homework1

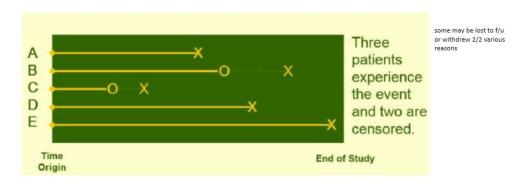
Question #1a, 3.2

- 3.2 A large number of disease-free individuals were enrolled in a study beginning January 1, 1970, and were followed for 30 years to assess the age at which they developed breast cancer. Individuals had clinical exams every 3 years after enrollment. For four selected individuals described below, discuss in detail, the types of censoring and truncation that are represented.
- (a) A healthy individual, enrolled in the study at age 30, never developed breast cancer during the study.
- a. type I, right censored since event did not occur prior to some prespecified time
- (b) A healthy individual, enrolled in the study at age 40, was diagnosed with breast cancer at the fifth exam after enrollment (i.e., the disease started sometime between 12 and 15 years after enrollment).
- b. interval censoring since event time is only known to occur between some interval follow up times
- (c) A healthy individual, enrolled in the study at age 50, died from a cause unrelated to the disease (i.e., not diagnosed with breast cancer at any time during the study) at age 61.
- c. Here if we assume the death was unrelated to breast cancer then is likely random censoring since the time at which the patient would have died from breast cancer is unknown

before had fixed censoring, in type I and type II it looked like the investigator had a little bit of control over the type of censoring in real life people may be lost (e.g. move, lost to f/u etc..)

Random censoring:

Random censoring occurs when follow-up is terminated for reasons that are not under the control of the investigator. (e.g. withdrawals, loss to follow-up)



In most studies of interest, the censoring scheme is a combination of random and Type I censoring: real life has both types

- Some patients are lost to follow-up during the study and become randomly censored their censoring time can be thought of as a random variable
- Other patients may be type I censored when the study period ends
- (d) An individual, enrolled in the study at age 42, moved away from the community at age 55 and was never diagnosed with breast cancer during the period of observation.
- d. This again seems to be random censoring since patient dropped out of study with no apparent reasons related to the disease process of breast cancer.
- (e) Confining your attention to the four individuals described above, write down the likelihood for this portion of the study.
- e. Likelihood for four observations would be the product of the four likelihoods using the constructed likelihood function obs 1; i=1

$$L = \prod_{i=1}^{n} [b(t_i)]^{\delta_i} \exp[-H(t_i)]$$

 $[h(t_1)]exp[-H(t_1)]$

obs 2; i=2

$$\prod_{i \in I} [S_i(L_i) - S_i(R_i)]$$

$1-[h(t_2)]exp[-H(t_2)]-[h(t_2)]exp[-H(t_2)]$

obs 3; i=3

Textbook section 3.5 example 3.10 seems to suggest that the formula for random censoring is:

$$L \propto \prod_{i=1}^{n} [f(t_i)]^{\delta_t} [S(t_i)]^{1-\delta_t}. \tag{3.5.6}$$

which appears nearly identical to that for right censoring

$$L \propto \prod_{i=1}^{n} [f_i(t)]^{\delta} [S_i(t)]^{1-\delta}$$

however the lecture notes mention that for right censoring we use fi not f So for random censoring do we use f(t3) or f3(t) or are they the same

Assuming the same then random censoring seems similar to right censoring for the calculation

$[h(t_3)]exp[-H(t_3)]$

This is further backed up by lecture notes

Does Type of Censoring Affect Survival Methods?

- Standard methods of survival analysis do not distinguish among Type I,
 Type II, or Random censoring

 theory behind this is explained in pages 75-77 of textbook
- With Type I or Type II censoring, standard methods are not biased
- The analytic tools we will use assume that, if random censoring occurs, it is *non-informative* or unrelated to the reason for failure.

make assumption in our class that random censoring may occur but it is not related to the cause of failure, if it does contribute then need more sophisticated form of analysis obs 4; i=4

 $[h(t_4)]exp[-H(t_4)]$

Thus, for e the final likelihood is:

 $[h(t_1)] \exp[-H(t_1)] * 1-[h(t_2)] \exp[-H(t_2)] - [h(t_2)] \exp[-H(t_2)] * [h(t_3)] \exp[-H(t_3)] * [h(t_4)] \exp[-H(t_4)]$

Question #1b, 3.3

An investigator, performing an animal study designed to evaluate the effects of vegetable and vegetable-fiber diets on mammary carcinogenesis risk, randomly assigned female Sprague-Dawley rats to five dietary groups (control diet, control diet plus vegetable mixture, 1; control diet plus vegetable mixture, 2; control diet plus vegetable-fiber mixture, 1; and control diet plus vegetable-fiber mixture, 2). Mammary tumors were induced by a single oral dose (5 mg dissolved in 1.0 ml. corn oil) of 7,12-dimethylbenz(α)anthracene (DMBA) administered by intragastric intubation, i.e., the starting point for this study is when DMBA was given.

Starting 6 weeks after DMBA administration, each rat was examined once weekly for 14 weeks (post DMBA administration) and the time (in days) until onset of the first palpable tumor was recorded. We wish to make an inference about the marginal distribution of the time until a tumor is detected. Describe, in detail, the types of censoring that are represented by the following rats.

(a) A rat who had a palpable tumor at the first examination at 6 weeks after intubation with DMBA.

Left censoring since to have a palpable tumor carcinogenesis must have been occurring before the DBMA was administered.

The definition of left censoring is that the event of interest occurred before the study started but it is unknown when the event occurred

(b) A rat that survived the study without having any tumors.

Right censored since we only know that the subject did not have the event during the observation period

(c) A rat which did not have a tumor at week 12 but which had a tumor at week 13 after inturbation with DMBA.

Interval censoring since the event occurred between follow-up time points but it is not clear at what point between 12 and 13 weeks post DMBA administration when the tumor development occurred

(d) A rat which died (without tumor present and death was unrelated to the occurrence of cancer) at day 37 after intubation with DMBA.

Random censoring since the death of the subject is not though to be related to the disease process of tumor formation. The time that the subject would have developed the tumor is unknown

(e) Confining our attention to the four rats described above, write down the likelihood for this portion of the study.

obs 1; i=1

$$(1-S_i(C_i))$$

 $1-[h(t_1)]exp[-H(t_1)]$

obs 2; i=2

$$L = \prod_{i=1}^{n} [b(t_i)]^{\delta_i} \exp[-H(t_i)]$$

 $[h(t_2)]exp[-H(t_2)]$

obs 3; i=3

$$\prod_{i \in I} [S_i(L_i) - S_i(R_i)]$$

 $1-[h(t_3)]exp[-H(t_3)]-[h(t_3)]exp[-H(t_3)]$

obs 4; i=4

As mentioned before, the textbook section 3.5 example 3.10 seems to suggest that the formula for random censoring is

$$L \propto \prod_{i=1}^{n} [f(t_i)]^{\delta_t} [S(t_i)]^{1-\delta_t}. \tag{3.5.6}$$

 $[h(t_4)]exp[-H(t_4)]$

Thus for e the final likelihood is:

 $1-[h(t_1)] exp[-H(t_1)]^*[h(t_2)] exp[-H(t_2)]^* \\ 1-[h(t_3)] exp[-H(t_3)]-[h(t_3)] exp[-H(t_3)]^*[h(t_4)] exp[-H(t_4)]$

Question #1c, 3.6 a

3.6 The following data consists of the times to relapse and the times to death following relapse of 10 bone marrow transplant patients. In the sample patients 4 and 6 were alive in relapse at the end of the study and patients 7–10 were alive, free of relapse at the end of the study. Suppose the time to relapse had an exponential distribution with hazard rate λ and the time to death in relapse had a Weibull distribution with parameters θ and α .

| Patient | Relapse Time (months) | Death Time (months) |
|------------------|--------------------------|------------------------------------|
| 1 | 5 | 11 |
| 2 | 8 | 12 |
| 3 | 12 | 15 |
| 3 4 5 6 | 24 | 33 ⁺ 45 |
| 5 | 32 | 45 |
| 6 | 17 | 28 ⁺ 16 ⁺ |
| 7 | 16 ⁺ | 16+ |
| 8 | 17+ | 17+ |
| 9 | 19+ | 19+ |
| 10 | 30+ | 30+ |

⁺ Censored observation

(a) Construct the likelihood for the relapse rate λ .

D= # that relapsed during study period = 6 PT = total time in months = 180 Likelihood for relapse rate

$$\hat{\lambda} = \frac{D}{PT}$$

> sum(c(5,8,12,24,32,17,16,17,19,30))

[1] 180

Thus lambda_hat = D/PT = 6/180=0.0333/month

Question #1d

Give examples of right censored, left censored, interval-censored, left-truncated data from your field of study. These examples should not be from the Klein & Moeschberger textbook or from the video lectures.

Agriculture is an industry which I grew up in and remain interested in so am providing examples from this context

Right censored

An example of right censored observations in agriculture could be time to hybrid seed germination where at the end of the study some seeds with specific hybrid genotypes have not germinated.

Left censored

Study at what age calves wean from their mothers with a bull present in the herd but find that at time of study initiation some calves have already weaned from their mother but don't know when.

Left truncated

Study at what age vaccinated animals become infected where animals randomly enter the study at different ages and times.

Interval censured

Visit orchards every two weeks to study at what age trees first produce fruit and find trees which produced between previous and current visit but do not know exactly when produced between those timepoints.

Question #2

Researchers wish to explore the efficacy of triple-drug combinations of antiretroviral therapy for treatment of HIV-infected patients. Because of limitations on potency and the continuing emergence of drug resistance seen with the use of currently available antiretroviral agents in monotherapy and two-drug regimens, triple combination regimens should represent a more promising approach to maximize antiviral activity, maintain long-term efficacy, and reduce the incidence of drug resistance. Towards this end, investigators performed a randomized study comparing AZT + zalcitabine (ddC) versus AZT + zalcitabine (ddC) + saquinavir. The data, time from administration of treatment (in days) until the CD4 count reached a pre-specified level, is given below for the two groups:

```
AZT + zalcitabine (ddC):
4+, 6, 11, 12, 32, 35, 38+, 39, 45, 49, 75, 80, 84, 85, 87, 102, 180+
AZT + zalcitabine (ddC) + saquinivir:
2, 3, 4, 12, 22, 48, 51+, 56+, 80, 85, 90, 94+, 160, 171, 180, 180+, 238
```

- a. For both groups separately, construct a data layout (similar to what was done in lecture slides) containing the unique, ordered event times, the number of events that occurred at those unique event times, the number of censored observations in the relevant interval, the number in the risk set at that time, and the Kaplan-Meier estimate of the survival curve at that time. What is the median survival time in the two groups? Will you be comfortable reporting the mean survival time in the two groups?
 - a) KM table using custom function in R for azt_ddc group

| • | orderedEventTimes_tj | eventsAtEventTime_ej | censoredObservationsInInterval_cj | inRiskSetAtTime_nj | kaplanMeirSurvivalCurveAtTime_s_tj- | c_tj-1 | kaplanMeirSurvivalCurveAtTime_s_tj |
|----|----------------------|----------------------|-----------------------------------|--------------------|-------------------------------------|--------|------------------------------------|
| 1 | 0 | 0 | 0 | 17 | - | 17/17 | 1 |
| 2 | 6 | 1 | 0 | 16 | 1 | 15/16 | 0.938 |
| 3 | 11 | 1 | 0 | 15 | 0.938 | 14/15 | 0.875 |
| 4 | 12 | 1 | 0 | 14 | 0.875 | 13/14 | 0.812 |
| 5 | 32 | 1 | 0 | 13 | 0.812 | 12/13 | 0.75 |
| 6 | 35 | 1 | 0 | 12 | 0.75 | 11/12 | 0.688 |
| 7 | 39 | 1 | 0 | 10 | 0.688 | 9/10 | 0.619 |
| 8 | 45 | 1 | 0 | 9 | 0.619 | 8/9 | 0.55 |
| 9 | 49 | 1 | 0 | 8 | 0.55 | 7/8 | 0.481 |
| 10 | 75 | 1 | 0 | 7 | 0.481 | 6/7 | 0.412 |
| 11 | 80 | 1 | 0 | 6 | 0.412 | 5/6 | 0.343 |
| 12 | 84 | 1 | 0 | 5 | 0.343 | 4/5 | 0.274 |
| 13 | 85 | 1 | 0 | 4 | 0.274 | 3/4 | 0.206 |
| 14 | 87 | 1 | 0 | 3 | 0.206 | 2/3 | 0.137 |
| 15 | 102 | 1 | 0 | 2 | 0.137 | 1/2 | 0.068 |

b) KM table using built in function in R for azt_ddc group

```
## Call: survfit(formula = Surv(as.numeric(sub("+", "", azt_ddc, fixed = TRUE)),
       ifelse(grepl("+", azt_ddc, fixed = TRUE), 0, 1)) ~ 1, conf.type = "log-log")
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
                          0.9375 0.0605
                                               0.63235
                                                              0.991
       6
             16
                      1
##
      11
             15
                          0.8750 0.0827
                                               0.58598
                                                              0.967
##
             14
                          0.8125 0.0976
                                               0.52460
##
      12
                                                              0.935
      32
             13
                          0.7500 0.1083
                                               0.46343
                                                              0.898
##
      35
             12
                          0.6875 0.1159
                                               0.40460
                                                              0.856
##
                          0.6188 0.1230
##
      39
             10
                                               0.33929
                                                              0.808
      45
              9
                          0.5500 0.1271
                                               0.27933
                                                              0.756
##
##
      49
                          0.4813 0.1285
                                               0.22410
                                                              0.699
      75
                          0.4125 0.1272
                                               0.17339
                                                              0.639
##
              6
                          0.3438 0.1232
                                               0.12728
                                                              0.575
##
      80
              5
                          0.2750 0.1162
                                               0.08617
                                                              0.507
##
      84
```

```
85
                         0.2063 0.1055
                                             0.05082
                                                             0.433
##
##
     87
                         0.1375 0.0900
                                             0.02265
                                                             0.354
              3
    102
                         0.0688 0.0662
                                             0.00443
                                                             0.267
              2
##
                      1
```

