#### TMA4275 LIFETIME ANALYSIS

Slides 11: Lifetime regression - parametric models and the semiparametric Cox-model

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#### CENSORED AND TRUNCATED DATA

- An observation is **right censored** at y: Unit is in our data, we know T > y. Contribution to L: P(T > y) = R(y).
- An observation is **left censored** at y: Unit is in our data, we know T < y. Contribution to L: P(T < y) = F(y).
- An observation is **right truncated** at *y*:
   Unit is in our data only if *T* ≤ *y*. We do not know about the units with *T* > *y*.
   Contribution to *L* of observed failure at *t*:
  - $\Delta^{-1}P(t \leq T \leq t + \Delta | T \leq y) \approx f(t)/F(y).$
- An observation is **left truncated** at y: Unit is in our data only if  $T \ge y$ . We do not know about the units with T < y.
  - Contribution to *L* of observed failure at *t*:
  - $\Delta^{-1}P(t \leq T \leq t + \Delta | T \geq y) \approx f(t)/R(y).$



#### **EXAMPLES OF LEFT TRUNCATION**

- Ultrasonic inspection of material. Signal amplitude only trusted when above limit  $\tau$ . Condition for being in the data set is  $T > \tau$ .
- Life data with pretest screening. Electronic component is burn-in tested for 1000 hours. Only the ones that passed this test are observed later. The number of components failing at burn-in is unknown. Condition for being in the data set is T>1000.

#### **EXAMPLES OF RIGHT TRUNCATION**

- Casting for automobile engine mounts. Pore size distribution below 10 microns only are recorded (other units are immediately discarded). Condition for being in the data set is  $\mathcal{T} < 10$  microns.
- Study group of individuals with AIDS diagnosis before July 1, 1986, and known date of HIV-infection (due to blood-transfu-sion). Let  $T_i = \text{time from HIV-infection to AIDS diagnosis for } i\text{th individual}$ . Then condition for being in the data set is that  $T_i \leq v_i$  where  $v_i$  is time from HIV-infection of the ith individual to July 1, 1986. (Kalbfleisch and Lawless, 1989)

# ANALYSIS OF LIFETIMES WITH COVARIATES

#### - SURVIVAL REGRESSION

**Until now:** Typically, *n* units observed, potential lifetimes

$$T_1, \ldots, T_n$$
 i.i.d.  $\sim R(t)$ 

and we have right censored data:

$$(y_i, \delta_i); \quad i = 1, \cdots, n$$

where  $y_i$  is observation time and  $\delta_i$  is censoring status for *i*th unit.

Often there exist more information which may help explain the lifetime - called *covariates* or *explanatory variables*.

This means that data are  $(y_i, \delta_i, x_i)$ , where  $x_i$  gives the values of one or more covariates/explanatory variables.

## **EXAMPLE: COMPUTER PROGRAM**

#### - EXECUTION TIME vs SYSTEM LOAD

Computer data: 17 observations of the pair (T, x) (no censorings), where T is time to complete a computationally intensive task, x is information on load from the Unix uptime command.

*Goal:* Make predictions needed for scheduling subsequent steps in a multi-step computational process.

Seconds (T)	Load (x)	Seconds (T)	Load (x)
123	2,74	110	,60
704	5,47	213	2,10
184	2,13	284	3,10
113	1,00	317	5,86
94	,32	142	1,18
76	,31	127	,57
78	,51	96	1,10
98	,29	111	1,89
240	,96	∢ □ )	· ◆♬ ▶ ◆ 壹 ▶ ◆

#### COVARIATES/EXPLANATORY VARIABLES IN RELIABILITY

Useful covariates explain/predict why some units fail quickly and some units survive a long time:

- Continuous variables like stress, temperature, voltage, and pressure.
- Discrete variables like number of hardening treatments or number of simultaneous users of a system.
- Categorical variables like manufacturer, design, and location.

