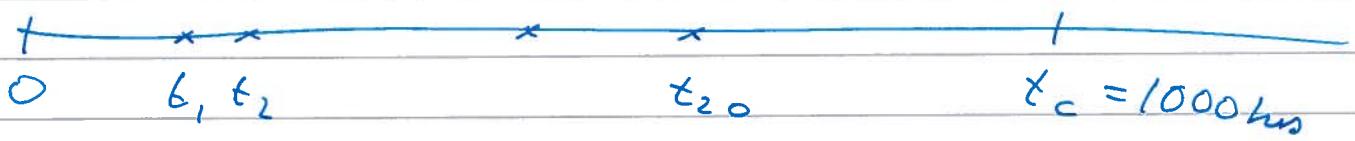


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2012

problem 1

$$n = 100$$



$$F(t; \theta) = 1 - e^{-\frac{t}{\theta}}$$

a) Likelihood:  $\prod_{i: \text{failed}} \frac{1}{\theta} e^{-\frac{t_i}{\theta}} \cdot \prod_{i: \text{cured}} e^{-\frac{t_c}{\theta}}$

$$\begin{aligned} &= \frac{1}{\theta^r} e^{-\frac{\sum_{i=1}^{20} t_i}{\theta}} \cdot e^{-\frac{(n-r)t_c}{\theta}} \\ &= \frac{e^{-\frac{\sum_{i=r+1}^{20} t_i + (n-r)t_c}{\theta}}}{\theta^r} \end{aligned}$$

so b) Log-Likelihood = 0

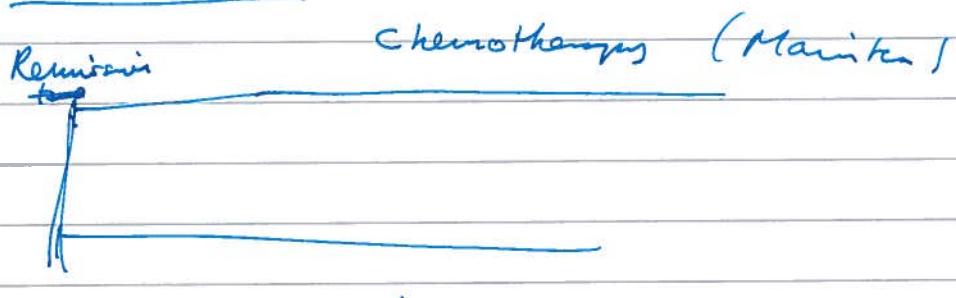
$$l(\theta) = -\frac{\sum_{i=r+1}^{20} t_i + (n-r)t_c}{\theta} - r \ln \theta \quad \cancel{\text{dL}}$$

-2-

Q2.  $\ell'(\theta) = \frac{\sum b_i + (n-r)t_c}{\theta^2} - \frac{r}{\theta} = 0$

$$\Rightarrow \hat{\theta} = \frac{\sum b_i + (n-r)t_c}{r} \left( \begin{array}{l} \text{tot. obs} \\ \text{\# failed} \end{array} \right)$$

Problem 2 :

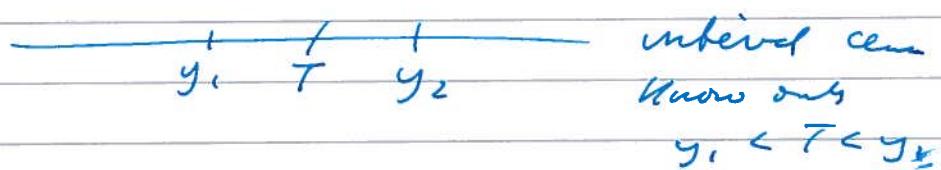


No treatment

a) Right cens

Left cens

Interval cens



Causes? end of trial, event has not occurred  
• ~~Moving~~ Dropping out (moving, e.g.)