Call: survfit(formula = Surv(as.numeric(sub("+", "", azt_ddc, fixed = TRUE)), ifelse(grepl("+", azt_ddc, fixed = TRUE), 0, 1)) ~ 1, conf.type = "log-log") n events median 0.95LCL 0.95UCL

17 14 **49** 32 85

median of 49

The median survival time is defined as the time at t such that S(t) = 1/2. For AZT + DDC it is 49.

a) KM table using custom function in R for azt ddc sag group

| • | ${\sf orderedEventTimes_tj} ^{\scriptsize \scriptsize \diamondsuit}$ | eventsAtEventTime_ej | censoredObservationsInInterval_cj | inRiskSetAtTime_nj [‡] | $kaplan Meir Survival Curve At Time_s_tj-1 \\ \hspace*{2em} ^{\scriptsize \scriptsize \bigcirc}$ | c_tj-1 [‡] | kaplanMeirSurvivalCurveAtTime |
|----|--|----------------------|-----------------------------------|---------------------------------|--|----------------------------|-------------------------------|
| 1 | 0 | 0 | 0 | 17 | - | 17/17 | 1 |
| 2 | 2 | 1 | 0 | 17 | 1 | 16/17 | 0.941 |
| 3 | 3 | 1 | 0 | 16 | 0.941 | 15/16 | 0.882 |
| 4 | 4 | 1 | 0 | 15 | 0.882 | 14/15 | 0.823 |
| 5 | 12 | 1 | 0 | 14 | 0.823 | 13/14 | 0.764 |
| 6 | 22 | 1 | 0 | 13 | 0.764 | 12/13 | 0.705 |
| 7 | 48 | 1 | 0 | 12 | 0.705 | 11/12 | 0.646 |
| 8 | 51 | 1 | 0 | 11 | 0.646 | 10/11 | 0.587 |
| 9 | 56 | 1 | 0 | 10 | 0.587 | 9/10 | 0.528 |
| 10 | 80 | 1 | 0 | 9 | 0.528 | 8/9 | 0.469 |
| 11 | 85 | 1 | 0 | 8 | 0.469 | 7/8 | 0.41 |
| 12 | 90 | 1 | 0 | 7 | 0.41 | 6/7 | 0.351 |
| 13 | 94 | 1 | 0 | 6 | 0.351 | 5/6 | 0.292 |
| 14 | 160 | 1 | 0 | 5 | 0.292 | 4/5 | 0.234 |
| 15 | 171 | 1 | 0 | 4 | 0.234 | 3/4 | 0.176 |
| 16 | 180 | 2 | 0 | 3 | 0.176 | 1/3 | 0.059 |
| 17 | 238 | 1 | 0 | 1 | 0.059 | 0/1 | 0 |

b) KM table using built in function in R for azt_ddc_saq

| ## | time | n.rısk | n.event | survival | sta.err | lower 95% Cl | upper 95% CI |
|----|------|--------|---------|----------|---------|--------------|--------------|
| ## | 2 | 17 | 1 | 0.9412 | 0.0571 | 0.65018 | 0.991 |
| ## | 3 | 16 | 1 | 0.8824 | 0.0781 | 0.60598 | 0.969 |

| ## | 4 | 15 | 1 | 0.8235 | 0.0925 | 0.54713 | 0.939 |
|----|-----|----|---|--------|--------|---------|-------|
| ## | 12 | 14 | 1 | 0.7647 | 0.1029 | 0.48828 | 0.904 |
| ## | 22 | 13 | 1 | 0.7059 | 0.1105 | 0.43148 | 0.866 |
| ## | 48 | 12 | 1 | 0.6471 | 0.1159 | 0.37715 | 0.823 |
| ## | 51 | 11 | 1 | 0.5882 | 0.1194 | 0.32537 | 0.778 |
| ## | 56 | 10 | 1 | 0.5294 | 0.1211 | 0.27617 | 0.730 |
| ## | 80 | 9 | 1 | 0.4706 | 0.1211 | 0.22960 | 0.680 |
| ## | 85 | 8 | 1 | 0.4118 | 0.1194 | 0.18576 | 0.626 |
| ## | 90 | 7 | 1 | 0.3529 | 0.1159 | 0.14483 | 0.570 |
| ## | 94 | 6 | 1 | 0.2941 | 0.1105 | 0.10712 | 0.511 |
| ## | 160 | 5 | 1 | 0.2353 | 0.1029 | 0.07308 | 0.449 |
| ## | 171 | 4 | 1 | 0.1765 | 0.0925 | 0.04348 | 0.383 |
| ## | 180 | 3 | 2 | 0.0588 | 0.0571 | 0.00391 | 0.235 |
| ## | 238 | 1 | 1 | 0.0000 | NaN | NA | NA |

n events median 0.95LCL 0.95UCL

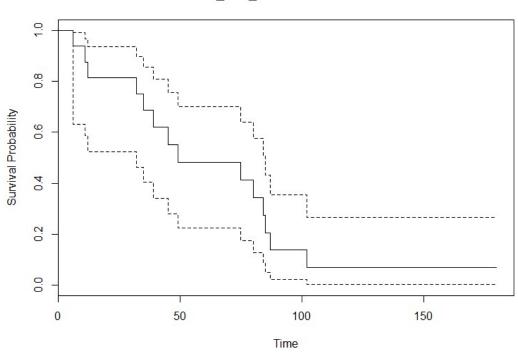
17 17 **80** 12 160

median of 80

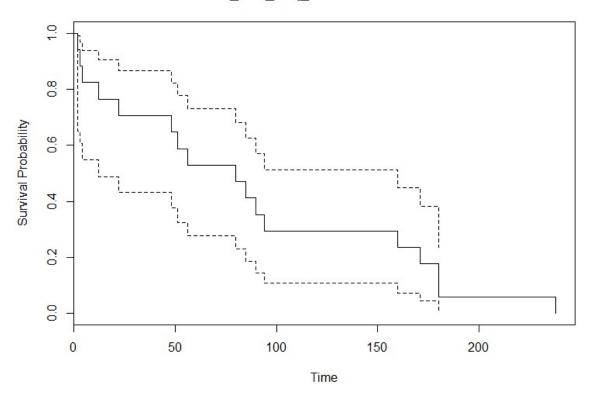
If the survival curve is not continuous at 1/2 (if the survival function is a step function, for example), then the median is taken to be the smallest t such that S(t) <= 1/2. Thus for AZT + DDC + SAQ the median survival would be 80.

The **mean** survival time is only defined if the survival curve goes to zero that is S(inf)=0. Thus, if the last subject is in the study is censored then the survival function will not go to 0 and the area under the curve cannot be calculated. In other words, and in theory, the mean survival cannot be computed when the Kaplan-Meier survival curve does not reach zero. Therefore, I would not feel comfortable reporting the mean for the azt_ddc group since the last observation is censored. I would feel more comfortable reporting the mean survival time in the azt_ddc_saq group since the last observation is not censored and the survival function does go to zero.

azt_ddc_KM survival curve



azt_ddc_saq_KM survival curve



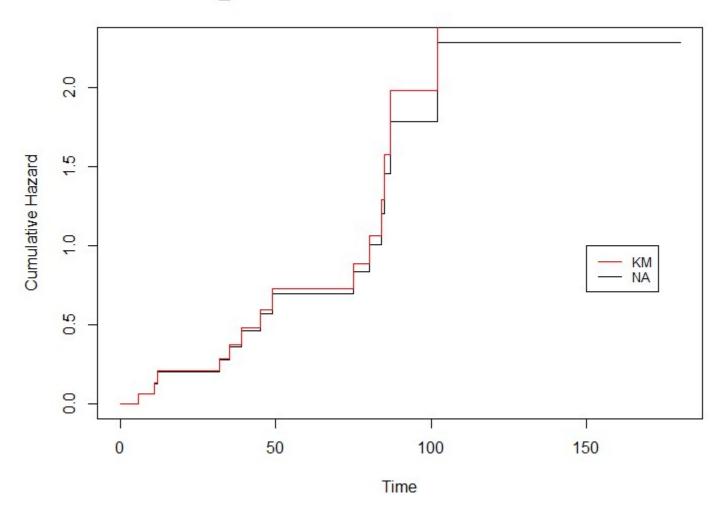
c. For any one group, compute the Nelson Aalen estimates of the cumulative hazard function. Graphically compare $\widehat{H}(t)$ versus $\widetilde{H}(t)$ and comment on this comparison.

cumulative hazard times using Nelson-Aalen technique for azt_ddc group

| orderedEventTimes_tj | eventsAtEventTime_ej | inRiskSetAtTime_nj | cumulativeHazardRate_ht |
|----------------------|----------------------|--------------------|-------------------------|
| 0 | 0 | 17 | 0 |

| 6 | 1 | 16 | 0.062 |
|-----|---|----|-------|
| 11 | 1 | 15 | 0.129 |
| 12 | 1 | 14 | 0.2 |
| 32 | 1 | 13 | 0.277 |
| 35 | 1 | 12 | 0.36 |
| 39 | 1 | 10 | 0.46 |
| 45 | 1 | 9 | 0.571 |
| 49 | 1 | 8 | 0.696 |
| 75 | 1 | 7 | 0.839 |
| 80 | 1 | 6 | 1.006 |
| 84 | 1 | 5 | 1.206 |
| 85 | 1 | 4 | 1.456 |
| 87 | 1 | 3 | 1.789 |
| 102 | 1 | 2 | 2.289 |

azt_ddc NA vs. KM cumulative hazard curves

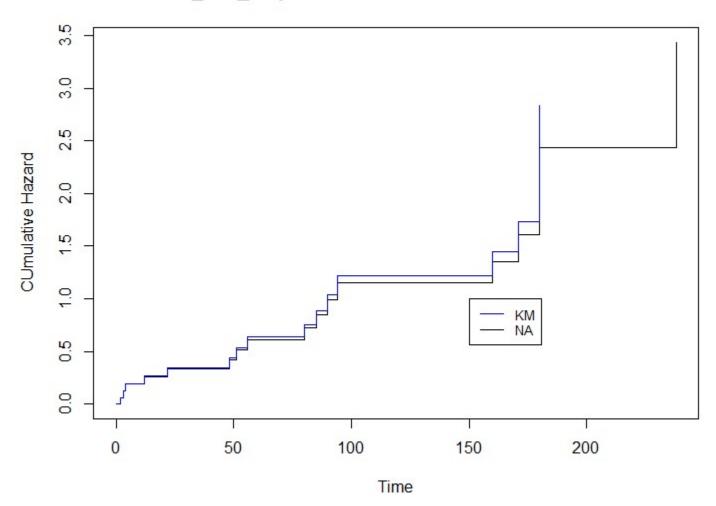


As described in the lecture and text the Nelson-Aalen technique underestimates hazard and overestimates survival compared to the Kaplan-Meier technique

Cumulative hazard times using Nelson-Aalen technique for azt_ddc_saq group

| orderedEventTimes_tj | eventsAtEventTime_ej | inRiskSetAtTime_nj | cumulativeHazardRate_ht |
|----------------------|----------------------|--------------------|-------------------------|
| 2 | 1 | 17 | 0.059 |
| 3 | 1 | 16 | 0.122 |
| 4 | 1 | 15 | 0.189 |
| 12 | 1 | 14 | 0.26 |
| 22 | 1 | 13 | 0.337 |
| 48 | 1 | 12 | 0.42 |
| 51 | 1 | 11 | 0.511 |
| 56 | 1 | 10 | 0.611 |
| 80 | 1 | 9 | 0.722 |
| 85 | 1 | 8 | 0.847 |
| 90 | 1 | 7 | 0.99 |
| 94 | 1 | 6 | 1.157 |
| 160 | 1 | 5 | 1.357 |
| 171 | 1 | 4 | 1.607 |
| 180 | 2 | 3 | 2.274 |
| 238 | 1 | 1 | 3.274 |

azt_ddc_saq NA vs. KM cumulative hazard curves



Again as described in the lecture and text, the Nelson-Aalen technique underestimates hazard and overestimates survival compared to the Kaplan-Meier technique

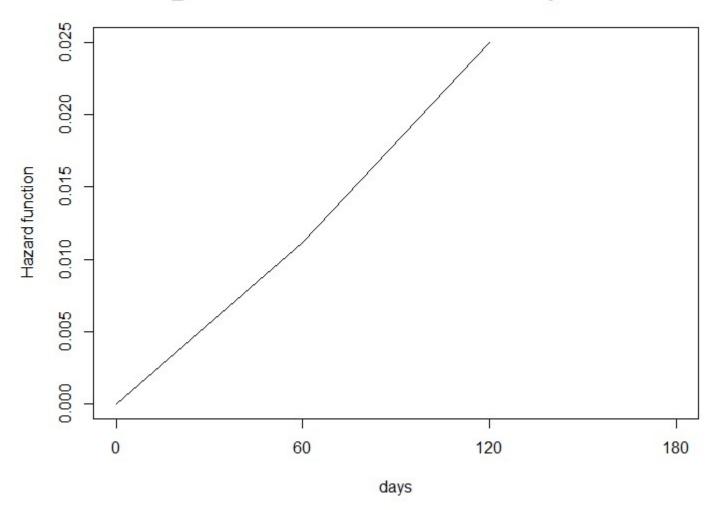
d. Use **PROC LIFETEST** to produce the Life-Table estimate of the survival function and to plot the life-table estimates of the hazard function based on interval widths of 60 days. What does this tell you about the hazard function for the two groups?

