## Homework4

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```
# import data
df = haven::read_dta("./data/umaru.dta")
```

# 1. Parametric/Accelerated Failure Time Models

a.

```
Variables: age, nonwhite, treat, site, ivdrug
Models: Exponential, Weibull, Log-logistic, Log-normal, Generalized Gamma
```

```
# define the survival object
surv_object <- Surv(time = df$time, event = df$censor)</pre>
# fit AFT models with different distributions
aft_exponential <-
 flexsurvreg(surv_object ~ age + nonwhite + treat + site + ivdrug, data = df, dist = "exponential")
aft_weibull <-
 flexsurvreg(surv_object ~ age + nonwhite + treat + site + ivdrug, data = df, dist = "weibull")
aft llogis <-
  flexsurvreg(surv_object ~ age + nonwhite + treat + site + ivdrug, data = df, dist = "llogis")
aft_lognormal <-
  flexsurvreg(surv_object ~ age + nonwhite + treat + site + ivdrug, data = df, dist = "lognormal")
aft_gen_gamma <-
  flexsurvreg(surv_object ~ age + nonwhite + treat + site + ivdrug, data = df, dist = "gengamma")
# check the results
# aft_exponential
# aft_weibull
# aft_llogis
# aft_lognormal
# aft_gen_gamma
```

i. Values of the -2 log L, the total number of parameters (including shape and scale for  $\epsilon$ ) and the AIC for each of these models

```
# extract the log-likelihood, total parameters, and AIC
results <- data.frame(
   Model = c("Exponential", "Weibull", "Log-logistic", "Log-normal", "Generalized Gamma"),
   "-2LogL" = c(
        -2 * aft_exponential$loglik,</pre>
```

```
-2 * aft_weibull$loglik,
    -2 * aft_llogis$loglik,
    -2 * aft_lognormal$loglik,
    -2 * aft_gen_gamma$loglik
  ),
  "Total Parameters" = c(
    aft_exponential$npars,
    aft weibull$npars,
    aft_llogis$npars,
    aft_lognormal$npars,
    aft_gen_gamma$npars
  ),
  AIC = c(
    aft_exponential$AIC,
    aft_weibull$AIC,
    aft_llogis$AIC,
    aft_lognormal$AIC,
    aft_gen_gamma$AIC
  )
)
colnames(results) <- c("Model", "-2 Log L", "Total Parameters", "AIC")</pre>
results |>
 kable()
```

| Model             | -2 Log L | Total Parameters | AIC      |
|-------------------|----------|------------------|----------|
| Exponential       | 6180.608 | 6                | 6192.608 |
| Weibull           | 6179.805 | 7                | 6193.805 |
| Log-logistic      | 6127.191 | 7                | 6141.191 |
| Log-normal        | 6137.951 | 7                | 6151.951 |
| Generalized Gamma | 6137.219 | 8                | 6153.219 |
|                   |          |                  |          |

The Log-logistic model has the lowest AIC (6141.191), making it the best-fitting model based on AIC.

#### ii.

Given the AIC, Weibull model does not provide an improved fit compared to the exponential model. I will confirm this using likelihood ratio test.

```
# extract log-likelihoods
logL_exp <- aft_exponential$loglik
logL_weib <- aft_weibull$loglik

# compute LRT statistic
lrt_stat <- -2 * (logL_exp - logL_weib)

# df
degf <- aft_weibull$npars - aft_exponential$npars
# p-value</pre>
```

```
p_value <- pchisq(lrt_stat, df = degf, lower.tail = FALSE)

# output results
cat("LRT Statistic:", lrt_stat)
## LRT Statistic: 0.8022232
cat("Degrees of Freedom:", degf)
## Degrees of Freedom: 1
cat("p-value:", p_value)
## p-value: 0.3704295</pre>
```

Given p-value > 0.05, we fail to reject the null hypothesis and conclude that Weibull model does not improve fit compared to the exponential model.

#### iii.

Exponential model and Weibull model are nested within the generalized gamma model. Generalized gamma model vs exponential model

```
# extract log-likelihoods
logL_exp <- aft_exponential$loglik
logL_ggamma <- aft_gen_gamma$loglik

# compute LRT statistic
lrt_stat <- -2 * (logL_exp - logL_ggamma)

# df
degf <- aft_gen_gamma$npars - aft_exponential$npars

# p-value
p_value <- pchisq(lrt_stat, df = degf, lower.tail = FALSE)

# output results
cat("LRT Statistic:", lrt_stat)
## LRT Statistic: 43.38879
cat("Degrees of Freedom:", degf)
## Degrees of Freedom: 2
cat("p-value:", p_value)
## p-value: 3.786546e-10</pre>
```

Given p-value < 0.05, we reject the null hypothesis and conclude that generalized gamma model provides a better fit compared to the exponential model.

Generalized gamma model vs Weibull model

```
# extract log-likelihoods
logL_exp <- aft_exponential$loglik
logL_weib <- aft_weibull$loglik

# compute LRT statistic
lrt_stat <- -2 * (logL_weib - logL_ggamma)
# df</pre>
```

```
degf <- aft_gen_gamma$npars - aft_weibull$npars

# p-value
p_value <- pchisq(lrt_stat, df = degf, lower.tail = FALSE)

# output results
cat("LRT Statistic:", lrt_stat)
## LRT Statistic: 42.58657
cat("Degrees of Freedom: 1
cat("p-value:", p_value)
## p-value: 6.762209e-11</pre>
```

Given p-value < 0.05, we reject the null hypothesis and conclude that generalized gamma model provides a better fit compared to the Weibull model.

#### b.

Time ratio:  $\phi = e^{\beta}$ 