-3

### b) Maintenance

$$R_{RM}(t) = \prod_{t_i \leq t} \frac{n_i - d_i}{n_i}$$

Maintenance gen

<u><math>R_M(t)</math></u>	<u><math>n_i</math></u>	<u><math>d_i</math></u>	<u><math>\frac{n_i - d_i}{n_i}</math></u>	<u><math>P_m</math></u>	<u>PwL. Total</u>
9	9	1	$\frac{9-1}{9} = \frac{8}{9} = 0.89$		0.889
13	8	1	$\frac{8-1}{8} = \frac{7}{8}$	$\frac{8}{9} \cdot \frac{7}{8} = \frac{7}{9}$	0.778
23	6	1	$\frac{6-1}{6} = \frac{5}{6}$	Mulitp.	0.698
34	4	1	$\frac{4-1}{4} = \frac{3}{4}$		0.486
55	2	1	$\frac{2-1}{2} = \frac{1}{2}$		0.243

For the other:

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## Distribution Analysis: Y by t

Variable: Y  
t = 1

Censoring Information Count  
Uncensored value 5  
Right censored value 4

Censoring value: d = 0

### Nonparametric Estimates

#### Characteristics of Variable

	Standard	95,0% Normal CI
Mean (MTTF)	Error	Lower Upper
63,4352	26,5864	11,3268 115,544

Median = 34  
IQR = 32 Q1 = 23 Q3 = 55

### Kaplan-Meier Estimates

Time	Number at Risk	Number Failed	Survival Probability	Standard Error	95,0% Normal CI
9	9	1	0,888889	0,104757	0,683570 1,000000
13	8	1	0,777778	0,138580	0,506166 1,000000
23	6	1	0,648148	0,165347	0,324074 0,97222
34	4	1	0,486111	0,187271	0,119066 0,85316
55	2	1	0,243056	0,195718	0,000000 0,62666

## Distribution Analysis: Y by t

Variable: Y  
t = 2

Censoring Information Count  
Uncensored value 7  
Right censored value 1

Censoring value: d = 0

### Nonparametric Estimates

#### Characteristics of Variable

	Standard	95,0% Normal CI
Mean (MTTF)	Error	Lower Upper
24,0313	5,74427	12,7727 35,2898

Median = 20  
IQR = 30 Q1 = 13 Q3 = 43

### Kaplan-Meier Estimates

Y.

Time	Number at Risk	Number Failed	Survival Probability	Standard Error	95,0% Normal CI	
5	8	1	0,87500	0,116927	0,645828	1,00000
13	7	2	0,62500	0,171163	0,289526	0,96047
20	4	1	0,46875	0,186521	0,103176	0,83432
21	3	1	0,31250	0,178152	0,000000	0,66167
43	2	1	0,15625	0,141921	0,000000	0,43441
45	1	1	0,00000	0,000000	0,000000	0,00000