Used R function "lifetab" to create table and graphs below instead of PROC LIFETEST

life-table azt_ddc

| ^ | nsubs ‡ | nlost ‡ | nrisk ‡ | nevent [‡] | surv [‡] | pdf [‡] | hazard [‡] |
|--------|---------|---------|---------|---------------------|-------------------|------------------|---------------------|
| 0-4 | 17 | 1 | 16.5 | 0 | 1.00000 | 0.000000000 | 0.00000000 |
| 4-6 | 16 | 0 | 16.0 | 1 | 1.00000 | 0.031250000 | 0.03225806 |
| 6-11 | 15 | 0 | 15.0 | 1 | 0.93750 | 0.012500000 | 0.01379310 |
| 11-12 | 14 | 0 | 14.0 | 1 | 0.87500 | 0.062500000 | 0.07407407 |
| 12-32 | 13 | 0 | 13.0 | 1 | 0.81250 | 0.003125000 | 0.00400000 |
| 32-35 | 12 | 0 | 12.0 | 1 | 0.75000 | 0.020833333 | 0.02898551 |
| 35-38 | 11 | 1 | 10.5 | 0 | 0.68750 | 0.000000000 | 0.00000000 |
| 38-39 | 10 | 0 | 10.0 | 1 | 0.68750 | 0.068750000 | 0.10526316 |
| 39-45 | 9 | 0 | 9.0 | 1 | 0.61875 | 0.011458333 | 0.01960784 |
| 45-49 | 8 | 0 | 8.0 | 1 | 0.55000 | 0.017187500 | 0.03333333 |
| 49-75 | 7 | 0 | 7.0 | 1 | 0.48125 | 0.002644231 | 0.00591716 |
| 75-80 | 6 | 0 | 6.0 | 1 | 0.41250 | 0.013750000 | 0.03636364 |
| 80-84 | 5 | 0 | 5.0 | 1 | 0.34375 | 0.017187500 | 0.0555556 |
| 84-85 | 4 | 0 | 4.0 | 1 | 0.27500 | 0.068750000 | 0.28571429 |
| 85-87 | 3 | 0 | 3.0 | 1 | 0.20625 | 0.034375000 | 0.20000000 |
| 87-102 | 2 | 0 | 2.0 | 1 | 0.13750 | 0.004583333 | 0.0444444 |

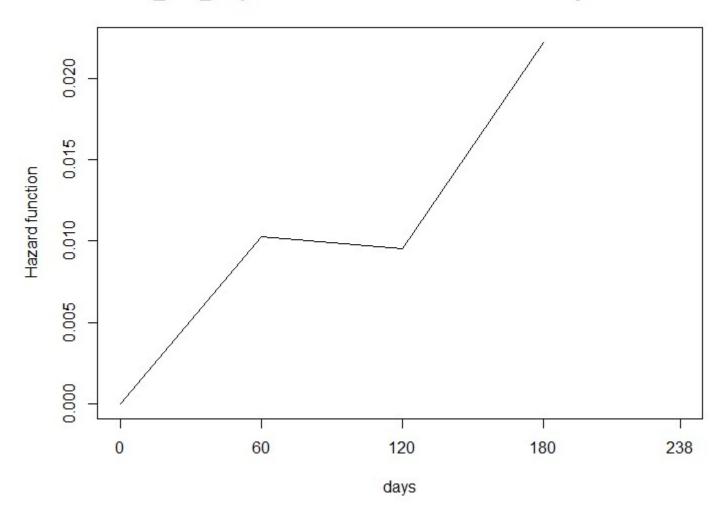
azt_ddc Life Table Hazard Function in 60 day windows



life-table azt_ddc_saq

| ^ | nsubs ÷ | nlost ‡ | nrisk [‡] | nevent ‡ | surv [‡] | pdf [‡] | hazard ‡ | se.surv ÷ | se.pdf | se.hazard |
|---------|---------|---------|--------------------|----------|-------------------|------------------|-------------|------------|--------------|-------------|
| 0-2 | 17 | 0 | 17 | 1 | 1.0000000 | 0.0294117647 | 0.030303030 | 0.00000000 | 0.0285336029 | 0.030289114 |
| 2-3 | 16 | 0 | 16 | 1 | 0.9411765 | 0.0588235294 | 0.064516129 | 0.05706721 | 0.0570672059 | 0.064482553 |
| 3-4 | 15 | 0 | 15 | 1 | 0.8823529 | 0.0588235294 | 0.068965517 | 0.07814249 | 0.0570672059 | 0.068924503 |
| 4-12 | 14 | 0 | 14 | 1 | 0.8235294 | 0.0073529412 | 0.009259259 | 0.09245944 | 0.0071334007 | 0.009252906 |
| 12-22 | 13 | 0 | 13 | 1 | 0.7647059 | 0.0058823529 | 0.008000000 | 0.10287937 | 0.0057067206 | 0.007993597 |
| 22-48 | 12 | 0 | 12 | 1 | 0.7058824 | 0.0022624434 | 0.003344482 | 0.11051017 | 0.0021948925 | 0.003341319 |
| 48-51 | 11 | 0 | 11 | 1 | 0.6470588 | 0.0196078431 | 0.031746032 | 0.11590404 | 0.0190224020 | 0.031710018 |
| 51-56 | 10 | 0 | 10 | 1 | 0.5882353 | 0.0117647059 | 0.021052632 | 0.11936462 | 0.0114134412 | 0.021023453 |
| 56-80 | 9 | 0 | 9 | 1 | 0.5294118 | 0.0024509804 | 0.004901961 | 0.12105782 | 0.0023778002 | 0.004893473 |
| 80-85 | 8 | 0 | 8 | 1 | 0.4705882 | 0.0117647059 | 0.026666667 | 0.12105782 | 0.0114134412 | 0.026607341 |
| 85-90 | 7 | 0 | 7 | 1 | 0.4117647 | 0.0117647059 | 0.030769231 | 0.11936462 | 0.0114134412 | 0.030678062 |
| 90-94 | 6 | 0 | 6 | 1 | 0.3529412 | 0.0147058824 | 0.045454545 | 0.11590404 | 0.0142668015 | 0.045266327 |
| 94-160 | 5 | 0 | 5 | 1 | 0.2941176 | 0.0008912656 | 0.003367003 | 0.11051017 | 0.0008646546 | 0.003346155 |
| 160-171 | 4 | 0 | 4 | 1 | 0.2352941 | 0.0053475936 | 0.025974026 | 0.10287937 | 0.0051879278 | 0.025707619 |
| 171-180 | 3 | 0 | 3 | 2 | 0.1764706 | 0.0130718954 | 0.111111111 | 0.09245944 | 0.0086824989 | 0.068041382 |

azt_ddc_saq Life Table Hazard Function in 60 day windows



The life table plots demonstrate that the addition of the anti-retroviral medication Saquinavir reduces the hazard function of death when compared to the group with only two medications azt and ddc

e. For any one group, use **PROC LIFETEST** to produce 95% confidence intervals and 95% confidence band using the various approaches you learned in class. Comment on your results.

Used R functions "survfit" and "confband" to generate values below

| ^ | time ‡ | n.risk ‡ | n.event | survival | std.err | lower 95% CI | upper \$ 95% CI | lower 95% CB | upper \$ 95% CB |
|----|--------|----------|---------|----------|------------|-----------------|-----------------|-----------------|-----------------|
| 1 | 6 | 16 | 1 | 0.93750 | 0.06051536 | 0.82608827 | 1.0000000 | 0.643728905 | 1.0000000 |
| 2 | 11 | 15 | 1 | 0.87500 | 0.08267973 | 0.72707141 | 1.0000000 | 0.510922015 | 0.9999794 |
| 3 | 12 | 14 | 1 | 0.81250 | 0.09757809 | 0.64209217 | 1.0000000 | 0.425344054 | 0.9985377 |
| 4 | 32 | 13 | 1 | 0.75000 | 0.10825318 | 0.56519837 | 0.9952258 | 0.355310460 | 0.9924805 |
| 5 | 35 | 12 | 1 | 0.68750 | 0.11587810 | 0.49408619 | 0.9566271 | 0.295073863 | 0.9808050 |
| 6 | 39 | 10 | 1 | 0.61875 | 0.12300557 | 0.41908199 | 0.9135481 | 0.242188699 | 0.9636890 |
| 7 | 45 | 9 | 1 | 0.55000 | 0.12710724 | 0.34966048 | 0.8651249 | 0.190995454 | 0.9389813 |
| 8 | 49 | 8 | 1 | 0.48125 | 0.12847323 | 0.28519127 | 0.8120921 | 0.146211957 | 0.9084306 |
| 9 | 75 | 7 | 1 | 0.41250 | 0.12719172 | 0.22540390 | 0.7548949 | 0.107343563 | 0.8721711 |
| 10 | 80 | 6 | 1 | 0.34375 | 0.12318011 | 0.17030262 | 0.6938476 | 0.074193951 | 0.8301174 |
| 11 | 84 | 5 | 1 | 0.27500 | 0.11615588 | 0.12017052 | 0.6293141 | 0.046800569 | 0.7819227 |
| 12 | 85 | 4 | 1 | 0.20625 | 0.10551909 | 0.07566808 | 0.5621798 | 0.025408255 | 0.7268739 |
| 13 | 87 | 3 | 1 | 0.13750 | 0.08999783 | 0.03812115 | 0.4959517 | 0.010431689 | 0.6636443 |
| 14 | 102 | 2 | 1 | 0.06875 | 0.06624337 | 0.01040174 | 0.4544011 | 0.002239724 | 0.5896384 |

As expected the lower and upper CIs (confidence intervals) are in general tighter than the CB (confidence band)

f. Read about the *Redistribute to the Right Algorithm* given in "Theoretical Notes #2" on Page # 102 of the textbook. Then study the example workout given on Page #103. Show a similar workout for the 'AZT + zalcitabine + saquinivir' treatment group data given in the problem statement.

Act_ddc_saq

| ac_saq | observation ‡ | decrement ÷ | action ‡ | S_t ÷ |
|--------|---------------|-------------|------------------------------------|-------|
| 2 | 2 | 0.059 | survival is 1-(0.0588235294117647) | 0.941 |
| 3 | 3 | 0.118 | survival is 1-(0.117647058823529) | 0.882 |
| 4 | 4 | 0.176 | survival is 1-(0.176470588235294) | 0.824 |
| 5 | 12 | 0.235 | survival is 1-(0.235294117647059) | 0.765 |
| 6 | 22 | 0.294 | survival is 1-(0.294117647058824) | 0.706 |
| 7 | 48 | 0.353 | survival is 1-(0.352941176470588) | 0.647 |
| 8 | 51 | 0.412 | survival is 1-(0.411764705882353) | 0.588 |
| 9 | 56 | 0.471 | survival is 1-(0.470588235294118) | 0.529 |
| 10 | 80 | 0.529 | survival is 1-(0.529411764705882) | 0.471 |
| 11 | 85 | 0.588 | survival is 1-(0.588235294117647) | 0.412 |
| 12 | 90 | 0.647 | survival is 1-(0.647058823529412) | 0.353 |
| 13 | 94 | 0.706 | survival is 1-(0.705882352941177) | 0.294 |
| 14 | 160 | 0.765 | survival is 1-(0.764705882352941) | 0.235 |
| 15 | 171 | 0.824 | survival is 1-(0.823529411764706) | 0.176 |
| 16 | 180 | 0.882 | survival is 1-(0.882352941176471) | 0.118 |
| 17 | 238 | 0.941 | survival is 1-(0.941176470588235) | 0.059 |

In table above there were no censored values so the survival function was simply reduced by 1/n for each observation (azt_ddc_saq)

| • | observation [‡] | decrement [‡] | action | S_t * | |
|----|---|----------------------------------|--|--------------|--|
| 2 | 4+ | 0.000 | next time survival is 1-(0 + 0.0588235294117647 + 0.0588 | 1.000 | |
| 3 | 6 | 0.062 | survival is 1-(0.0625) | 0.938 | |
| 4 | 4 11 0.125 s | | survival is 1-(0.125) | 0.875 | |
| 5 | 12 | 0.188 | survival is 1-(0.1875) | 0.812 | |
| 6 | 5 32 0.250 survival is 1-(0.25) | | | 0.750 | |
| 7 | 35 0.312 survival is 1-(0.3125) | | | 0.688 | |
| 8 | 38+ 0.000 next time survival is 1-(0.3125 + 0.0625 + 0.0625*1/10) | | | 0.688 | |
| 9 | 39 0.381 survival is 1-(0.38125) | | survival is 1-(0.38125) | 0.619 | |
| 10 | 45 | 15 0.450 survival is 1-(0.45) | | 0.550 | |
| 11 | 49 | 0.519 survival is 1-(0.51875) | | 0.481 | |
| 12 | 75 | 0.588 | 0.588 survival is 1-(0.5875) | | |
| 13 | 80 | 80 0.656 survival is 1-(0.65625) | | 0.344 | |
| 14 | 84 | 34 0.725 survival is 1-(0.725) | | 0.275 | |
| 15 | 85 | 0.794 survival is 1-(0.79375) | | 0.206 | |
| 16 | 87 | 7 0.862 survival is 1-(0.8625) | | 0.138 | |
| 17 | 102 | 0.931 | 0.931 survival is 1-(0.93125) | | |
| 18 | 180+ | 1.000 | survival is 1-(1) | 0.000 | |

For completeness this is the output for azt_ddc without saq

4.7 Consider a hypothetical study of the mortality experience of diabetics. Thirty diabetic subjects are recruited at a clinic and followed until death or the end of the study. The subject's age at entry into the study and their age at the end of study or death are given in the table below. Of interest is estimating the survival curve for a 60- or for a 70-year-old diabetic.