Regression model relates failure time distribution to covariates  $\mathbf{x} = (x_1, \dots, x_k)$ :

$$P(T \le t) = F(t) = F(t; \mathbf{x})$$

#### WHY REGRESSION MODELS IN RELIABILITY?

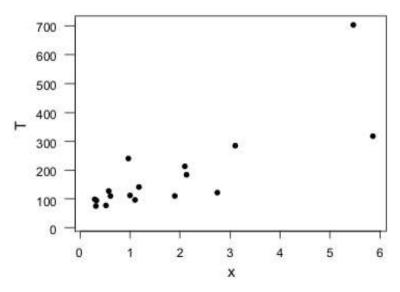
- Want to find factors which explain the reliability of an item
- Want to exclude factors which do not influence the reliability
- Obtain new knowledge about failure mechanisms
- Make better predictions for reliability of an item

#### EXPLANATORY VARIABLES IN MEDICAL RESEARCH

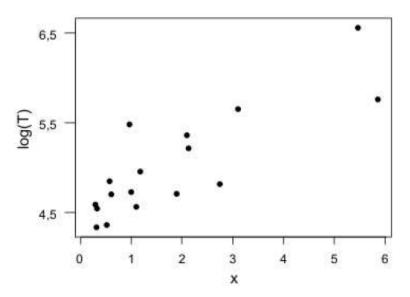
A typical medical example would include explanatory variables, often called prognostic factors or covariates, such as

- treatment assignment
- patient characteristics such as
  - age at start of study
  - gender
  - presence of other diseases at start of study

## PLOT OF COMPUTER DATA - $(x_i, T_i)$

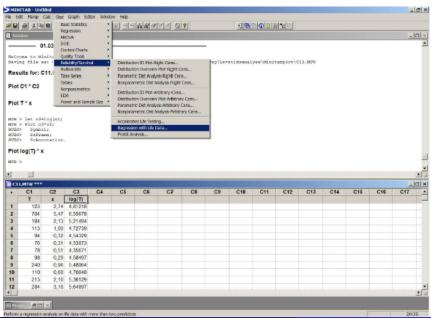


## PLOT OF COMPUTER DATA - $(x_i, ln(T_i))$



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#### **COMPUTER DATA - MINITAB**



#### REGRESSION MODELING - COMPUTER DATA

First analysis of computer data:

Simple linear regression - well known from basic statistics courses:

$$T = \beta_0 + \beta_1 x + E$$
, where  $E \sim N(0, \sigma)$  (error)

i.e.  $T \sim N(\beta_0 + \beta_1 X, \sigma)$ 

Plots suggest that it's better to take log of the data as response:

In 
$$T = \beta_0 + \beta_1 x + E$$
,  $E \sim N(0, \sigma)$ 

and we can, after taking log of all the lifetimes, use ordinary simple regression "as in basic course". Now

In 
$$T \sim N(\beta_0 + \beta_1 x, \sigma)$$

which means that

$$T \sim \text{lognormal}(\beta_0 + \beta_1 x, \sigma)$$

#### MINITAB ANALYSIS - LOGNORMAL

MINITAB does this for us in Relibility -> Survival -> Regression.

Can choose lognormal, Weibull, log-logistic etc.

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Regression with Life Data: T versus x

Response Variable: T

Censoring Information Count Uncensored value 17

Estimation Method: Maximum Likelihood Distribution: Lognormal base e
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#### Regression Table

		Standard			95,0%	Normal CI
Predictor	Coef	Error	Z	P	Lower	Upper
Intercept	4,4936	0,1112	40,39	0,000	4,2756	4,7116
x	0,29075	0,04595	6,33	0,000	0,20069	0,38080
Scale	0,31247	0,05359			0,22327	0,43730

Log-Likelihood = -89,498

Anderson-Darling (adjusted) Goodness-of-Fit

Standardized Residuals = 0,8356; Cox-Snell Residuals = 0,8170

#### ESTIMATED MODEL - LOGNORMAL

$$\ln T = 4.4936 + 0.29075 \times +0.31247 \ W, \quad W \sim N(0,1)$$

$$\hat{\beta}_0 = 4.4936$$
: "Intercept"  $\hat{\beta}_1 = 0.29075$ : "x"

 $\hat{\sigma} = 0.3147$ : "Scale"

*Note:* We could have done this by ordinary regression using  $\ln T$  as response.since there is no censoring.