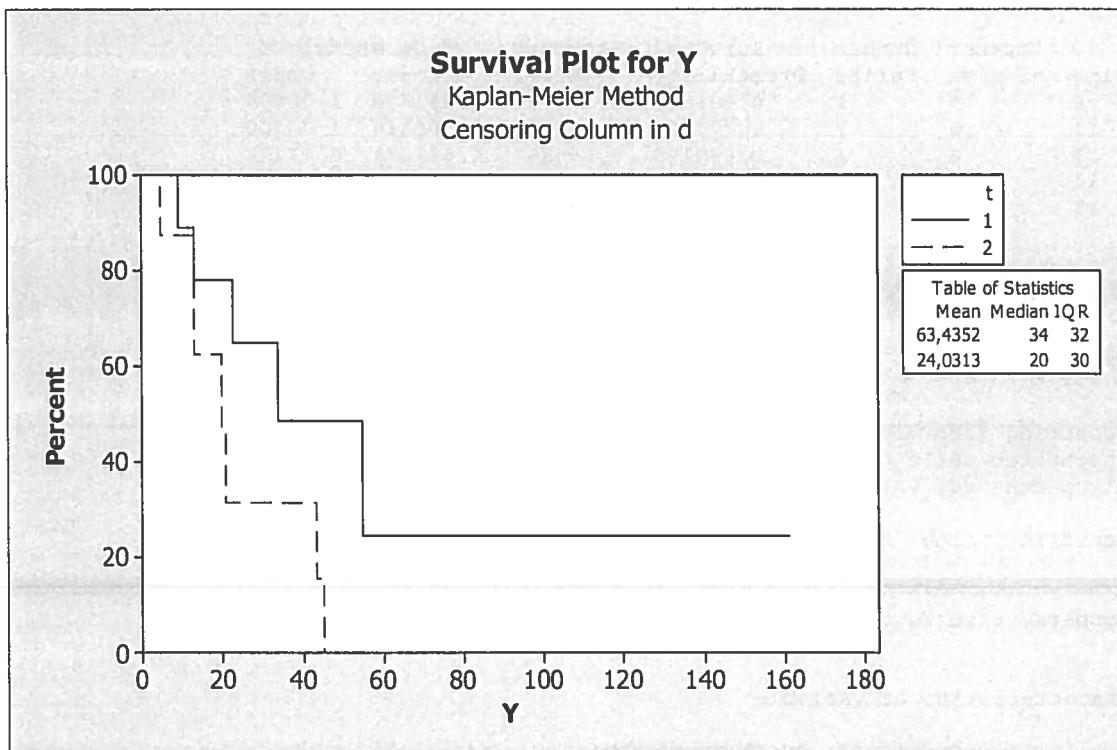
### Distribution Analysis: Y by t

Comparison of Survival Curves

#### Test Statistics

Method	Chi-Square	DF	P-Value
Log-Rank	3,05050	1	0,081
Wilcoxon	1,78571	1	0,181

### Nonparametric Survival Plot for Y



Logrank test:

$$H_0: R_M(t) = R_C(t) \text{ for all } t$$

$$\text{vs. } H_1: \neq \text{ (at least for some } t).$$

Most complete

Slides 2 :

Time	#Risk <sub>M</sub>	#Risk <sub>C</sub>	#Risk	Fail <sub>M</sub>	Fail <sub>C</sub>	Fail	EM	EC
5	9	8	17	0	1	1	$\frac{1}{17} \cdot 9$	$\frac{1}{17} \cdot 8$
9	9	7	16	1	0	1	$\frac{1}{16} \cdot 9$	$\frac{1}{16} \cdot 7$
13	8	7	15	1	2	3	$\frac{3}{15} \cdot 8$	$\frac{3}{15} \cdot 7$
⋮	⋮	⋮	⋮	⋮	⋮	⋮	⋮	⋮
							7.73	4.27

So: # Logrank statistic is

$$\frac{(5-7.73)^2}{7.73} + \frac{(7-4.27)^2}{4.27} = 2.71$$

Compare to  $\chi^2_1$ :  
 $P(\chi^2_1 > 2.71) = 0.10$   
so do not reject

## TMA4275 Lifetime analysis Spring 2009

\*

### The logrank test for comparison of survival functions

Bo Lindqvist

The logrank test is a test for equality of survival functions of two or more groups of units.

Consider here the case of two groups. Suppose that we have censored data of the usual form  $(Y_{1i}, \delta_{1i})$  for group 1 and similarly  $(Y_{2i}, \delta_{2i})$  for group 2. Let  $R_1(t)$  and  $R_2(t)$  be the true underlying survival functions for the two groups. The logrank test is a test for the null hypothesis

$$H_0 : R_1(t) = R_2(t) \text{ for all } t$$

versus the alternative hypothesis

$$H_1 : R_1(t) \neq R_2(t) \text{ for at least one } t$$

Let  $T_{(j)}$  for  $j = 1, \dots, k$  be the distinct times of observed failures when considering the two groups together. For each time  $T_{(j)}$  let  $N_{1j}$  and  $N_{2j}$  be the number of subjects "at risk" (have not yet had a failure or been censored) immediately before  $T_{(j)}$  in the two groups. Let  $N_j = N_{1j} + N_{2j}$  be the total number at risk at time  $T_{(j)}$ .

Now let  $O_{1j}$  and  $O_{2j}$  be the observed number of failures in the two groups at time  $T_{(j)}$ , and define  $O_j = O_{1j} + O_{2j}$ .