USING LEFT TRUNCATION AND RIGHT CENSORING

(a) Since the diabetics needed to survive long enough from birth until the study began, the data is left truncated. Construct a table showing the number of subjects at risk, Y, as a function of age.

Left truncated and right censored approach:

| a. | ^ | tj ÷ | ej ‡ | cj ‡ | nj ‡ | c_tj- [‡] | s_tj |
|----|----|------|------|------|------|--------------------|--------|
| | 1 | 0 | 0 | 0 | 21 | | 1 |
| | 2 | 60 | 1 | 0 | 5 | 4/5 | 0.8 |
| | 3 | 62 | 1 | 0 | 9 | 8/9 | 0.7111 |
| | 4 | 63 | 1 | 0 | 10 | 9/10 | 0.64 |
| | 5 | 65 | 2 | 0 | 10 | 8/10 | 0.512 |
| | 6 | 66 | 1 | 0 | 10 | 9/10 | 0.4608 |
| | 7 | 68 | 2 | 0 | 13 | 11/13 | 0.3899 |
| | 8 | 69 | 2 | 2 | 14 | 12/14 | 0.3342 |
| | 9 | 70 | 2 | 0 | 13 | 11/13 | 0.2828 |
| | 10 | 71 | 2 | 0 | 14 | 12/14 | 0.2424 |
| | 11 | 72 | 2 | 1 | 14 | 12/14 | 0.2078 |
| | 12 | 73 | 1 | 1 | 13 | 12/13 | 0.1918 |
| | 13 | 74 | 1 | 1 | 12 | 11/12 | 0.1758 |
| | 14 | 76 | 1 | 1 | 11 | 10/11 | 0.1598 |
| | 15 | 77 | 1 | 0 | 10 | 9/10 | 0.1438 |

where nj is the $\mbox{\tt\#}$ at risk Y as a function of age tj. For example at age 62 there are 9 at risk

(b) Estimate the conditional survival function for the age of death of a diabetic patient who has survived to age 60.

Using above table conditional survival for the age of death for a diabetic patient who has survived to age 60 is 0.8

(c) Estimate the conditional survival function for the age of death of a diabetic patient who has survived to age 70.

Using above table conditional survival for the age of death for a diabetic patient who has survived to age 70 is 0.283

(d) Suppose an investigator incorrectly ignored the left truncation and simply treated the data as right censored. Repeat parts a-c.

USING RIGHT CENSORING WHLE IGNORING LEFT CENSORING

(a) Since the diabetics needed to survive long enough from birth until the study began, the data is left truncated. Construct a table showing the number of subjects at risk, Y, as a function of age.

Right censored approach while ignoring left truncation:

a.

| | • | orderedEventTimes_tj | eventsAtEventTime_ej | censoredObservationsInInterval_cj | inRiskSetAtTime_nj | kaplanMeirSurvivalCurveAtTime_s_tj-1 | ¢ c_tj-1 | kaplanMeirSurvivalCurveAtTime_s_tj |
|--|----|----------------------|----------------------|-----------------------------------|--------------------|--------------------------------------|----------|------------------------------------|
| | 1 | 0 | 0 | 0 | 30 | - | 30/30 | 1 |
| | 2 | 60 | 1 | 0 | 30 | 1 | 29/30 | 0.97 |
| | 3 | 62 | 1 | 0 | 29 | 0.97 | 28/29 | 0.94 |
| | 4 | 63 | 1 | 0 | 28 | 0.94 | 27/28 | 0.91 |
| | 5 | 65 | 2 | 0 | 27 | 0.91 | 25/27 | 0.84 |
| | 6 | 66 | 1 | 0 | 25 | 0.84 | 24/25 | 0.81 |
| | 7 | 68 | 2 | 0 | 24 | 0.81 | 22/24 | 0.74 |
| | 8 | 69 | 2 | 2 | 22 | 0.74 | 20/22 | 0.67 |
| | 9 | 70 | 2 | 0 | 18 | 0.67 | 16/18 | 0.6 |
| | 10 | 71 | 2 | 0 | 16 | 0.6 | 14/16 | 0.52 |
| | 11 | 72 | 2 | 1 | 14 | 0.52 | 12/14 | 0.45 |
| | 12 | 73 | 1 | 1 | 11 | 0.45 | 10/11 | 0.41 |
| | 13 | 74 | 1 | 1 | 9 | 0.41 | 8/9 | 0.36 |
| | 14 | 76 | 1 | 1 | 7 | 0.36 | 6/7 | 0.31 |
| | 15 | 77 | 1 | 0 | 5 | 0.31 | 4/5 | 0.25 |

where inRiskSetAtTIme_nj is the # at risk Y as a function of age tj. For example, at age 63 there are 28 at risk

(b) Estimate the conditional survival function for the age of death of a

diabetic patient who has survived to age 60.

Using above table conditional survival for the age of death for a diabetic patient who has survived to age 60 is 0.97

(c) Estimate the conditional survival function for the age of death of a diabetic patient who has survived to age 70.

Using above table conditional survival for the age of death for a diabetic patient who has survived to age 70 is 0.6

R CODE AND OUTPUT BELOW

```
azt ddc=c("4+",6,11,12,32,35,"38+",39,45,49,75,80,84,85,87,102,"180+")
azt_ddc_saq=c(2,3,4,12,22,48,51,56,80,85,90,94,160,171,180,180,238)
kmTable=data.frame()
getKMTable = function(censoredTimesVector,censorSymbol){
  #get numeric representation of censor vector
  censoredTimesVectorNumeric=as.numeric(sub(censorSymbol, '', censoredTimesVector, fixed=TRUE))
  #count number of actual rows in KM table
  cnt n=length(censoredTimesVectorNumeric)
 #create first row of KM table
  kmTable=setNames(data.frame(matrix(nrow=1,c(0,0,0,cnt_n,as.character("-"),as.character(paste0(cnt_n,"/",c
nt_n)),1)),stringsAsFactors=FALSE),c("orderedEventTimes_tj","eventsAtEventTime_ej",
  "censoredObservationsInInterval cj", "inRiskSetAtTime nj", "kaplanMeirSurvivalCurveAtTime s tj-1", "c tj-1",
"kaplanMeirSurvivalCurveAtTime s tj"))
  censoredTimesVectorNumeric=sort(censoredTimesVectorNumeric)
  for (i in 1:max(censoredTimesVectorNumeric)){
   if(i %in% censoredTimesVectorNumeric){
      #create empty row to fill in
      kmTableRow=setNames(data.frame(matrix(NA,nrow=1,ncol=length(names(kmTable)))),names(kmTable))
      kmTableRow$orderedEventTimes tj=i
      #count how many events at time
      kmTableRow$eventsAtEventTime_ej=length(which(censoredTimesVector==i))
     #count how many censured at time
      kmTableRow$censoredObservationsInInterval cj=length(which(censoredTimesVector==paste0(i,censorSymbol)
))
      kmTableRow$inRiskSetAtTime nj=cnt n
      #sum events and number censored at time
      loss=kmTableRow$eventsAtEventTime ej+kmTableRow$censoredObservationsInInterval cj
      prevSurv=kmTable[dim(kmTable)[1],c("kaplanMeirSurvivalCurveAtTime s tj")]
      kmTableRow[c("kaplanMeirSurvivalCurveAtTime s tj-1")]=prevSurv
      kmTableRow[c("c tj-1")]=paste0((cnt n-loss),"/",cnt n)
      #kmTableRow$kaplanMeirSurvivalCurveAtTime s tj=round((cnt n-loss)/length(censoredTimesVectorNumeric),
3)
```

```
kmTableRow$kaplanMeirSurvivalCurveAtTime_s_tj=round((cnt_n-loss)/cnt_n*as.numeric(prevSurv),3)
      #update count
      cnt_n=cnt_n-loss
      if(kmTableRow$censoredObservationsInInterval_cj==0){
        #add row to kmtable
        kmTable=rbind(kmTable,kmTableRow)
    }
  kmTable
azt ddc KM=getKMTable(azt ddc,"+")
azt_ddc_saq_KM=getKMTable(azt_ddc_saq,"+")
show(azt_ddc_KM)
      orderedEventTimes_tj eventsAtEventTime_ej
##
## 1
## 2
                         6
                                               1
## 3
                        11
                        12
                                               1
## 4
                        32
                                               1
## 5
## 6
                        35
                                               1
## 7
                        39
## 8
                        45
## 9
                        49
                                               1
## 10
                        75
                                               1
## 11
                        80
                                               1
## 12
                        84
## 13
                        85
                        87
## 14
                                               1
## 15
                       102
##
      censoredObservationsInInterval_cj inRiskSetAtTime_nj
## 1
                                                         17
## 2
                                       0
                                                         16
## 3
                                                         15
                                       0
## 4
                                                         14
```

```
## 5
                                                         13
## 6
                                       0
                                                         12
## 7
                                                         10
## 8
                                                          9
## 9
                                                          8
## 10
                                                          7
                                                          6
## 11
                                                          5
## 12
## 13
                                                          4
## 14
                                                          3
                                       0
                                                          2
## 15
      kaplanMeirSurvivalCurveAtTime_s_tj-1 c_tj-1
##
## 1
                                           17/17
## 2
                                          1 15/16
## 3
                                     0.938 14/15
## 4
                                     0.875 13/14
## 5
                                     0.812 12/13
## 6
                                      0.75 11/12
## 7
                                     0.688
                                             9/10
## 8
                                     0.619
                                              8/9
## 9
                                      0.55
                                              7/8
## 10
                                     0.481
                                               6/7
## 11
                                     0.412
                                               5/6
## 12
                                              4/5
                                     0.343
## 13
                                     0.274
                                               3/4
## 14
                                     0.206
                                               2/3
## 15
                                     0.137
                                              1/2
      kaplanMeirSurvivalCurveAtTime_s_tj
##
## 1
## 2
                                   0.938
## 3
                                   0.875
                                   0.812
## 4
                                    0.75
## 5
## 6
                                   0.688
                                   0.619
## 7
## 8
                                    0.55
```

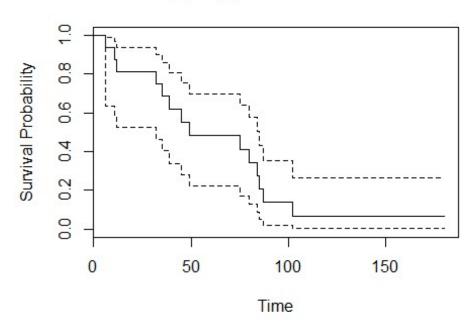
```
## 9
                                    0.481
## 10
                                    0.412
## 11
                                    0.343
## 12
                                    0.274
## 13
                                    0.206
## 14
                                    0.137
## 15
                                    0.068
show(azt_ddc_saq_KM)
      orderedEventTimes_tj eventsAtEventTime_ej
##
## 1
## 2
                          2
                                               1
## 3
                          3
                                               1
## 4
                          4
## 5
                        12
## 6
                        22
                                               1
## 7
                        48
                                               1
## 8
                        51
                                               1
## 9
                        56
                                               1
## 10
                        80
## 11
                        85
## 12
                        90
## 13
                        94
## 14
                       160
## 15
                       171
                                               1
## 16
                       180
                                               2
## 17
                       238
      censoredObservationsInInterval_cj inRiskSetAtTime_nj
##
## 1
                                                          17
## 2
                                       0
                                                          17
## 3
                                                          16
## 4
                                                         15
## 5
                                                          14
## 6
                                                         13
## 7
                                                         12
```

```
## 8
                                                         11
## 9
                                      0
                                                         10
## 10
                                                          9
## 11
                                                          8
## 12
                                                          7
                                                          6
## 13
                                                          5
## 14
## 15
                                                          4
## 16
                                      0
                                                          3
## 17
                                                          1
      kaplanMeirSurvivalCurveAtTime_s_tj-1 c_tj-1
##
## 1
                                         - 17/17
## 2
                                         1 16/17
## 3
                                     0.941 15/16
## 4
                                     0.882 14/15
## 5
                                     0.823 13/14
## 6
                                     0.764 12/13
## 7
                                     0.705 11/12
## 8
                                     0.646 10/11
## 9
                                     0.587
                                             9/10
## 10
                                     0.528
                                              8/9
                                     0.469
                                              7/8
## 11
## 12
                                      0.41
                                               6/7
## 13
                                              5/6
                                     0.351
## 14
                                     0.292
                                              4/5
                                     0.234
## 15
                                              3/4
## 16
                                     0.176
                                              1/3
## 17
                                     0.059
                                               0/1
      kaplanMeirSurvivalCurveAtTime_s_tj
##
## 1
## 2
                                   0.941
## 3
                                   0.882
## 4
                                   0.823
## 5
                                   0.764
## 6
                                   0.705
## 7
                                   0.646
```

```
## 8
                                   0.587
## 9
                                   0.528
## 10
                                   0.469
## 11
                                   0.41
## 12
                                   0.351
                                  0.292
## 13
## 14
                                  0.234
## 15
                                  0.176
## 16
                                   0.059
## 17
                                       0
library(survival)
## Warning: package 'survival' was built under R version 3.5.2
#numeric times and censor list (0 for not censored 1 for censored)
Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grep1("+",azt_ddc,fixed=TRUE),0,1))
                            32 35
                                       38+ 39
                                                45 49
                                                         75
                                                               80
## [1]
                  11 12
                                                                    84
                                                                         85
## [15] 87 102 180+
azt_ddc_KM_R=survfit(Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grep1("+",azt_ddc,fixed=TRUE),0
,1))~1,conf.type="log-log")
summary(azt ddc KM R)
## Call: survfit(formula = Surv(as.numeric(sub("+", "", azt ddc, fixed = TRUE)),
      ifelse(grep1("+", azt_ddc, fixed = TRUE), 0, 1)) ~ 1, conf.type = "log-log")
##
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
       6
             16
                         0.9375 0.0605
                                              0.63235
                                                             0.991
##
             15
                                              0.58598
##
      11
                         0.8750 0.0827
                                                             0.967
                     1 0.8125 0.0976
                                              0.52460
##
      12
             14
                                                             0.935
                     1 0.7500 0.1083
      32
             13
                                             0.46343
                                                             0.898
##
                     1 0.6875 0.1159
      35
                                             0.40460
                                                             0.856
##
             12
      39
             10
                         0.6188 0.1230
                                              0.33929
                                                             0.808
##
                         0.5500 0.1271
      45
##
             9
                                              0.27933
                                                             0.756
##
      49
             8
                          0.4813 0.1285
                                              0.22410
                                                             0.699
```

