Weibull AFT Regression Functions in ${\sf R}$

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Weibull accelerated failure time regression can be performed in R using the **survreg** function. The results are not, however, presented in a form in which the Weibull distribution is usually given. Accelerated failure time models are usually given by

$$\log T = Y = \mu + \gamma^T \mathbf{z} + \sigma W,$$

where \mathbf{z} are set of covariates, and W has the extreme value distribution. Given tranformations

$$\begin{aligned} \alpha &= 1/\sigma, \\ \lambda &= \exp(-\mu/\sigma) \\ \boldsymbol{\beta} &= -\boldsymbol{\gamma}/\sigma, \end{aligned}$$

we have a Weibull model with baseline hazard of

$$h(x|\mathbf{z}) = (\alpha \lambda t^{\alpha - 1}) \exp(\boldsymbol{\beta}^T \mathbf{z}).$$

Further, the survreg function generally gives $\log \sigma$, rather than σ as output. The function WeibullReg (along with ConvertWeibull) solves this problem. Hazard ratios $(\exp(\beta_i))$ are additionally produced.

The function also produces the "event time ratio" (also known as "acceleration factor", $\exp(-\beta_i/\alpha) = \exp \gamma_i$), as discussed in Carroll (2003). This ratio quantifies the relative difference in time it takes to achieve the *p*th percentile between two levels of a covariate. The *p*th percentile of the (covariate-adjusted) Weibull distribution occurs at

$$t_p = \left[\frac{-\log p}{\lambda e^{\beta^T \mathbf{z}}}\right]^{1/\alpha}.$$

Then the ratio of times for a covariate with value z_1 versus values z_0 , with parameter estimate β , can then be computed as:

$$\frac{t_B}{t_A} = \left[\frac{-\log p}{\lambda e^{\beta z_1}}\right]^{1/\alpha} \left[\frac{\lambda e^{\beta z_0}}{-\log p}\right]^{1/\alpha}$$
$$= \exp\left\{\frac{\beta(z_0 - z_1)}{\alpha}\right\}.$$

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Thus, if we are comparing treatment B to treatment A, where the parameter estimate for treatment B is β_{trt} , then the ETR is $\exp\{-\beta_{trt}/\alpha\}$.

For example if the ETR for treatments A vs B is 1.2, then the amount of time it takes for p percent of patients with treatment A to have the event is predicted to be about 20% longer than it takes for the same percentage of patients with treatment B to experience an event. (That is, treatment B is worse.) For this reason, the ETR can also be called an "acceleration factor."

Additionally, a function WeibullDiag has been provided to check the adequacy of the Weibull Model.

1 WeibullReg

The WeibullReg function performs Weibull AFT regression on survival data, returning a list which contains:

formula the regression formula,

coef the coefficient table,

HR a table with the hazard rates (with confidence intervals) for each of the covariates,

ETR a table with the Event Time Ratios (with confidence intervals) for each of the covariates, and

summary the summary table from the original survreg model.

Such tables can also be produced using the streg function in stata with the following options: 1) the nohr option gives coef, 2) without any options gives HR, 3) the tr option gives ETR, and 4) the time option produces summary, the original output from survreg. While proc lifereg in SAS can also perform parametric regression for survival data, its output must also be transformed.

The following example reproduces Tables 12.1 and 12.2 from Klein and Moeschberger (2003), on the larynx data set.

```
stage2 0.16692694 0.46112943
stage3 0.66289534 0.35550887
stage4 1.74502788 0.41476410
age 0.01973646 0.01424135
```

\$HR

HRLBUBstage21.1816680.47860962.917491stage31.9404020.96667863.894946stage45.7260612.539850412.909334age1.0199330.99185731.048802

\$ETR

ETRLBUBstage20.86268630.38808791.917678stage30.55623830.29711131.041364stage40.21350900.10476190.435140age0.98268790.95838201.007610

\$summary

n= 90

```
Call:
survreg(formula = formula, data = data, dist = "weibull")
             Value Std. Error
                                 z
                                         р
(Intercept) 3.5288
                       0.9041 3.90 9.5e-05
                       0.4076 -0.36 0.717
stage2
           -0.1477
stage3
           -0.5866
                       0.3199 -1.83 0.067
           -1.5441
stage4
                       0.3633 -4.25 2.1e-05
age
           -0.0175
                       0.0128 -1.37 0.172
Log(scale) -0.1223
                       0.1225 -1.00 0.318
Scale= 0.885
Weibull distribution
Loglik(model) = -141.4 Loglik(intercept only) = -151.1
       Chisq= 19.37 on 4 degrees of freedom, p= 0.00066
```

The hazard rates produced with the Weibull regression model are similar to what is obtained with Cox proportional hazards regression:

```
> summary(coxph(Surv(time, death) ~ stage2 + stage3 + stage4 + age, data=larynx))$conf.int
```

exp(coef) exp(-coef) lower .95 upper .95 stage2 1.150320 0.8693233 0.4646755 2.847656 stage3 1.901003 0.5260381 0.9459343 3.820364

Number of Newton-Raphson Iterations: 5

stage4 5.506778 0.1815944 2.4085976 12.590147 age 1.019213 0.9811488 0.9911247 1.048098

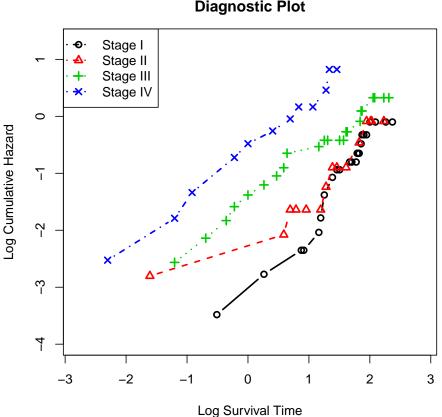
Most of the work of the function is actually performed by ConvertWeibull. These functions require the survival package in R. Formulas for the variance estimates come from Klein and Moeschberger (2003) (Equations 12.2.13-18, with some modifications since R gives $\log \sigma$).

2 WeibullDiag

The WeibullDiag function produces a diagnostic plot for Weibull AFT regression, similar to what is present Klein and Moeschberger (2003), Figure 12.2. It plots log Time versus the log of the estimated cumulative hazard estimate. If the Weibull model has adequate fit, then the plots for each of the covariates should be roughly linear and parallel. This function, at the moment, only works for one nominal covariate (or a set covariates that defines one factor). For example, if disease stage can be divided into 4 categories, one covariate can be used with levels 1:4, or alternately, 3 binary covariates. (The labels for the two types of covariates may vary, however.) It returns a table with the data used to create the plot.

This function requires both survival and prodlim.

```
> fm <- Surv(time, death) ~ stage
> diagWR <- WeibullDiag(fm, larynx, labels=c("Stage I", "Stage II", "Stage III", "Stage IV
```





References

- CARROLL, K. (2003). On the use and utility of the weibull model in the analysis of survival data. *Controlled Clinical Trials*, **24** 682–701.
- KLEIN, J. and MOESCHBERGER, M. (2003). Survival analysis: techniques for censored and truncated data. Springer.