Homework 2

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Due date: Thursday, October 11

1. Show that (algebraically) in the absence of censoring $\hat{S}_{\text{KM}}(t) = \hat{S}_{\text{e}}(t)$.

$$\hat{S}_{\text{KM}}(t) = \prod_{t_{(i) \le t}} \frac{n_i - d_i}{n_i} = \frac{n_1 - d_1}{n_1} \cdot \frac{n_2 - d_2}{n_2} \cdot \frac{n_3 - d_3}{n_3} \cdot \cdot \cdot \frac{n_t - d_t}{n_t} = \frac{n_1 - d_1}{n_1} \cdot \frac{n_2 - d_2}{n_1 - d_1} \cdot \frac{n_3 - d_3}{n_2 - d_2} \cdot \cdot \cdot \cdot \frac{n_t - d_t}{n_{t-1} - d_{t-1}}$$

$$= \frac{n_t - d_t}{n_1} = \frac{n_1 - \sum_{t_{(i)} \le t} d_i}{n_1} = \frac{n_{t+1}}{n_1} = \hat{S}_{e}(t)$$

2. In the absence of censoring, show that the Greenwood Formula (page 30 on note 2) can be reduced to

$$\frac{\hat{S}_{\text{KM}}(t) \times \{1 - \hat{S}_{\text{KM}}(t)\}}{n}.$$

You might assume there are no ties among the observations.

$$Var(\hat{S}_{\text{KM}}(t)) = Var(\frac{n_{t+1}}{n}) = \frac{Var(n_{t+1})}{n^2} = \frac{n \cdot (1 - \frac{n_{t+1}}{n}) \cdot \frac{n_{t+1}}{n}}{n^2} = \frac{\hat{S}_{\text{KM}}(t) \times \{1 - \hat{S}_{\text{KM}}(t)\}}{n}$$

Since $Var(n_{t+1})$ follows binomial distribution with parameters $(1 - \frac{n_{t+1}}{n})$ and $(\frac{n_{t+1}}{n})$.

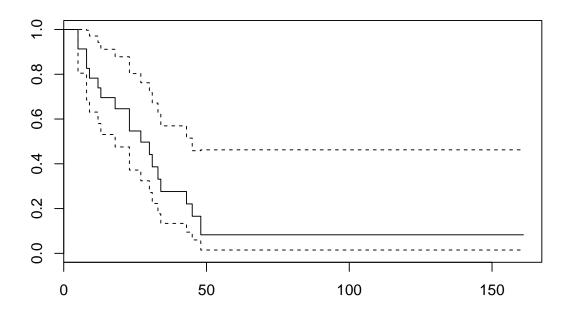
3. Consider the Leukemia data from the survival package:

In here, each row represent one patient. aml is the observed survival time, status is the censoring indicator (1 = event, 0 = censored), and x is the treatment indicator. We will ignore the treatment indicator for now.

- a. Plot the Kaplan-Meier survival curve for the data.
- > library(survival)

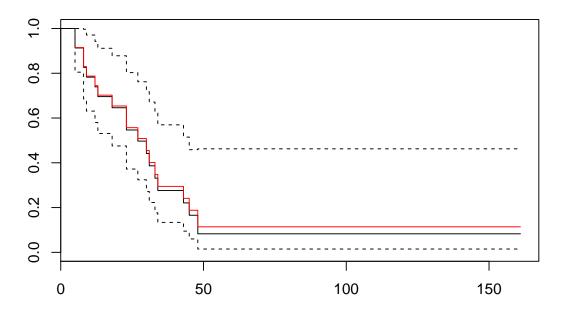
Warning: package 'survival' was built under R version 3.4.4

- > km <- survfit(Surv(time, status) ~ 1, dat = aml)
- > plot(km)



b. Add the Nelson-Aalen survival curve to the Kaplan-Meier plot from (3a).

```
> cox <- coxph(Surv(time, status) ~ 1, dat = aml)
> H0 <- basehaz(cox)
> plot(km)
> lines(H0$time, exp(-H0$hazard), 's', col = 2)
```



4. The expected survival time for the Leukemia data in #(3) does not exist because the last observation is a censored event. An alternative is to look instead of looking at the expected survival time, an alternative is to look at the restricted mean survival time. Compute E(T|T<161) based on the survival curve in (3a).