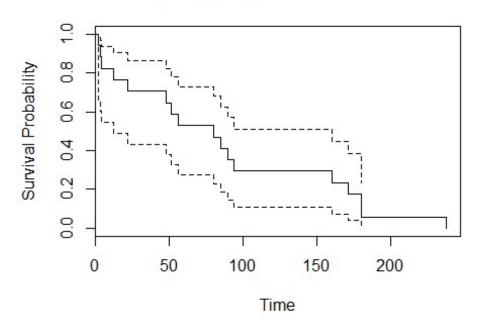
```
##
      75
                          0.4125 0.1272
                                              0.17339
                                                              0.639
                          0.3438 0.1232
##
      80
                                              0.12728
                                                              0.575
##
      84
              5
                      1 0.2750 0.1162
                                              0.08617
                                                              0.507
##
      85
              4
                          0.2063 0.1055
                                              0.05082
                                                              0.433
                          0.1375 0.0900
##
      87
              3
                                              0.02265
                                                              0.354
     102
              2
                          0.0688 0.0662
                                              0.00443
                                                              0.267
##
azt ddc sag KM R = survfit(Surv(as.numeric(sub("+","",azt ddc sag,fixed=TRUE)),ifelse(grepl("+",azt ddc sag
,fixed=TRUE),0,1))~1,conf.type="log-log")
summary(azt ddc saq KM R)
## Call: survfit(formula = Surv(as.numeric(sub("+", "", azt_ddc_saq, fixed = TRUE)),
       ifelse(grep1("+", azt_ddc_saq, fixed = TRUE), 0, 1)) ~ 1,
##
##
       conf.type = "log-log")
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
                          0.9412 0.0571
##
       2
             17
                                              0.65018
                                                              0.991
       3
##
             16
                          0.8824 0.0781
                                              0.60598
                                                              0.969
                          0.8235 0.0925
                                              0.54713
                                                              0.939
##
       4
             15
             14
                          0.7647 0.1029
                                              0.48828
                                                              0.904
##
      12
                          0.7059 0.1105
##
      22
             13
                                              0.43148
                                                              0.866
##
      48
             12
                          0.6471 0.1159
                                              0.37715
                                                              0.823
##
      51
             11
                          0.5882 0.1194
                                              0.32537
                                                              0.778
##
      56
             10
                          0.5294 0.1211
                                              0.27617
                                                              0.730
                          0.4706 0.1211
##
      80
              9
                                              0.22960
                                                              0.680
##
      85
              8
                          0.4118 0.1194
                                              0.18576
                                                              0.626
      90
              7
                          0.3529 0.1159
##
                                              0.14483
                                                              0.570
                      1 0.2941 0.1105
                                              0.10712
      94
              6
                                                              0.511
##
              5
                          0.2353 0.1029
##
    160
                      1
                                              0.07308
                                                              0.449
                          0.1765 0.0925
##
    171
              4
                      1
                                              0.04348
                                                              0.383
                          0.0588 0.0571
                                              0.00391
     180
              3
                                                              0.235
##
##
     238
              1
                      1
                          0.0000
                                     NaN
                                                   NA
                                                                 NA
plot(azt ddc KM R,xlab="Time",ylab="Survival Probability",main="azt ddc KM survival curve")
```

azt_ddc_KM survival curve



plot(azt_ddc_saq_KM_R,xlab="Time",ylab="Survival Probability",main="azt_ddc_saq_KM survival curve")

azt_ddc_saq_KM survival curve



For both groups separately, construct a data layout (similar to what was done in lecture slides) containing the unique, ordered event times, the number of events that occurred at those unique event times, the number of censored observations in the relevant interval, the number in the risk set at that time, and the Kaplan-Meier estimate of the survival curve at that time. What is the median survival time in the two groups? Will you be comfortable reporting the mean survival time in the two groups?

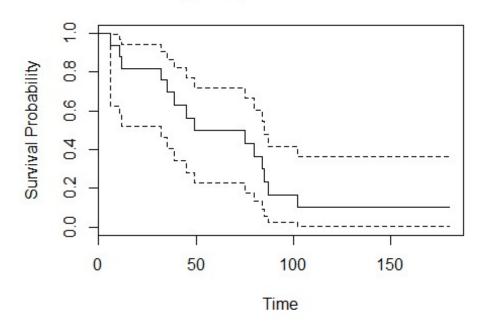
```
#6MP as test case
leukemia_6MP = c(10,7,"32+",23,22,6,16,"34+","32+","25+","11+","20+","19+",6,"17+","35+",6,13,"9+","6+","10
+")
getNATable = function(censoredTimesVector,censorSymbol){
    #get numeric representation of censor vector
```

```
censoredTimesVectorNumeric=as.numeric(sub(censorSymbol, '', censoredTimesVector, fixed=TRUE))
  #count number of actual rows in NA table
  cnt n=length(censoredTimesVectorNumeric)
  #create first row of NA table
  naTable=setNames(data.frame(matrix(nrow=1,c(0,0,cnt_n,0,0,as.character(paste0(cnt_n,"/",cnt_n)),0,0)),str
ingsAsFactors=FALSE),c("orderedEventTimes tj","eventsAtEventTime ej","inRiskSetAtTime nj","censoredObservat
ionsInInterval cj", "cumulativeHazardRate ht", "d Y ratio", "cumulativeHazardEstimatedVariance vt", "nelsonAale
nSurvivalCurveAtTime s tj"))
  censoredTimesVectorNumeric=sort(censoredTimesVectorNumeric)
  sumCensoredInInterval=0
  for (i in 1:max(censoredTimesVectorNumeric)){
   if(i %in% censoredTimesVectorNumeric){
      #create empty row to fill in
      naTableRow=setNames(data.frame(matrix(NA, nrow=1, ncol=length(names(naTable)))), names(naTable))
      naTableRow$orderedEventTimes tj=i
      #count how many events at time
      naTableRow$eventsAtEventTime ej=length(which(censoredTimesVector==i))
     #running total of censured between censured time intervals
      naTableRow$censoredObservationsInInterval cj=length(which(censoredTimesVector==pasteO(i,censorSymbol)
))
      sumCensoredInInterval=sumCensoredInInterval+naTableRow$censoredObservationsInInterval ci
      naTableRow$inRiskSetAtTime nj=cnt n
      naTableRow$d Y ratio=paste0((naTableRow$eventsAtEventTime ej),"/",cnt n)
      naTableRow$cumulativeHazardRate ht=round(as.numeric(naTable[dim(naTable)[1],c("cumulativeHazardRate h
t")])+naTableRow$eventsAtEventTime ej/cnt n,3)
      naTableRow$cumulativeHazardEstimatedVariance_vt=round(as.numeric(naTable[dim(naTable)[1],c("cumulativ
eHazardEstimatedVariance vt")])+naTableRow$eventsAtEventTime ej/(cnt n)^2,3)
      naTableRow$nelsonAalenSurvivalCurveAtTime s tj=round(exp(-naTableRow$cumulativeHazardRate ht),3)
      #sum events and number censored at time
      loss=naTableRow$eventsAtEventTime ej+naTableRow$censoredObservationsInInterval cj
      #update count
      cnt n=cnt n-loss
     #add row to na table if at least one uncensored variable
      if (i %in% censoredTimesVector){
        naTableRow$censoredObservationsInInterval_cj=sumCensoredInInterval
        naTable=rbind(naTable,naTableRow)
```

```
sumCensoredInInterval=0
   }
  naTable
#adapted from http://sas-and-r.blogspot.com/2010/05/example-739-nelson-aalen-estimate-of.html
getCumulativeHazardNA = function(time, event) {
   na.fit = survfit(coxph(Surv(time,event)~1), type="aalen")
   jumps = c(0, na.fit$time, max(time))
   # need to be careful at the beginning and end
   surv = c(1, na.fit$surv, na.fit$surv[length(na.fit$surv)])
   # apply appropriate transformation
   neglogsurv = -log(surv)
   # create placeholder of correct length
   naest = numeric(length(time))
   for (i in 2:length(jumps)) {
      naest[which(time>=jumps[i-1] & time<=jumps[i])] =</pre>
         neglogsurv[i-1] # snaq the appropriate value
   return(sort(unique(naest)))
#TEST SET VALIDATED BY TABLE 4.2 in text page 95
# Leukemia 6MP NA=getNATable(leukemia 6MP,"+")
# leukemia_6MP_NA_R=survfit(Surv(as.numeric(sub("+","",leukemia_6MP,fixed=TRUE)),ifelse(grepl("+",leukemia_
6MP, fixed=TRUE), 0, 1))~1, conf.type="log-log", type="fh")
# summary(Leukemia 6MP NA R)
# plot(leukemia 6MP NA R,xlab="Time",ylab="Survival Probability",main="leukemia 6MP NA R survival curve")
# Leukemia_6MP_NA_CH_R=getCumulativeHazardNA(as.numeric(sub("+","",leukemia_6MP,fixed=TRUE)),ifelse(grepl("
+", Leukemia 6MP, fixed=TRUE), 0, 1))
azt ddc NA=getNATable(azt ddc,"+")
azt_ddc_NA_R=survfit(Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grep1("+",azt_ddc,fixed=TRUE),0
```

```
,1))~1,conf.type="log-log",type="fh")
summary(azt ddc NA R)
## Call: survfit(formula = Surv(as.numeric(sub("+", "", azt_ddc, fixed = TRUE)),
       ifelse(grep1("+", azt_ddc, fixed = TRUE), 0, 1)) ~ 1, conf.type = "log-log",
##
       type = "fh")
##
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
                      1
                           0.939 0.0606
                                              0.62304
                                                              0.992
##
       6
             16
##
      11
             15
                           0.879 0.0830
                                              0.58170
                                                              0.970
                           0.818 0.0983
                                              0.52281
##
      12
             14
                      1
                                                              0.940
                           0.758 0.1094
##
      32
             13
                      1
                                              0.46341
                                                              0.905
      35
                           0.697 0.1175
##
             12
                                              0.40600
                                                              0.865
                           0.631 0.1254
                                                              0.820
##
      39
             10
                      1
                                              0.34186
                           0.564 0.1304
                                              0.28289
##
      45
              9
                      1
                                                              0.772
                           0.498 0.1330
                                              0.22843
                                                              0.720
##
      49
              8
                      1
      75
              7
                           0.432 0.1331
                                              0.17823
                                                              0.664
##
                      1
      80
              6
                           0.366 0.1310
                                              0.13234
##
                      1
                                                              0.606
##
      84
              5
                      1
                           0.299 0.1264
                                              0.09106
                                                             0.545
      85
              4
##
                           0.233 0.1192
                                              0.05506
                                                              0.481
                      1
              3
                      1
##
      87
                           0.167 0.1093
                                              0.02560
                                                              0.417
                           0.101 0.0976
              2
                      1
##
     102
                                               0.00539
                                                              0.367
plot(azt_ddc_NA_R,xlab="Time",ylab="Survival Probability",main="azt_ddc_NA survival curve")
```