#### MINITAB SURVIVAL REGRESSION

We could have done the above analysis by *ordinary linear regression* using  $\ln T$  as response, since there was *no censoring*.

But MINITAB survival regression does more:

- can have other distributions
- can have censored observations

#### PARAMETRIC SURVIVAL REGRESSION

Data:  $(y_1, \delta_1, x_1), (y_2, \delta_2, x_2), \dots, (y_n, \delta_n, x_n).$ 

*Model*: For an observation unit with covariate value x there is a potential lifetime T such that

$$\ln T = \beta_0 + \beta_1 x + \sigma U$$

where  $U \sim N(0,1)$  for lognormal, or has another standard distribution for other families.

Recall for log-logation-scale families: In  $T=\mu+\sigma U$ . The new feature is that  $\mu$  depends on x. Thus the data are no longer identically distributed but we assume

- In  $T_i = \beta_0 + \beta_1 x_i + \sigma U_i$ ;  $i = 1, \dots, n$ , where  $x_1, \dots, x_n$  may vary between units, and  $U_1, \dots, U_n$  are i.i.d., e.g., N(0,1), Gumbel(0,1) or Logistic(0,1).
- Can also have censoring, so we observe only  $Y_i = \min(T_i, C_i)$ , where  $C_i$  is a censoring time.

The extension implies estimation of Bo. By a instead of earlier usa a second

#### LIKELIHOOD FUNCTION

We need the density and survival function of an observation (T,x):

$$f(t; \beta_0, \beta_1, \sigma) = \phi(\frac{\ln t - \overbrace{\beta_0 - \beta_1 x}^{\mu \text{ before}}}{\sigma}) \frac{1}{\sigma t}$$

$$R(t; \beta_0, \beta_1, \sigma) = 1 - \Phi(\frac{\ln t - \overbrace{\beta_0 - \beta_1 x}^{\mu \text{ before}}}{\sigma})$$

So likelihood is

$$L(\beta_0, \beta_1, \sigma) = \prod_{i:\delta_i = 1} \Phi\left(\frac{\log y_i - \beta_0 - \beta_1 x_i}{\sigma}\right) \frac{1}{\sigma y_i} \cdot \prod_{i:\delta_i = 0} \left(1 - \Phi\left(\frac{\log y_i - \beta_0 - \beta_1 x_i}{\sigma}\right)\right)$$

This is maximized w.r.t parameters,  $\beta_0$ ,  $\beta_1$ ,  $\sigma$  (MINITAB!)



#### OBSERVED INFORMATION AND STANDARD ERRORS

$$I(\hat{\beta}_{0}, \hat{\beta}_{1}, \hat{\sigma}) = \begin{bmatrix} -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \beta_{0}^{2}} & -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \beta_{0}\partial \beta_{1}} & -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \beta_{0}\partial \sigma} \\ & \cdot & -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \beta_{1}^{2}} & -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \beta_{1}\partial \sigma} \\ & \cdot & \cdot & -\frac{\partial^{2}I(\beta_{0}, \beta_{1}, \sigma)}{\partial \sigma^{2}} \end{bmatrix}$$

inserted the estimated parameters. Further,

$$I(\hat{\beta}_0, \hat{\beta}_1, \hat{\sigma})^{-1} = \begin{bmatrix} \widehat{Var}\hat{\beta}_0 & \cdot & \cdot \\ \cdot & \widehat{Var}\hat{\beta}_1 & \cdot \\ \cdot & \cdot & \widehat{Var}\hat{\sigma} \end{bmatrix}$$

where as usual the entries outside the diagonal are estimated covariances.

