000    14.000    0.4864   0.0961    0.3302
## upper 95% CI
##    0.7164
```

Given the output, individuals with Aneuploid Tumor (`type = 1`) have an estimated 1-year (52 weeks) survival rate of 0.654 (95% CI: 0.537 - 0.797) and individuals with Diploid Tumor (`type = 2`) have that of 0.4864 (95% CI: 0.3302 - 0.7164).