Under the null hypothesis the probability of failure at  $T_{(j)}$  would be the same for the two groups, reasonably estimated by  $O_j/N_j$ . Hence since the number at risk for the two groups are respectively  $N_{1j}$  and  $N_{2j}$ , the expected number of failures in the two groups at  $T_{(j)}$  would be respectively

$$E_{1j} = \frac{O_j}{N_j} N_{1j}$$

and

$$E_{2j} = \frac{O_j}{N_j} N_{2j}$$

Let now  $O_1 = \sum_{j=1}^k O_{1j}$  and  $O_2 = \sum_{j=1}^k O_{2j}$  be the total observed number of failures for each of the two groups, and let  $E_1 = \sum_{j=1}^k E_{1j}$  and  $E_2 = \sum_{j=1}^k E_{2j}$  be the corresponding total expected number of failures.

The logrank statistic compares expected and observed values in the two groups, and is given by

$$\frac{(O_1 - E_1)^2}{E_1} + \frac{(O_2 - E_2)^2}{E_2}$$

which is approximately chi-square distributed with 1 degree of freedom if the null hypothesis holds and the number of observations is large enough. We

therefore reject at significance level  $\alpha$  if this is larger than the upper  $\alpha$  quantile of this distribution (e.g. equal to 3.84 if  $\alpha = 0.05$ ).

Note that the logrank statistic for the case of two groups is often given in a slightly different way in the literature. However, the above statistic has the advantage of being easily generalized to the case of comparing more than two survival functions (degrees of freedom is then always the number of groups minus 1).

The logrank test is based on the same assumptions as the Kaplan-Meier estimator.

An example is given in the slides (page 68 in 2009).

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d) Cox-model

$$\hat{\lambda}(t; x) = \hat{\lambda}_0(t) \cdot e^{\beta x}$$

$x=1$  for Markt  
 $x=0$  for Control.

Mo R was used to estimate

$$\hat{\beta} = -1.06, \text{ Testing } H_0: \beta = 0 \text{ vs. } \beta \neq 0$$

with  
p-value 0.093

& likelihood ratio test  
2.92.

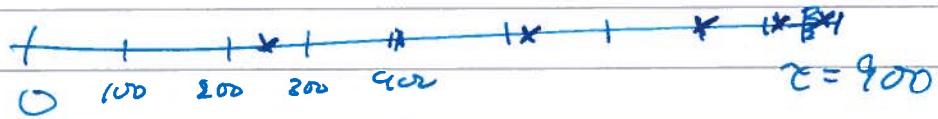
MINITAB; Weibull regression

$$\hat{\alpha} = 1.26691$$

$$\hat{\beta} = -1.06502$$

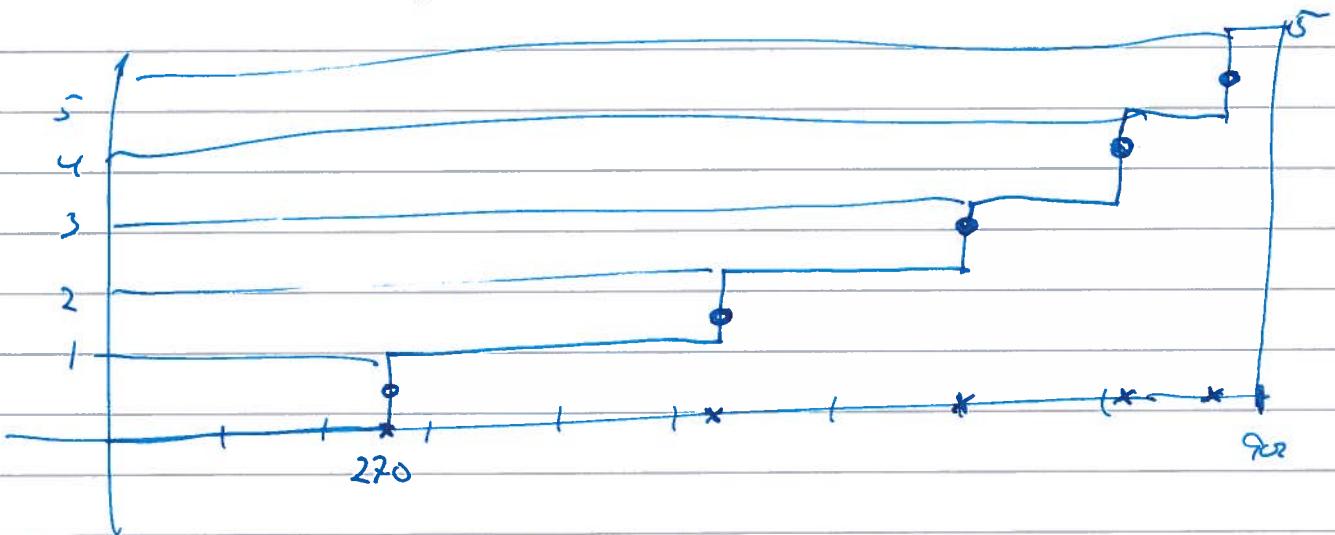
$$\begin{aligned} \text{But } \hat{\beta} &= \alpha - \alpha \hat{\beta} = -1.26691 \cdot (-1.06502) \\ &= \underline{\underline{-1.34}} \end{aligned}$$