#### ESTIMATES FOR LOGNORMAL MODEL FOR COMPUTER DATA

• Lognormal ML estimates for the computer time experiment were  $\hat{\theta} = (\hat{\beta}_0, \hat{\beta}_1, \hat{\sigma}) = (4.49, .290, .312)$  and an estimate of the variance-covariance matrix for  $\hat{\theta}$  is

$$\hat{\Sigma}_{\hat{\theta}} = \begin{bmatrix} .012 & -.0037 & 0 \\ -.0037 & .0021 & 0 \\ 0 & 0 & .0029 \end{bmatrix}.$$

 Normal-approximation confidence interval for the computer execution time regression slope is

$$\begin{split} & [\hat{\beta_1}, \quad \tilde{\beta_1}] = \hat{\beta}_1 \pm z_{(.975)} \widehat{\text{se}}_{\hat{\beta}_1} = .290 \pm 1.96 (.046) = [.20, \quad .38] \\ & \text{where } \widehat{\text{se}}_{\hat{\beta}_1} = \sqrt{.0021} = .046 \ . \end{split}$$



#### PARMETRIC MULTIPLE SURVIVAL REGRESSION

In general we may have more than one covariate.

In 
$$T = \beta_0 + \beta_1 x_1 + \dots + \beta_k x_k + \sigma U$$

$$= \beta_0 + \beta' \mathbf{x} + \sigma U$$
where  $\beta = \begin{pmatrix} \beta_1 \\ \beta_2 \\ \vdots \\ \beta_k \end{pmatrix}$ ;  $\mathbf{x} = \begin{pmatrix} x_1 \\ x_2 \\ \vdots \\ x_k \end{pmatrix}$ 

#### With data from *n* units:

$$(Y_i, \delta_i, x_{1i}, \cdots, x_{ki})$$
 or  $(Y_i, \delta_i, \mathbf{x}_i)$  for  $i = 1, 2, \dots, n$ .

In 
$$T_i = \beta_0 + \beta_1 x_{1i} + \dots + \beta_k x_{ki} + \sigma U$$
  
=  $\beta_0 + \beta' \mathbf{x}_i + \sigma U_i$ 

where  $U_1, U_2, \ldots, U_n$  are i.i.d  $\sim \Phi$ . We can extend the observed information matrix to  $(\beta_0, \cdots, \beta_k, \sigma)$ 

#### MODEL CHECKING WITH PROBABILTY PLOTS

Recall model:

$$In T_i = \beta_0 + \boldsymbol{\beta}' \mathbf{x}_i + \sigma U_i$$

which implies

$$U_i = \frac{\ln T_i - \beta_0 - \boldsymbol{\beta}' \mathbf{x}_i}{\sigma}$$

Recall also that  $U_1, U_2, \cdot, U_n$  are i.i.d  $\sim \Phi$ , and define *standardized* residuals (S-Residuals in MINITAB) by

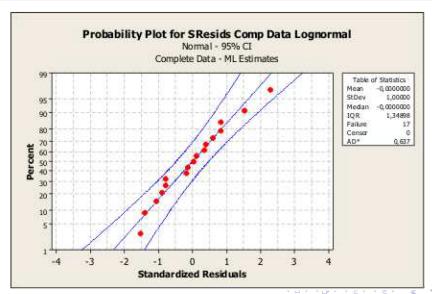
$$\hat{U}_i = \frac{\ln Y_i - \hat{\beta}_0 - \hat{\boldsymbol{\beta}}' \mathbf{x}_i}{\hat{\sigma}}$$

where  $Y_i$  are the *observed* times, either  $T_i$  or  $C_i$ .

Now the  $\hat{U}_i$  should behave like a *right censored* set from N(0,1), Gumbel(0,1), Logistic(0,1) etc.

MINITAB plots the  $\hat{U}_i$  in an ordinary probability plot for these distributions ("**Probability Plot for SResids**")

# STANDARD RESIDUALS: LOGNORMAL MODEL FOR **COMPUTER DATA**



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#### **COX-SNELL RESIDUALS**

Recall: If T has survival function R(t) and cumulative hazard Z(t), then  $Z(T) = -\ln R(T) \sim \text{expon}(1)$ 

Application here: Since In  $T_i = \beta_0 + \beta' \mathbf{x}_i + \sigma U_i$ , we have

$$R_{T_i}(t) = 1 - \Phi\left(\frac{\ln t - \beta_0 - \boldsymbol{\beta}' \mathbf{x}_i}{\sigma}\right)$$

and hence

$$V_i \equiv -\ln R_{\mathcal{T}_i}(\mathcal{T}_i) = -\ln \left[1 - \Phi\left(rac{\ln \mathcal{T}_i - eta_0 - oldsymbol{eta}'\mathbf{x}_i}{\sigma}
ight)
ight] \sim \mathsf{expon}(1)$$