```
# function to compute time ratio and confidence interval
tr ci <- function(model) {</pre>
  # extract coefficient and standard error for ivdrug
  beta <- model$res["ivdrug", "est"]</pre>
  se <- model$res["ivdrug", "se"]</pre>
  # compute time ratio (phi) and 95% confidence interval
  phi <- exp(beta)</pre>
  ci_lower <- exp(beta - 1.96 * se)</pre>
  ci\_upper \leftarrow exp(beta + 1.96 * se)
  # return as a named vector
  c("Time Ratio (phi)" = phi,
    "95% CI Lower" = ci_lower,
    "95% CI Upper" = ci_upper)
}
# apply the function to all models
results <- data.frame(</pre>
  Model = c("Exponential", "Weibull", "Log-logistic", "Log-normal", "Generalized Gamma"),
  rbind(
    tr_ci(aft_exponential),
    tr_ci(aft_weibull),
    tr_ci(aft_llogis),
    tr_ci(aft_lognormal),
    tr_ci(aft_gen_gamma)
  )
)
colnames(results) = c("Model", "Time Ratio (phi)", "95% CI Lower", "95% CI Upper")
results |>
 kable()
```

| Time Ratio (phi) | 95% CI Lower                                     | 95% CI Upper  |
|------------------|--|---|
| 1.5210016        | 1.2386592  | 1.8677018   |
| 0.6527657        | 0.5274755  | 0.8078157   |
| 0.6610930        | 0.5254482  | 0.8317545   |
| 0.6547605        | 0.5161506  | 0.8305934   |
| 0.6520866        | 0.5154833  | 0.8248900   |
|                  | 1.5210016<br>0.6527657<br>0.6610930<br>0.6547605 | 1.5210016       1.2386592         0.6527657       0.5274755         0.6610930       0.5254482         0.6547605       0.5161506 |

In the Wibull model, the estimated  $\phi$  for the ivdrug covariate is 0.653 (95%CI: 0.527-0.808). On average, individuals with IV drug use have survival times that are about 65.3% of those without IV drug use, after adjusting for other covariates. This association is statistically significant.

c.

```
Exponential model
\beta_{HR} = \beta_{AFT}
HR = e^{\beta_{HR}} = e^{\beta_{AFT}}
95\%\text{CI} = e^{\beta_{HR} \pm 1.96 \times SE_{HR}}
# extract AFT beta and SE for IV drug use (exponential model)
beta_aft_exp <- aft_exponential$res["ivdrug", "est"]</pre>
se_aft_exp <- aft_exponential$res["ivdrug", "se"]</pre>
# log-hazard ratio for exponential
beta_hr_exp <- beta_aft_exp</pre>
# HR and 95% CI
hr_exp <- exp(beta_hr_exp)</pre>
hr_exp_ci <- exp(c(beta_hr_exp - 1.96 * se_aft_exp, beta_hr_exp + 1.96 * se_aft_exp))</pre>
Webull model
\beta_{HR} = -\beta_{AFT}/\alpha, where \alpha = \text{shape parameter}
HR = e^{\beta_{HR}} = e^{\beta_{AFT}/\alpha}
95\%\text{CI} = e^{\beta_{HR} \pm 1.96 \times SE_{HR}}
# extract AFT beta and SE for IV drug use (Weibull model)
beta aft weib <- aft weibull$res["ivdrug", "est"]</pre>
se_aft_weib <- aft_weibull$res["ivdrug", "se"]</pre>
# extract Weibull shape parameter
alpha <- aft_weibull$res["shape", "est"]</pre>
# log-hazard ratio for Weibull
beta_hr_weib <- -beta_aft_weib / alpha</pre>
se_hr_weib <- se_aft_weib / alpha</pre>
# HR and 95% CI
hr_weib <- exp(beta_hr_weib)</pre>
hr_weib_ci <- exp(c(beta_hr_weib - 1.96 * se_hr_weib, beta_hr_weib + 1.96 * se_hr_weib))
```

Results:

```
# summarize results
results <- data.frame(
    Model = c("Exponential", "Weibull"),
    "Log-HR" = c(beta_hr_exp, beta_hr_weib),
    "HR" = c(hr_exp, hr_weib),
    "95% CI Lower" = c(hr_exp_ci[1], hr_weib_ci[1]),
    "95% CI Upper" = c(hr_exp_ci[2], hr_weib_ci[2])
)

colnames(results) = c("Model", "Log-HR", "HR", "95% CI Lower", "95% CI Upper")

results |>
    kable()
```

| Model               | Log-HR  | HR  | 95% CI Lower           | 95% CI Upper         |
|---------------------|---|---|------------------------|----------------------|
| Exponential Weibull | $\begin{array}{c} 0.4193691 \\ 0.4412566 \end{array}$ | $\begin{array}{c} 1.521002 \\ 1.554660 \end{array}$ | $1.238659 \\ 1.247057$ | 1.867702<br>1.938136 |

In the exponential model, the HR for IV drug use is 1.52~(95%~CI: 1.24-1.87), indicating that individuals with IV drug use have a 52% higher hazard of the event compared to those without IV drug use, holding other variables constant. Similarly, in the Weibull model, the HR is 1.55~(95%~CI: 1.25-1.94), showing consistent evidence of increased hazard associated with IV drug use.

d.

# 2. Hazard Rates and Survival from Parametric Models

a.

b.

c.

d.