Problem B).



a)

$$W(t) = N(t)$$



Laplace test:

$$W = \frac{\sum (S_i - \frac{C}{2})}{\tau \sqrt{\frac{n}{12}}} = \frac{(220-450) + (520-450) + (700-450) + (860-450)}{900 \sqrt{\frac{5}{12}}}$$

$$= \frac{-180 + 20 + 250 + 360 + 410}{900 \sqrt{\frac{5}{12}}} = 1.57$$

Not reject

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b)  $w(t) = e^{\alpha + \beta t}$

Log-linear model

$$\begin{aligned} W(t) &= \int_0^t e^{\alpha + \beta u} du = e^\alpha \int_0^t e^{\beta u} du \\ &= e^\alpha \cdot \left[ \frac{1}{\beta} e^{\beta u} \right]_0^t = \frac{e^\alpha}{\beta} (e^{\beta t} - 1) \end{aligned}$$

If  $\beta = 0$ : HPP with rate  $e^\alpha$

$$\underline{W(t) = e^\alpha t}$$

c) General

$$l = \sum_{i=1}^N \ln w(s_i) - W(c)$$

Here:

$$l(\alpha, \beta) = \sum_{i=1}^N (\alpha + \beta s_i) - \frac{e^\alpha}{\beta} (e^{\beta c} - 1)$$

$$= N\alpha + \beta \sum_s s_i - \frac{e^\alpha}{\beta} (e^{\beta c} - 1)$$

-10-

$$\frac{\partial l}{\partial \alpha} = N - \frac{e^\alpha}{\beta} (e^{\beta c} - 1) \quad (1)$$

$$\frac{\partial l}{\partial \beta} = S - e^\alpha \cdot \frac{\alpha e^{\beta c} \beta - (e^{\beta c} - 1)}{\beta^2} \quad (2)$$

~~$\beta - e^{\beta c}$~~

(1) = 0 gives

$$\rho N = \beta e^\alpha (e^{\beta c} - 1)$$

$$e^\alpha = \frac{\beta N}{\beta e^{\beta c} - 1} \quad \text{so ok.}$$

$$\alpha = \ln \left[ \frac{5 \cdot 0.003}{e^{0.003 \cdot 900} - 1} \right]$$

Put this into  
(ii) to solve.  
Or put  $\alpha = \ln(1)$   
into log-likelihood

to get the profile  
likelihood.

$$\hat{W}(t; \hat{\alpha}, \hat{\beta}) = \frac{e^{-6.83}}{0.003} \left( e^{0.003 \frac{t}{0.003}} - 1 \right)$$

$t$	$\hat{W}(t; \hat{\alpha}, \hat{\beta})$
270	0.450
520	1.31
700	2.58
810	3.72
860	4.38

900 : 5.0