Cox-Snell residuals are now defined as

$$\hat{V}_i = -\ln\left[1 - \Phi\left(\frac{\ln Y_i - \hat{eta}_0 - \hat{oldsymbol{eta}}'\mathbf{x}_i}{\hat{\sigma}}
ight)
ight] \ = -\ln\left[1 - \Phi\left( ext{standardized residuals}
ight)
ight]$$

which should behave as a set of right-censored observations from expon(1) if the model is correctly specified.

#### COX-SNELL RESIDUALS IN MINITAB

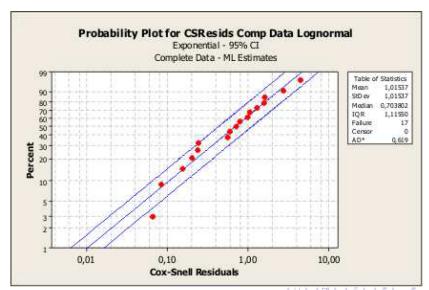
#### Recall definition:

$$\hat{V}_i = -\ln\left[1 - \Phi\left(rac{\ln Y_i - \hat{eta}_0 - \hat{eta}'\mathbf{x}_i}{\hat{\sigma}}
ight)
ight] \ = -\ln\left[1 - \Phi\left( ext{standardized residuals}
ight)
ight]$$

- MINITAB puts the Cox-Snell residuals  $\hat{V}_i$  into the usual exponential probability plot. ("CSResids").
- ullet Cox-Snell residuals are always exponentially dsitributed, while standardized residuals are distributed as the corresponding  $\Phi$  of the log-location-scale family.

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# COX-SNELL RESIDUALS: LOGNORMAL MODEL FOR COMPUTER DATA



#### SPECIAL CASE: WEIBULL REGRESSION

#### Recall:

$$\hat{V}_{i} = -\ln\left[1 - \Phi(\frac{\ln Y_{i} - \hat{\beta}_{0} - \hat{\boldsymbol{\beta}}^{'}\mathbf{x}_{i}}{\hat{\sigma}})\right] = -\ln\left[1 - \Phi(\hat{U}_{i})\right]$$

where the  $\hat{U}_i$  are the standardized residuals.

For the Weibull-distribution we have  $\Phi(w) = 1 - e^{-e^w}$ , so

$$-\ln(1-\Phi(w))=\ln(e^{-e^w})=e^w$$

Thus Cox-Snell residuals are

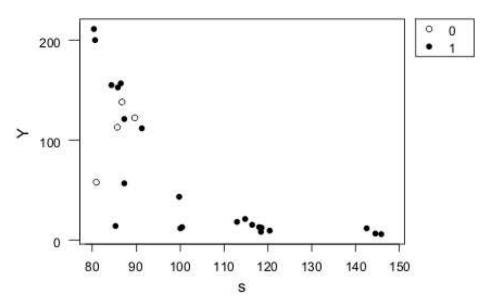
$$\hat{V}_i = e^{\hat{U}_i} = e^{\mathsf{Standardized}}$$
 residual

(Not so nice connection between SResid og CSResid for lognormal, e.g.)

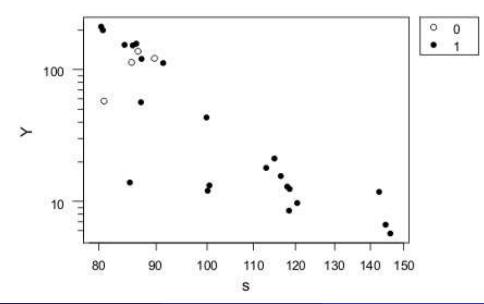
## EXAMPLE: SUPERALLOY DATA (Nelson, 1990)