azt_ddc_NA survival curve



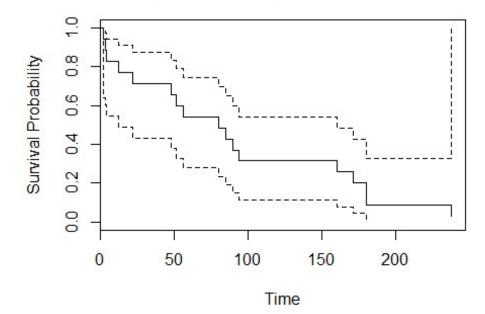
```
#cumulative hazard, confirmation of results
azt_ddc_NA_CH_R=getCumulativeHazardNA(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grep1("+",azt_ddc,f
ixed=TRUE),0,1))
show(azt_ddc_NA)
      orderedEventTimes_tj eventsAtEventTime_ej inRiskSetAtTime_nj
##
## 1
                                                                 17
## 2
                         6
                                               1
                                                                 16
## 3
                                                                 15
                        11
                                               1
## 4
                                                                 14
                        12
## 5
                        32
                                                                 13
                                               1
## 6
                        35
                                                                 12
                                               1
## 7
                        39
                                                                 10
```

```
## 8
                        45
                                                                   9
## 9
                        49
                                                                   8
                                               1
                        75
## 10
                                               1
## 11
                        80
                                               1
## 12
                        84
                                               1
## 13
                        85
## 14
                        87
## 15
                       102
      censoredObservationsInInterval_cj cumulativeHazardRate_ht d_Y_ratio
##
## 1
                                                                0
                                                                      17/17
## 2
                                                            0.062
                                                                       1/16
## 3
                                                            0.129
                                                                       1/15
## 4
                                                              0.2
                                                                       1/14
## 5
                                                            0.277
                                                                       1/13
## 6
                                                             0.36
                                                                       1/12
## 7
                                                             0.46
                                                                       1/10
## 8
                                                                        1/9
                                                            0.571
## 9
                                                            0.696
                                                                        1/8
## 10
                                                            0.839
                                                                        1/7
## 11
                                                            1.006
                                                                        1/6
## 12
                                                            1.206
                                                                        1/5
## 13
                                                            1.456
                                                                        1/4
## 14
                                       0
                                                            1.789
                                                                        1/3
## 15
                                                            2.289
                                                                        1/2
      cumulativeHazardEstimatedVariance vt
##
## 1
## 2
                                      0.004
## 3
                                      0.008
## 4
                                      0.013
                                      0.019
## 5
## 6
                                      0.026
                                      0.036
## 7
## 8
                                      0.048
## 9
                                      0.064
## 10
                                      0.084
## 11
                                      0.112
```

```
## 12
                                     0.152
                                     0.214
## 13
## 14
                                     0.325
## 15
                                     0.575
      nelsonAalenSurvivalCurveAtTime_s_tj
##
## 1
## 2
                                     0.94
## 3
                                    0.879
## 4
                                    0.819
## 5
                                    0.758
## 6
                                    0.698
                                    0.631
## 7
## 8
                                    0.565
                                    0.499
## 9
## 10
                                    0.432
## 11
                                    0.366
## 12
                                    0.299
## 13
                                    0.233
## 14
                                    0.167
## 15
                                    0.101
azt ddc sag NA=getNATable(azt ddc sag,"+")
azt_ddc_saq_NA_R = survfit(Surv(as.numeric(sub("+","",azt_ddc_saq,fixed=TRUE)),ifelse(grepl("+",azt_ddc_saq
,fixed=TRUE),0,1))~1,conf.type="log-log",type="fh")
summary(azt_ddc_saq_NA_R)
## Call: survfit(formula = Surv(as.numeric(sub("+", "", azt ddc saq, fixed = TRUE)),
       ifelse(grepl("+", azt ddc saq, fixed = TRUE), 0, 1)) ~ 1,
##
       conf.type = "log-log", type = "fh")
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
##
       2
             17
                      1
                          0.9429 0.0572
                                               0.6417
                                                              0.992
       3
##
             16
                          0.8857 0.0784
                                               0.6021
                                                              0.971
                          0.8286 0.0930
       4
##
             15
                                               0.5455
                                                              0.943
                          0.7715 0.1038
##
      12
             14
                                               0.4883
                                                              0.910
##
      22
             13
                          0.7144 0.1118
                                               0.4328
                                                              0.874
```

```
0.834
      48
             12
                          0.6573 0.1177
                                                0.3795
##
      51
             11
                          0.6001 0.1218
                                                0.3287
                                                               0.791
##
      56
             10
                          0.5430 0.1242
                                                0.2802
                                                               0.746
##
##
      80
              9
                          0.4859 0.1250
                                                0.2343
                                                               0.698
                                                0.1908
                                                               0.649
##
      85
                          0.4288 0.1243
##
      90
                          0.3717 0.1221
                                                0.1501
                                                              0.597
                          0.3147 0.1182
      94
              6
                                                0.1124
                                                               0.543
##
                          0.2576 0.1126
                                                0.0780
                                                              0.486
##
     160
              5
     171
##
                          0.2006
                                  0.1051
                                                0.0476
                                                               0.428
##
     180
                          0.0872 0.0846
                                                0.0049
                                                               0.327
     238
              1
                          0.0321
                                      Inf
                                                   NaN
                                                              1.000
##
plot(azt_ddc_saq_NA_R,xlab="Time",ylab="Survival Probability",main="azt_ddc_saq_NA survival curve")
```

azt_ddc_saq_NA survival curve

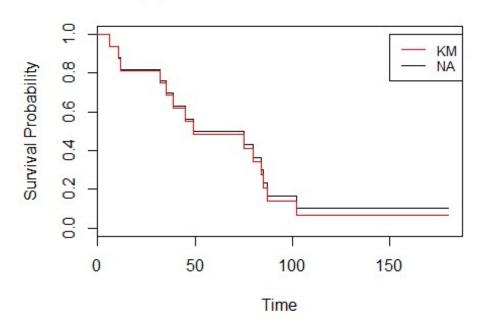


```
#cumulative hazard, confirmation of results
azt_ddc_saq_NA_CH_R=getCumulativeHazardNA(as.numeric(sub("+","",azt_ddc_saq,fixed=TRUE)),ifelse(grep1("+",a
zt_ddc_saq,fixed=TRUE),0,1))
show(azt_ddc_saq_NA)
      orderedEventTimes tj eventsAtEventTime ej inRiskSetAtTime nj
##
## 1
                                                                   17
## 2
                          2
                                                1
                                                                   17
## 3
                                                1
                                                                   16
## 4
                          4
                                                1
                                                                   15
## 5
                                                                   14
                                                1
                         12
## 6
                                                                   13
                         22
                                                1
                         48
                                                                   12
## 7
                                                1
## 8
                         51
                                                1
                                                                   11
## 9
                         56
                                                1
                                                                   10
                                                                    9
## 10
                         80
                                                1
## 11
                         85
                                                1
                                                                    8
## 12
                         90
                                                1
                                                                    7
## 13
                         94
                                                1
                                                                    6
## 14
                        160
                                                1
                                                                    5
## 15
                                                1
                        171
                                                2
## 16
                        180
## 17
                        238
      censoredObservationsInInterval_cj cumulativeHazardRate_ht d_Y_ratio
##
## 1
                                                                       17/17
                                                            0.059
                                                                        1/17
## 2
                                       0
                                       0
                                                                        1/16
## 3
                                                             0.122
## 4
                                                             0.189
                                                                        1/15
                                                             0.26
## 5
                                                                        1/14
## 6
                                       0
                                                            0.337
                                                                        1/13
                                       0
                                                             0.42
                                                                        1/12
## 7
## 8
                                       0
                                                            0.511
                                                                        1/11
## 9
                                                            0.611
                                                                        1/10
                                                                         1/9
                                       0
                                                             0.722
## 10
## 11
                                                            0.847
                                                                         1/8
                                       0
## 12
                                                             0.99
                                                                         1/7
```

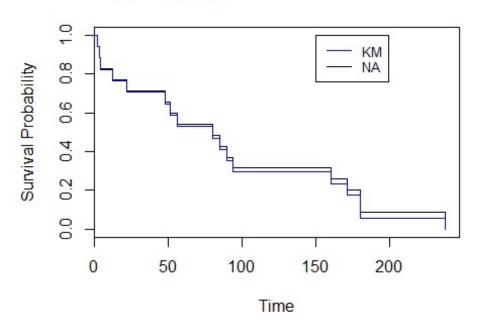
```
## 13
                                                                       1/6
                                                           1.157
## 14
                                       0
                                                           1.357
                                                                       1/5
## 15
                                       0
                                                           1.607
                                                                       1/4
## 16
                                       0
                                                           2.274
                                                                       2/3
## 17
                                                           3.274
                                                                       1/1
##
      cumulativeHazardEstimatedVariance vt
## 1
## 2
                                      0.003
## 3
                                      0.007
## 4
                                      0.011
## 5
                                      0.016
## 6
                                      0.022
## 7
                                      0.029
## 8
                                      0.037
## 9
                                      0.047
## 10
                                      0.059
## 11
                                      0.075
## 12
                                      0.095
## 13
                                      0.123
## 14
                                      0.163
## 15
                                      0.226
## 16
                                      0.448
## 17
                                      1.448
      nelsonAalenSurvivalCurveAtTime_s_tj
##
## 1
## 2
                                    0.943
## 3
                                    0.885
## 4
                                    0.828
## 5
                                    0.771
## 6
                                    0.714
## 7
                                    0.657
## 8
                                      0.6
## 9
                                    0.543
## 10
                                    0.486
                                    0.429
## 11
## 12
                                     0.372
```

```
## 13
                                    0.314
                                    0.257
## 14
                                      0.2
## 15
## 16
                                    0.103
## 17
                                    0.038
#compare NA to KM survival curve graphically
#azt ddc
plot(survfit(Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grepl("+",azt_ddc,fixed=TRUE),0,1))~1,c
onf.type="none", type="fh"), xlab="Time", ylab="Survival Probability", main="azt_ddc NA vs. KM survival curves"
lines(survfit(Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grep1("+",azt_ddc,fixed=TRUE),0,1))~1,
conf.type="none",type="kaplan-meier"),col="red")
legend(150, 1, legend=c("KM", "NA"),
       col=c("red", "black"), lty=1,cex=0.8)
```