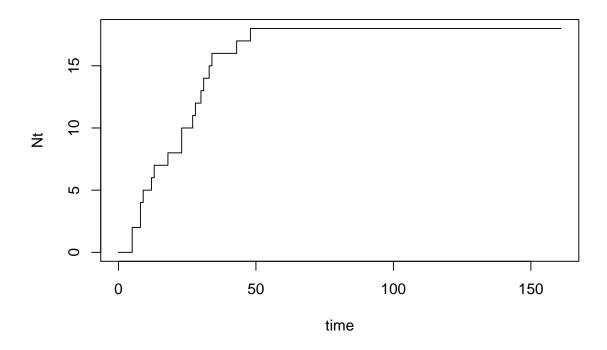
I calculate the integration of the survival function. (I cannot figure out how to do it automatically in R, so I typed in each interval by hand.) The result is shown as follows:

```
> diff_time<-c(5,3,1,3,1,0,5,5,4,3,0,1,2,1,9,2,3,113,0)
> Surv_rate<-c(1,km$surv)
> Surv_rate
```

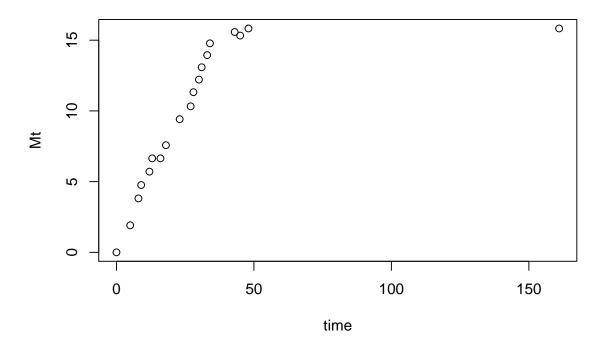
- [1] 1.00000000 0.91304348 0.82608696 0.78260870 0.73913043 0.69565217
- [7] 0.69565217 0.64596273 0.54658385 0.49689441 0.49689441 0.44168392
- [13] 0.38647343 0.33126294 0.27605245 0.22084196 0.16563147 0.08281573
- [19] 0.08281573
- > sum(diff_time*Surv_rate)
- [1] 36.36439
- 5. Let $N_i(t)$ be the number of events over time interval (0,t] for the *i*th patient in #(3). Let $N(t) = \sum_{i=1}^{n} N_i(t)$ be the aggregated counting process.

```
> time <- c(0, H0\$time)
```

- > Nt <- c(0,2,4,5,6,7,7,8,10,11,12,13,14,15,16,17,17,18,18)
- > plot(time,Nt, type="s")



- b. Plot M(t), where $M(t)=N(t)-\hat{H}(t)$ and $\hat{H}(t)$ is the Nelson-Aalen estimator for the cumulative hazard function.
- > Ht <- c(0, H0\$hazard)
- > Mt<- Nt-Ht
- > plot(time,Mt)

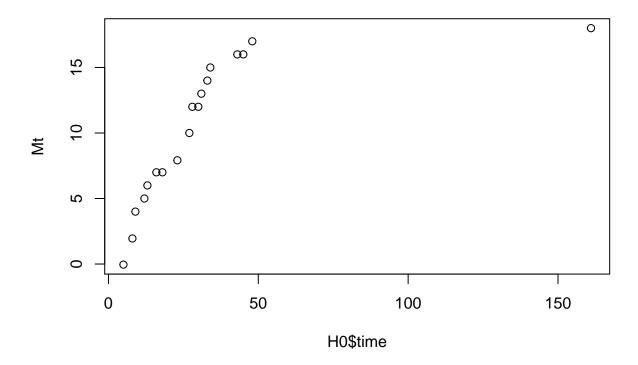


• Note on 5b: After giving some thought, I think it is more meaningful to plot $dM(t) = dN(t) - \hat{h}(t)dt$. Both plots will receive full credit for 5b.

Personally, I don't like this statement. A plot of dM(t) will look nicer since all the points are near 0. However, if we examine carefully in this case, we will find almost all points of dM(t) are above 0, which exactly explains why this M(t) function is (almost) increasing. But ideally, we should have dM(t) randomly dispersed around zero. As a result, a plot of dM(t) will still be abnormal, though it looks more preferable at first glance.

Based on our discussion in Tuesday class, I directly calculate M(t) from the definition: $M(t) = N(t) - \int h(t)Y(t)dt$, and use $\Delta H(t)$ as the approximation for $\int h(t)dt$. That gives the following result:

```
> Ht2 <- c(0,H0$hazard[1:17])
> delta_Ht <- H0$hazard-Ht2
>
> Nt <- c(2,4,5,6,7,7,8,10,11,12,13,14,15,16,17,17,18,18)
> Mt <- Nt-delta_Ht*km$n.risk
>
> plot(H0$time,Mt)
```



The two graphs are almost the same, the difference (I guess) is due to approximation error.