Row	Pseudo-	k-Cycles	Status	(1=failed, 0=censored)
	stress			
i	s	Y	C	DATA DESCRIPTION:
1	80,3	211,629	1	Low-Cycle Fatigue Life of Nickel-Base
2	80,6	200,027	1	Superalloy Specimens
3	80,8	57,923	0	(in units of thousands of cycles
4	84,3	155,000	1	to failure).
5	85,2	13,949	1	
6	85,6	112,968	0	Data from Nelson (1990):
7	85,8	152,680	1	
8	86,4	156,725	1	SUPER ALLOY DATA
9	86,7	138,114	0	
10	87,2	56,723	1	
11	87,3	121,075	1	
12	89,7	122,372	0	
13	91,3	112,002	1	
14	99,8	43,331	1	
15	100,1	12,076	1	
16	100,5	13,181	1	
17	113,0	18,067	1	
18	114,8	21,300	1	
19	116,4	15,616	1	
20	118,0	13,030	1	
21	118,4	8,489	1	
22	118,6	12,434	1	
23	120,4	9,750	1	
24	142,5	11,865	1	
25	144,5	6,705	1	
26	145,9	5,733	1	

## SUPERALLOY DATA, PLOT (s, Y)



## SUPERALLOY DATA, PLOT $(\ln s, \ln Y)$



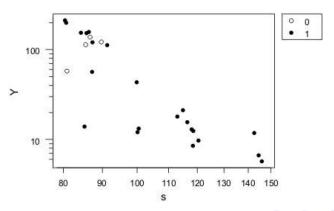
## SUPERALLOY DATA, MODEL 1

T =thousands of cycles to failure

s = pseudo-stress

 $x = \ln s = \log \text{ pseudo-stress.}$ 

**Model 1:** In  $T = \beta_0 + \beta_1 x + \sigma W$ , where W is standard Gumbel.



#### SUPERALLOY DATA, MODEL 1 - MINITAB

**Model 1:** In  $T = \beta_0 + \beta_1 x + \sigma W$ ,

Censoring Information Count
Uncensored value 22
Right censored value 4
Censoring value: C = 0

Estimation Method: Maximum Likelihood

Distribution: Weibull

Regression Table

		Standard			95,0%	Normal CI
Predictor	Coef	Error	Z	P	Lower	Upper
Intercept	31,432	2,008	15,65	0,000	27,496	35,368
x	-5,9600	0,4329	-13,77	0,000	-6,8085	-5,1116
Shape	2,2105	0,3894			1,5651	3,1221

Log-Likelihood = -97,155



#### SUPERALLOY DATA, MODEL 2 - MINITAB

**Model 2:** In 
$$T = \beta_0 + \beta_1 x + \beta_2 x^2 + \sigma W$$
,

Censoring Information Count
Uncensored value 22
Right censored value 4
Censoring value: C = 0

Estimation Method: Maximum Likelihood

Distribution: Weibull

#### Regression Table

		Standard			95,08	Normal CI
Predictor	Coef	Error	$\mathbf{z}$	P	Lower	Upper
Intercept	217,61	62,13	3,50	0,000	95,83	339,39
x	-85,52	26,55	-3,22	0,001	-137,55	-33,49
x*x	8,483	2,831	3,00	0,003	2,934	14,032
Shape	2,6685	0,4777			1,8789	3,7900

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Log-Likelihood = -93,382



## SUPERALLOY DATA, ANALYSIS

Model 2 is an example of *polynomial* survival regression:

$$\ln T = \beta_0 + \beta_1 \underbrace{x_1}_{x_1} + \beta_2 \underbrace{x_2}_{x_2} + \sigma W$$

Recall values of log likelihoods:

- Model with x only:
  - -97.155
- Model with x and x<sup>2</sup>:
   -93.382

Thus 2(difference of log-likelihoods) = 7.546 (significant at  $\alpha = 0.006$ )

Model 1: 
$$\ln T = 31.432 - 5.96 \ln s + \frac{1}{2.2105} U$$

Model 2: 
$$\ln T = 217.61 - 85.52 \ln s + 8.483 (\ln s)^2 + \frac{1}{2.6685} U$$