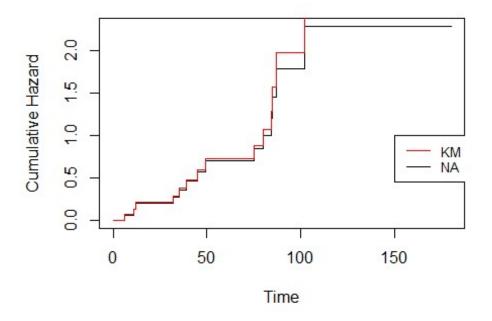
azt_ddc NA vs. KM survival curves



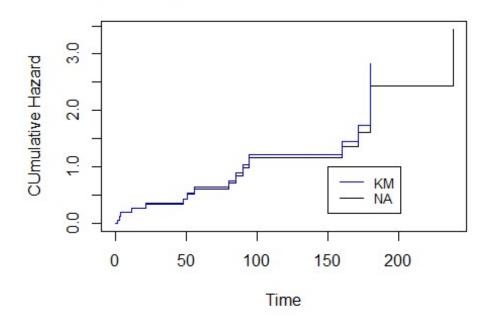
azt_ddc_saq NA vs. KM survival curves



azt_ddc NA vs. KM cumulative hazard curves



azt_ddc_saq NA vs. KM cumulative hazard curve



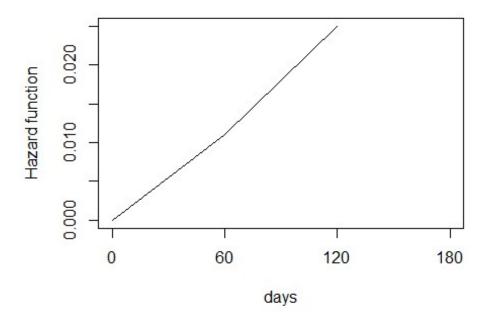
```
----- tidyverse conflicts() --
## x dplyr::filter() masks stats::filter()
## x dplyr::lag()
                    masks stats::lag()
getLifeTableInput = function(censoredTimesVector,censorSymbol){
  #get numeric representation of censor vector
  censoredTimesVectorNumeric=as.numeric(sub(censorSymbol, '', censoredTimesVector, fixed=TRUE))
 #count number of actual rows in KM table
  cnt n=length(censoredTimesVectorNumeric)
 #create first row of KM table
 lifeTableInputTable=setNames(data.frame(matrix(nrow=1,c(NA,NA,NA)),stringsAsFactors=FALSE),c("time","nlos
t", "nevent"))
  censoredTimesVectorNumeric=sort(censoredTimesVectorNumeric)
 for (i in 1:max(censoredTimesVectorNumeric)){
   if(i %in% censoredTimesVectorNumeric){
      #create empty row to fill in
     lifeTableInputRow=setNames(data.frame(matrix(NA,nrow=1,ncol=length(names(lifeTableInputTable)))),name
s(lifeTableInputTable))
      lifeTableInputRow$time=i
     #count how many events at time
     lifeTableInputRow$nevent=length(which(censoredTimesVector==i))
     #count how many censured at time
     lifeTableInputRow$nlost=length(which(censoredTimesVector==paste0(i,censorSymbol)))
     lifeTableInputTable=rbind(lifeTableInputTable,lifeTableInputRow)
   }
  na.omit(lifeTableInputTable)
azt_ddc_numeric=as.numeric(gsub("+","",azt_ddc,fixed=TRUE))
cuts ad=seq(0,max(azt ddc numeric),60)
#ensure no loss of upper bound when incrementing
if(max(azt ddc numeric)>cuts ad[length(cuts ad)]){cuts ad=cuts ad=c(cuts ad,(max(azt ddc numeric)))}
azt ddc lt raw=getLifeTableInput(azt ddc,"+")
lifetab dat=mutate(azt ddc lt raw,time cat = cut(time, cuts ad)) %>% group by(time cat) %>% summarize(ilost
=sum(nlost),ievent=sum(nevent))
```

```
azt ddc lt=lifetab(tis = c(0,azt ddc lt raw$time), ninit = length(azt ddc), nlost = azt ddc lt raw$nlost, n
event = azt ddc lt raw$nevent) %>% drop na(hazard)
show(azt_ddc_lt)
##
         nsubs nlost nrisk nevent
                                                  pdf
                                                          hazard
                                                                   se.surv
                                     surv
## 0-4
                   1 16.5
                                17
                   0 16.0
## 4-6
            16
                                1 1.00000 0.031250000 0.03225806 0.00000000
                   0 15.0
                                1 0.93750 0.012500000 0.01379310 0.06051536
## 6-11
            15
                   0 14.0
                                1 0.87500 0.062500000 0.07407407 0.08267973
## 11-12
            14
## 12-32
                   0 13.0
                                1 0.81250 0.003125000 0.00400000 0.09757809
            13
                   0 12.0
## 32-35
            12
                                1 0.75000 0.020833333 0.02898551 0.10825318
                   1 10.5
## 35-38
            11
                                0 0.68750 0.000000000 0.00000000 0.11587810
                   0 10.0
## 38-39
            10
                                1 0.68750 0.068750000 0.10526316 0.11587810
                      9.0
## 39-45
                                1 0.61875 0.011458333 0.01960784 0.12300557
## 45-49
                      8.0
                                1 0.55000 0.017187500 0.03333333 0.12710724
## 49-75
                      7.0
                                1 0.48125 0.002644231 0.00591716 0.12847323
## 75-80
                       6.0
                                1 0.41250 0.013750000 0.03636364 0.12719172
                       5.0
## 80-84
                                1 0.34375 0.017187500 0.05555556 0.12318011
## 84-85
                       4.0
                                1 0.27500 0.068750000 0.28571429 0.11615588
## 85-87
                       3.0
                                1 0.20625 0.034375000 0.20000000 0.10551909
## 87-102
                       2.0
                                1 0.13750 0.004583333 0.04444444 0.08999783
                       se.hazard
##
              se.pdf
## 0-4
                 NaN
                             NaN
## 4-6
         0.030257682 0.032241277
## 6-11
         0.012103073 0.013784901
## 11-12 0.060515365 0.074023251
## 12-32 0.003025768 0.003996799
## 32-35 0.020171788 0.028958098
## 35-38
                 NaN
## 38-39 0.066243366 0.105117263
## 39-45 0.011040561 0.019573890
## 45-49 0.016560842 0.033259177
## 49-75 0.002547822 0.005899627
## 75-80 0.013248673 0.036213062
## 80-84 0.016560842 0.055211555
## 84-85 0.066243366 0.282783805
```

```
## 85-87 0.033121683 0.195959179
## 87-102 0.004416224 0.041902624
azt_ddc_lt_60=lifetab(tis = cuts_ad, ninit = length(azt_ddc), nlost = lifetab_dat$ilost, nevent = lifetab_d
at$ievent)
azt_ddc_saq_numeric=as.numeric(gsub("+","",azt_ddc_saq,fixed=TRUE))
cuts ads=seq(0,max(azt ddc sag numeric),by=60)
#ensure no loss of upper bound when incrementing
if(max(azt ddc sag numeric)>cuts ads[length(cuts ads)]){cuts ads=c(cuts ads,(max(azt ddc sag numeric)))}
azt ddc sag lt raw=getLifeTableInput(azt ddc sag,"+")
lifetab dat=mutate(azt ddc sag lt raw,time cat = cut(time, cuts ads)) %>% group by(time cat) %>% summarize(
ilost=sum(nlost),ievent=sum(nevent))
azt ddc sag lt=lifetab(tis = c(0,azt ddc sag lt raw$time), ninit = length(azt ddc sag), nlost = azt ddc sag
lt raw$nlost, nevent = azt ddc sag lt raw$nevent) %>% drop na(hazard)
show(azt ddc sag lt)
           nsubs nlost nrisk nevent
##
                                                       pdf
                                                                hazard
                                         surv
## 0-2
                                  1 1.0000000 0.0294117647 0.030303030
              17
                          17
## 2-3
              16
                     0
                          16
                                  1 0.9411765 0.0588235294 0.064516129
## 3-4
              15
                     0
                          15
                                  1 0.8823529 0.0588235294 0.068965517
## 4-12
              14
                          14
                                  1 0.8235294 0.0073529412 0.009259259
## 12-22
              13
                          13
                                  1 0.7647059 0.0058823529 0.008000000
## 22-48
              12
                          12
                                  1 0.7058824 0.0022624434 0.003344482
## 48-51
              11
                     0
                          11
                                  1 0.6470588 0.0196078431 0.031746032
## 51-56
              10
                     0
                          10
                                  1 0.5882353 0.0117647059 0.021052632
## 56-80
               9
                     0
                           9
                                  1 0.5294118 0.0024509804 0.004901961
## 80-85
               8
                     0
                           8
                                  1 0.4705882 0.0117647059 0.026666667
## 85-90
               7
                     0
                           7
                                  1 0.4117647 0.0117647059 0.030769231
                           6
## 90-94
               6
                     0
                                  1 0.3529412 0.0147058824 0.045454545
               5
                     0
                                  1 0.2941176 0.0008912656 0.003367003
## 94-160
                           5
               4
                     0
## 160-171
                                  1 0.2352941 0.0053475936 0.025974026
## 171-180
                                  2 0.1764706 0.0130718954 0.111111111
                                     se.hazard
##
              se.surv
                            se.pdf
           0.00000000 0.0285336029 0.030289114
## 0-2
## 2-3
           0.05706721 0.0570672059 0.064482553
```

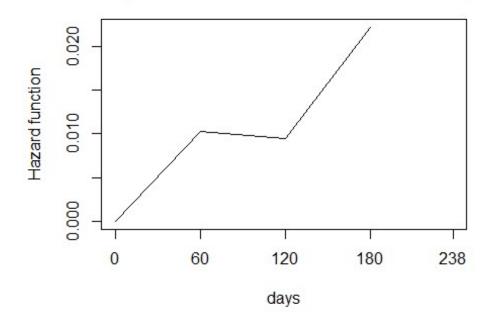
```
## 3-4
          0.07814249 0.0570672059 0.068924503
## 4-12
          0.09245944 0.0071334007 0.009252906
## 12-22
         0.10287937 0.0057067206 0.007993597
## 22-48
          0.11051017 0.0021948925 0.003341319
## 48-51
          0.11590404 0.0190224020 0.031710018
## 51-56 0.11936462 0.0114134412 0.021023453
## 56-80
         0.12105782 0.0023778002 0.004893473
## 80-85 0.12105782 0.0114134412 0.026607341
## 85-90 0.11936462 0.0114134412 0.030678062
## 90-94 0.11590404 0.0142668015 0.045266327
## 94-160 0.11051017 0.0008646546 0.003346155
## 160-171 0.10287937 0.0051879278 0.025707619
## 171-180 0.09245944 0.0086824989 0.068041382
azt_ddc_saq_lt_60=lifetab(tis = cuts_ads, ninit = length(azt_ddc_saq), nlost = lifetab_dat$ilost, nevent =
lifetab_dat$ievent)
#plot azt_ddc
plot(cuts ad,c(0,azt ddc lt 60$hazard),type='l',ylab="Hazard function",xlab="days",xaxt="n",main="azt ddc L
ife Table Hazard Function in 60 day windows")
axis(1, at = cuts ad, las=1)
```

azt_ddc Life Table Hazard Function in 60 day windc



```
#plot azt_ddc_saq
plot(cuts_ads,c(0,azt_ddc_saq_lt_60$hazard),type='l',ylab="Hazard function",xlab="days",xaxt="n",main="azt_
ddc_saq Life Table Hazard Function in 60 day windows")
axis(1, at = cuts_ads, las=1)
```

zt_ddc_saq Life Table Hazard Function in 60 day win

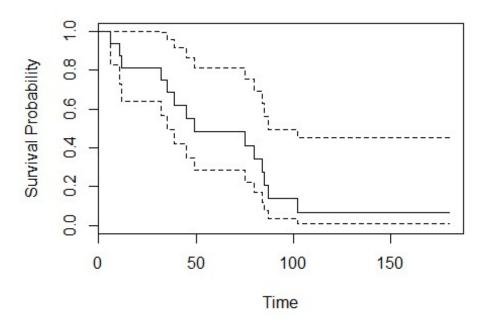


```
library(kmconfband)
## Warning: package 'kmconfband' was built under R version 3.5.2
#survival function confidence intervals for azt_ddc
azt_ddc_s=survfit(Surv(as.numeric(sub("+","",azt_ddc,fixed=TRUE)),ifelse(grepl("+",azt_ddc,fixed=TRUE),0,1)
>~1)
azt_ddc_s_ci=summary(azt_ddc_s)
azt_ddc_s_ci_df=data.frame(azt_ddc_s_ci$time,azt_ddc_s_ci$n.risk,azt_ddc_s_ci$n.event,azt_ddc_s_ci$surv,azt_ddc_s_ci$std.err,azt_ddc_s_ci$lower,azt_ddc_s_ci$upper)
#Lower and upper intervals and bounds
azt_ddc_s_ci_cb=setNames(cbind(azt_ddc_s_ci_df,confband(azt_ddc_s)[1:dim(azt_ddc_s_ci_df)[1],]),c("time","n.risk","n.event","survival","std.err","lower 95% CI","upper 95% CI","lower 95% CB","upper 95% CB"))
```

```
## The critical value required is 0.4404776

plot(azt_ddc_s,xlab="Time",ylab="Survival Probability",main="azt_ddc KM survival curves")
```