 $U \sim \mathsf{Gumbel}(0,1)$ 

#### PERCENTILES IN SURVIVAL REGRESSION

Recall that for log-location-scale families:

$$\ln t_p = \mu + \sigma \Phi^{-1}(p)$$

Thus, for an individual with covariate vector  $\mathbf{x}$ ,

$$\ln t_p(\mathbf{x}) = \beta_0 + \beta_1' \mathbf{x} + \sigma \Phi^{-1}(p)$$

so for Weibull regression:

$$\ln t_{p}(\mathbf{x}) = \beta_{0} + \beta_{1}^{'}\mathbf{x} + \frac{1}{\alpha}\ln(-\ln(1-p))$$

MINITAB computes these for given values of p and x

#### SUPERALLOY DATA, PERCENTILES

Regression with Life Data: Y versus  $\boldsymbol{\varkappa}$ 

Response Variable: Y

Table of Percentiles

				Standard	95,0%	Normal CI
Percent	S	x	Percentile	Error	Lower	Upper
10	80	4,3820	133,3747	34,0579	80,8565	220,0048
10	100	4,6052	16,7928	3,4263	11,2577	25,0494
10	120	4,7875	5,7830	1,2364	3,8034	8,7929
10	140	4,9416	3,6458	0,8760	2,2766	5,8386
50	80	4,3820	270,1879	56,0580	179,9121	405,7621
50	100	4,6052	34,0186	4,3027	26,5494	43,5891
50	120	4,7875	11,7151	1,5950	8,9713	15,2980
50	140	4,9416	7,3856	1,2828	5,2547	10,3807
90	80	4,3820	423,6933	90,4646	278,8097	643,8659
90	100	4,6052	53,3461	6,8162	41,5281	68,5272
90	120	4,7875	18,3709	2,4567	14,1351	23,8760
90	140	4,9416	11,5817	1,9813	8,2824	16,1952

#### SUPERALLOY DATA, COMPUTATION OF PERCENTILES

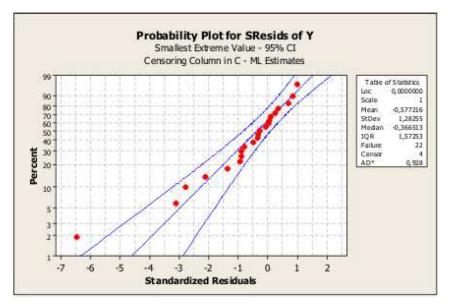
Example: 
$$p = 0.90$$
,  $s = 80$ ,  $x = \ln s = 4.3820$ ,  $x^2 = 4.3820^2$ ,  $\ln(-\ln(1 - 0.90)) = 0.8340$ 

Then

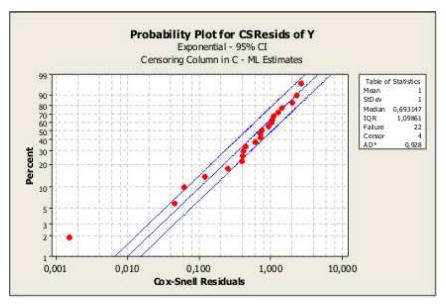
$$\begin{array}{lcl} \hat{t}_{\rho}(\mathbf{x}) & = & e^{217.61 - 85.52 \cdot 4.3820 + 8.483 \cdot 4.3820^2 + \frac{1}{2.6685} \cdot 0.8340} \\ & = & e^{6.0638} \\ & = & 430.02 \end{array}$$

(MINITAB gives 423.6933, probably rounding errors?)

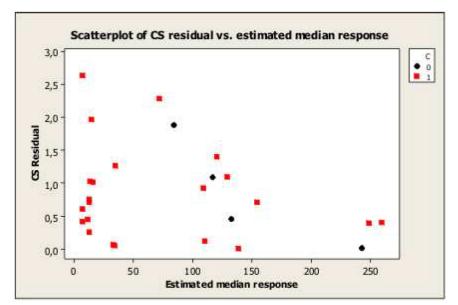
#### SUPERALLOY DATA, STANDARDIZED RESIDUALS



#### SUPERALLOY DATA, COX-SNELL RESIDUALS



#### SUPERALLOY DATA, COX-SNELL RESIDUAL PLOT



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