azt_ddc KM survival curves



```
rtr_example=c(3,4,"5+",6,"6+","8+",11,14,15,"16+")
decrement=0
#redistribute to right value
getRedistributeToRightTable = function(censoredTimesVector,censorSymbol){
    #get numeric representation of censor vector
    censoredTimesVectorNumeric=as.numeric(sub(censorSymbol,'',censoredTimesVector,fixed=TRUE))
    #sort to ensure when determine step# omit correct last element
    censoredTimesVector=censoredTimesVector[order(censoredTimesVectorNumeric)]
    #count number of actual rows in RTR table
```

```
cnt n=length(censoredTimesVectorNumeric)
  steps=length(which(grep1(censorSymbol,censoredTimesVector[1:length(censoredTimesVector)-1],fixed=TRUE)))
  #create first row of RTR table
  rtrTable=setNames(data.frame(matrix(nrow=1,c(0,(1/cnt_n),"",1)),stringsAsFactors=FALSE),c("observation","
decrement", "action", "S_t"))
  censoredTimesVectorNumeric=sort(censoredTimesVectorNumeric)
  decrement = 0
  base=1/cnt n
  i=0
  uniqueCensoredTimesVector=unique(censoredTimesVector)
  for (dataPoint in uniqueCensoredTimesVector){
    i=i+1
    #create empty row to fill in
   rtrTableRow=setNames(data.frame(matrix(NA, nrow=1, ncol=length(names(rtrTable)))), names(rtrTable))
    rtrTableRow$observation=dataPoint
    #count how many events or censures at unique dataPoint
   numerator=length(which(censoredTimesVector==dataPoint))
    #censured at time?
   if(length(which(censoredTimesVector==dataPoint & grep(censorSymbol,dataPoint,fixed=TRUE)))>0){
      #no change in survival function
      rtrTableRow$S t=rtrTable[dim(rtrTable)[1],c("S t")]
      #display decrement of 0
      rtrTableRow$decrement=0
      #update denominator
      denominator=length(censoredTimesVector)-max(which(dataPoint == censoredTimesVector))
      if(i==length(uniqueCensoredTimesVector)){
        rtrTableRow$action=paste0("survival is 1-(1)")
        rtrTableRow$S t=0
        rtrTableRow$decrement=1
      else{
        rtrTableRow$action=paste0("next time survival is 1-(",decrement," + ",base," + ",base,"*",numerator
,"/",denominator,")")
      if(length(uniqueCensoredTimesVector)>=(i+1) && (!grepl(censorSymbol,uniqueCensoredTimesVector[i+1],fi
xed=TRUE))){
```

```
#update decrement only if not proceeded by a censored observation
        decrement = decrement + base+(base*numerator/denominator)
      #update base
      base=base+(base*numerator/denominator)
   else{
        #update decrement and show action
        if (as.numeric(rtrTable[dim(rtrTable)[1],c("decrement")])==0)
          rtrTableRow$action=paste0("survival is 1-(",decrement,")")
          decrement = decrement
        else{
          decrement = decrement + base
          rtrTableRow$action=paste0("survival is 1-(",decrement,")")
        rtrTableRow$S t=1-decrement
       #update decrement display
        rtrTableRow$decrement=decrement
      #add row to rtrtable
      rtrTable=rbind(rtrTable,rtrTableRow)
  #eleminate initial bogus row
  rtrTable$decrement=round(as.numeric(rtrTable$decrement),3)
  rtrTable$S_t=round(as.numeric(rtrTable$S_t),3)
  rtrTable[-1,]
azt_ddc_saq_RTR=getRedistributeToRightTable(azt_ddc_saq,"+")
azt_ddc_RTR=getRedistributeToRightTable(azt_ddc,"+")
show(azt_ddc_saq_RTR)
##
      observation decrement
                                                        action
                                                               St
                      0.059 survival is 1-(0.0588235294117647) 0.941
## 2
```

```
## 3
                      0.118 survival is 1-(0.117647058823529) 0.882
## 4
                4
                      0.176 survival is 1-(0.176470588235294) 0.824
                      0.235 survival is 1-(0.235294117647059) 0.765
## 5
               12
                      0.294 survival is 1-(0.294117647058824) 0.706
## 6
               22
## 7
                      0.353 survival is 1-(0.352941176470588) 0.647
               48
## 8
               51
                      0.412 survival is 1-(0.411764705882353) 0.588
## 9
               56
                      0.471 survival is 1-(0.470588235294118) 0.529
## 10
               80
                      0.529 survival is 1-(0.529411764705882) 0.471
## 11
               85
                      0.588 survival is 1-(0.588235294117647) 0.412
## 12
                      0.647 survival is 1-(0.647058823529412) 0.353
               90
                      0.706 survival is 1-(0.705882352941177) 0.294
## 13
               94
## 14
                      0.765 survival is 1-(0.764705882352941) 0.235
              160
## 15
              171
                      0.824 survival is 1-(0.823529411764706) 0.176
## 16
                      0.882 survival is 1-(0.882352941176471) 0.118
              180
## 17
              238
                      0.941 survival is 1-(0.941176470588235) 0.059
show(azt ddc RTR)
      observation decrement
##
## 2
                      0.000
               4+
## 3
                6
                      0.062
## 4
               11
                      0.125
## 5
               12
                      0.188
## 6
               32
                      0.250
                      0.312
## 7
               35
## 8
              38+
                      0.000
## 9
               39
                      0.381
               45
                      0.450
## 10
## 11
                      0.519
               49
## 12
               75
                      0.588
                      0.656
## 13
               80
## 14
               84
                      0.725
                      0.794
## 15
               85
## 16
                      0.862
               87
## 17
              102
                      0.931
## 18
             180+
                      1.000
```

```
##
                                                                           action
## 2
      next time survival is 1-(0 + 0.0588235294117647 + 0.0588235294117647*1/16)
## 3
                                                          survival is 1-(0.0625)
## 4
                                                           survival is 1-(0.125)
## 5
                                                          survival is 1-(0.1875)
## 6
                                                            survival is 1-(0.25)
                                                          survival is 1-(0.3125)
## 7
                         next time survival is 1-(0.3125 + 0.0625 + 0.0625*1/10)
## 8
                                                         survival is 1-(0.38125)
## 9
                                                            survival is 1-(0.45)
## 10
## 11
                                                         survival is 1-(0.51875)
## 12
                                                          survival is 1-(0.5875)
                                                         survival is 1-(0.65625)
## 13
                                                           survival is 1-(0.725)
## 14
## 15
                                                         survival is 1-(0.79375)
## 16
                                                          survival is 1-(0.8625)
## 17
                                                         survival is 1-(0.93125)
## 18
                                                               survival is 1-(1)
        S t
##
## 2 1.000
## 3 0.938
## 4 0.875
## 5 0.812
## 6 0.750
## 7 0.688
## 8 0.688
## 9 0.619
## 10 0.550
## 11 0.481
## 12 0.412
## 13 0.344
## 14 0.275
## 15 0.206
## 16 0.138
## 17 0.069
## 18 0.000
```

```
rtr example RTR = getRedistributeToRightTable(rtr example,"+")
#column1 of book page 137 problem 4.7
entry_c1 = c(58,58,59,60,60,61,61,62,62,62,63,63,64,66,66)
exit_c1=c(60,63,69,62,65,72,69,73,66,65,68,74,71,68,69)
death c1 = c(1,1,0,1,1,0,0,0,1,1,1,0,1,1,1)
#column1 of book page 137 problem 4.7
entry c2=c(67,67,67,68,69,69,69,70,70,70,71,72,72,73,73)
exit_c2=c(70,77,69,72,79,72,70,76,71,78,79,76,73,80,74)
death_c2=c(1,1,1,1,0,1,1,0,1,0,0,1,1,0,1)
df2 lec = data.frame(cbind(entry c1,exit c1,death c1))
df2 prob = setNames(data.frame(cbind(c(entry c1,entry c2),c(exit c1,exit c2),c(death c1,death c2))),c("entr
y","exit","death"))
df2 lec censored noLT=c(60,63,"69+",62,65,"72+","69+","73+",66,65,68,"74+",71,68,69)
df2_prob_censored_noLT=c(60,63,"69+",62,65,"72+","69+","73+",66,65,68,"74+",71,68,69,70,77,69,72,"79+",72,7
0, "76+", 71, "78+", "79+", 76, 73, "80+", 74)
getKM LT Table = function(entryExitDeathVector,deathSymbol){
  #create first row of KM LT table
  km ltTable=setNames(data.frame(matrix(nrow=1,c(0,0,0,length(entryExitDeathVector),"",1)),stringsAsFactors
=FALSE),c("tj","ej","cj","nj","c tj-1","s tj"))
  #sort by exit
  orderedIndices=order(entryExitDeathVector$exit)
  entryExitDeathVector=entryExitDeathVector[orderedIndices,]
  for (time in unique(entryExitDeathVector$exit)){
   total at risk=length(which(entryExitDeathVector$entry<=time))</pre>
    gone=length(which(entryExitDeathVector$exit<time & entryExitDeathVector$death==deathSymbol))</pre>
    events=length(which(entryExitDeathVector$exit==time & entryExitDeathVector$death==deathSymbol))
   truncated_at_risk=total_at_risk-gone-as.numeric(km_ltTable$cj[length(km_ltTable$cj)])
   censored=length(which(entryExitDeathVector$exit==time & entryExitDeathVector$death!=deathSymbol))
   numerator=truncated at risk-events
   denominator=truncated at risk
   #print(paste0(time,": ",numerator,"/",denominator))
   #create empty row to fill in
   km ltTableRow=setNames(data.frame(matrix(NA,nrow=1,ncol=length(names(km ltTable)))),names(km ltTable))
```

```
km ltTableRow$tj=time
   #count how many events at time
   km ltTableRow$ej=events
   #count how many censured at time
   km ltTableRow$cj=censored
   km ltTableRow$nj=truncated at risk
   #sum events and number censored at time
   km_ltTableRow[c("c_tj-1")]=paste0(numerator,"/",denominator)
   km ltTableRow$s tj=round((numerator/denominator)*as.numeric(km_ltTable[dim(km_ltTable)[1],c("s_tj")]),4
   if (km ltTableRow$ej>0 | km ltTableRow$s tj==1){
     #add row to km lttable
     km ltTable=rbind(km ltTable,km ltTableRow)
  km_ltTable
df2 lec LT=getKM LT Table(df2 lec,1)
show(df2 lec LT)
## tj ej cj nj c_tj-1 s_tj
## 1 0 0 0 3
                            1
## 2 60 1 0 5
                   4/5
                          0.8
## 3 62 1 0 9 8/9 0.7111
## 4 63 1 0 10 9/10 0.64
## 5 65 2 0 10 8/10 0.512
## 6 66 1 0 10 9/10 0.4608
## 7 68 2 0 9
                 7/9 0.3584
## 8 69 1 2 7
                   6/7 0.3072
## 9 71 1 0 4
                   3/4 0.2304
getKMTableNoCensorRemoval = function(censoredTimesVector,censorSymbol){
  #get numeric representation of censor vector
  censoredTimesVectorNumeric=as.numeric(sub(censorSymbol,'',censoredTimesVector,fixed=TRUE))
 #count number of actual rows in KM table
  cnt n=length(censoredTimesVectorNumeric)
```

```
#create first row of KM table
  kmTable=setNames(data.frame(matrix(nrow=1,c(0,0,0,cnt_n,as.character("-"),as.character(paste0(cnt_n,"/",c
nt_n)),1)),stringsAsFactors=FALSE),c("orderedEventTimes_tj","eventsAtEventTime_ej",
  "censoredObservationsInInterval_cj", "inRiskSetAtTime_nj", "kaplanMeirSurvivalCurveAtTime s tj-1", "c tj-1",
"kaplanMeirSurvivalCurveAtTime s tj"))
  # orderedIndices=order(censoredTimesVectorNumeric)
 # censoredTimesVectorNumeric=censoredTimesVectorNumeric[orderedIndices]
  # censoredTimesVector=censoredTimesVector[orderedIndices]
  censoredTimesVectorNumeric=sort(censoredTimesVectorNumeric)
  for (i in 1:max(censoredTimesVectorNumeric)){
    if(i %in% censoredTimesVectorNumeric){
      #create empty row to fill in
      kmTableRow=setNames(data.frame(matrix(NA,nrow=1,ncol=length(names(kmTable)))),names(kmTable))
      kmTableRow$orderedEventTimes tj=i
      #count how many events at time
      kmTableRow$eventsAtEventTime ej=length(which(censoredTimesVector==i))
      #count how many censured at time
      kmTableRow$censoredObservationsInInterval cj=length(which(censoredTimesVector==pasteO(i,censorSymbol)
))
      kmTableRow$inRiskSetAtTime_nj=cnt_n
      #sum events and censored
      loss=kmTableRow$eventsAtEventTime ej+kmTableRow$censoredObservationsInInterval cj
      kmTableRow[c("kaplanMeirSurvivalCurveAtTime s tj-1")]=kmTable[dim(kmTable)[1],c("kaplanMeirSurvivalCu
rveAtTime s tj")]
      #TOOK LAZY WAY OUT AND JUST ADDED BACK IN THE CENSORED OBS - WILL DO CORRECT WAY LATER I Hope
      numerator=(cnt n-loss+kmTableRow$censoredObservationsInInterval cj)
      denominator=cnt n
      kmTableRow[c("c tj-1")]=paste0(numerator,"/",denominator)
      kmTableRow$kaplanMeirSurvivalCurveAtTime_s_tj=round(numerator/denominator*as.numeric(kmTable[dim(kmTa
ble)[1],c("kaplanMeirSurvivalCurveAtTime s tj")]),2)
      #update count
      cnt n=cnt n-loss
     #don't add a row when no events 0 should put this at top but no time :0
      if (kmTableRow$eventsAtEventTime ej>0 | cnt n==length(censoredTimesVectorNumeric)){
        #add row to kmtable
        kmTable=rbind(kmTable,kmTableRow)
```

```
kmTable
df2_lec_censored_noLT_KM=getKMTableNoCensorRemoval(df2_lec_censored_noLT,"+")
show(df2_lec_censored_noLT_KM)
     orderedEventTimes_tj eventsAtEventTime_ej
##
## 1
## 2
                       60
                                             1
## 3
                       62
## 4
                       63
## 5
                       65
## 6
                       66
## 7
                       68
                                             2
## 8
                       69
                                             1
                       71
## 9
     censoredObservationsInInterval cj inRiskSetAtTime nj
## 1
                                                       15
## 2
                                                       15
## 3
                                                       14
## 4
                                                       13
## 5
                                                       12
## 6
                                                       10
## 7
                                                        9
## 8
                                     2
                                                        7
## 9
                                                        4
     kaplanMeirSurvivalCurveAtTime s tj-1 c tj-1
## 1
                                        - 15/15
## 2
                                        1 14/15
## 3
                                     0.93 13/14
## 4
                                     0.86 12/13
                                     0.79 10/12
## 5
## 6
                                     0.66 9/10
## 7
                                     0.59 7/9
```

```
6/7
## 8
                                       0.46
                                               3/4
## 9
                                       0.39
     kaplanMeirSurvivalCurveAtTime s tj
##
## 1
## 2
                                     0.93
## 3
                                     0.86
## 4
                                     0.79
## 5
                                     0.66
## 6
                                     0.59
## 7
                                     0.46
## 8
                                     0.39
                                     0.29
## 9
```

4.7(a) Since the diabetics needed to survive long enough from birth until the study began, the data is left truncated. Construct a table showing the number of subjects at risk, Y, as a function of age.

```
#above code and data match output in lecture slide 33 output table - now try problem data
df2 prob LT=getKM LT Table(df2 prob,1)
show(df2 prob LT)
     tj ej cj nj c_tj-1
##
                         s_tj
## 1
      0
            0 3
                            1
## 2 60 1 0 5
                   4/5
                          0.8
## 3 62 1 0 9
                   8/9 0.7111
## 4 63 1 0 10
                         0.64
                  9/10
## 5 65 2 0 10
                  8/10 0.512
## 6 66 1 0 10
                  9/10 0.4608
## 7 68 2 0 13
                 11/13 0.3899
## 8 69 2 2 14 12/14 0.3342
## 9 70 2 0 13 11/13 0.2828
## 10 71 2 0 14 12/14 0.2424
## 11 72 2 1 14 12/14 0.2078
## 12 73 1 1 13 12/13 0.1918
## 13 74 1 1 12 11/12 0.1758
## 14 76 1 1 11 10/11 0.1598
                  9/10 0.1438
## 15 77 1 0 10
```

```
df2_prob_censored_noLT_KM=getKMTableNoCensorRemoval(df2_prob_censored_noLT,"+")
show(df2_prob_censored_noLT_KM)
      orderedEventTimes_tj eventsAtEventTime_ej
##
## 1
## 2
                        60
                                               1
## 3
                        62
## 4
                        63
## 5
                        65
## 6
                        66
## 7
                        68
## 8
                        69
## 9
                        70
                        71
## 10
## 11
                        72
                                               2
## 12
                        73
## 13
                        74
                        76
## 14
                                               1
## 15
                        77
      censoredObservationsInInterval_cj inRiskSetAtTime_nj
##
## 1
## 2
                                                         30
## 3
                                                         29
## 4
                                                         28
## 5
                                                         27
## 6
                                                         25
                                                         24
## 7
## 8
                                                         22
## 9
                                                         18
## 10
                                                         16
## 11
                                                         14
## 12
                                                         11
## 13
                                       1
                                                          9
## 14
                                                          7
                                       1
## 15
      kaplanMeirSurvivalCurveAtTime_s_tj-1 c_tj-1
##
```

```
## 1
                                           30/30
## 2
                                        1 29/30
## 3
                                     0.97 28/29
                                     0.94 27/28
## 4
## 5
                                     0.91 25/27
## 6
                                     0.84 24/25
## 7
                                     0.81 22/24
## 8
                                     0.74 20/22
## 9
                                     0.67 16/18
                                      0.6 14/16
## 10
## 11
                                     0.52 12/14
## 12
                                     0.45 10/11
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## 13
                                             8/9
## 14
                                     0.36
                                             6/7
## 15
                                     0.31
                                             4/5
##
     kaplanMeirSurvivalCurveAtTime_s_tj
## 1
                                      1
## 2
                                   0.97
## 3
                                   0.94
## 4
                                   0.91
## 5
                                   0.84
## 6
                                   0.81
## 7
                                   0.74
## 8
                                   0.67
## 9
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## 10
                                   0.52
## 11
                                   0.45
## 12
                                   0.41
## 13
                                   0.36
## 14
                                   0.31
## 15
                                   0.25
```