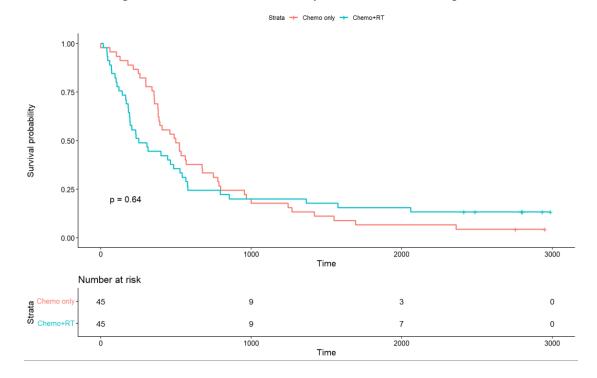
HW 4: Diagnose the validation of Cox PH model for the gastric data

Gastric cancer data are shown in "gastric cancer data.csv", where Treatment= 0 if chemotherapy only; 1 if chemotherapy with radiotherapy.

Please answer the following questions and summarize and explain your results. Computer outputs without any interpretations are not acceptable.

1. Please draw Kaplan-Meier curves stratified by treatments and interpret the result.



Log-rank test:

 H_0 : The two survival curves are identical, i.e. $S_1(t) = S_2(t)$ for all t.

 H_1 : The two survival curves are not identical, i.e. $S_1(t) \neq S_2(t)$ for some t.

Treatment=0 (化療)與 Treatment=1 (化療+放療)的生存曲線,在 1000 天左右的位置生存曲線交叉,整體上差異不大。根據 Log-rank 檢定,p 值= 0.64 (> 0.05),兩組之間的生存差異不顯著。因此,我們沒有足夠的證據認為放射治療能顯著改變病人的生存時間。

2. Please fit the data with Cox PH model and interpret the result.

```
h(t \mid \text{Treatment}) = h_0(t) \exp(\beta \cdot \text{Treatment})

H_0: \beta_{\text{Treatment}} = 0 \quad (HR = 1)

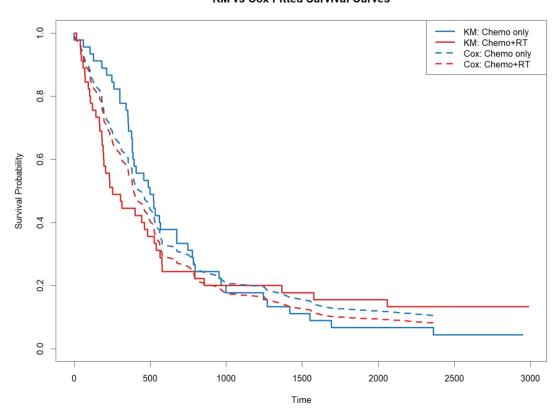
H_1: \beta_{\text{Treatment}} \neq 0 \quad (HR \neq 1)
```

```
summary(cox1)
Call:
coxph(formula = surv_obj ~ Treatment, data = df)
  n= 90, number of events= 82
            coef exp(coef) se(coef)
                                         z Pr(>|z|)
Treatment 0.1051
                    1.1109
                              0.2233 0.471
                                               0.638
          exp(coef) exp(-coef) lower .95 upper .95
              1.111
                         0.9002
                                   0.7171
Treatment
                                               1.721
Concordance= 0.562 (se = 0.031)
Likelihood ratio test= 0.22
                              on 1 df,
                                         p = 0.6
Wald test
                     = 0.22
                              on 1 df,
                                         p = 0.6
Score (logrank) test = 0.22
                              on 1 df,
                                         p = 0.6
```

根據 Cox 比例風險模型的結果:治療方式(化放療 vs. 化療)對病人存活時間的影響不顯著。化放療組的估計風險比 (HR = 1.11) 稍高於化療組,但差異並不顯著 (p = 0.638)。表示根據資料,加上放射治療並沒有明顯降低或提高死亡風險。因此,無法拒絕虛無假設,治療方式沒有顯著影響生存風險。

3. Following Question 2, draw the fitted survival curve on the same figure of Question 1 and compare with Kaplan-Meier curves. Use the result to conduct model checking.

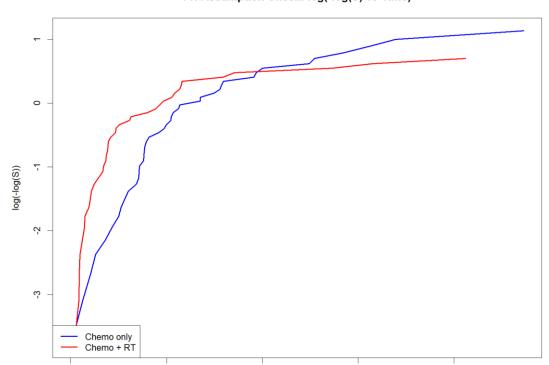
KM vs Cox Fitted Survival Curves



Cox 擬合曲線與 KM 曲線有明顯差距,有可能是因為 Cox hazard ratio 隨時間變化,比例風險假設可能不成立,所以 Cox 曲線擬合得不好。

4. Use the log-log plots to diagnose Cox model.

0



PH Assumption Check: log(-log(S) vs Time)

從 log-log plot 觀察到兩組曲線不太平行且在後期有交叉,加放療組曲線下降至低於單純化療組,顯示比例風險假設不成立。這與 Cox 擬合 KM 曲線 距離較遠的現象一致,表示 Cox 模型在此資料中對生存函數的估計不好, Cox 模型用固定危險比描述 Treatment 效果,會錯估後期加放療組的事件 風險 (生存機會)。

1000

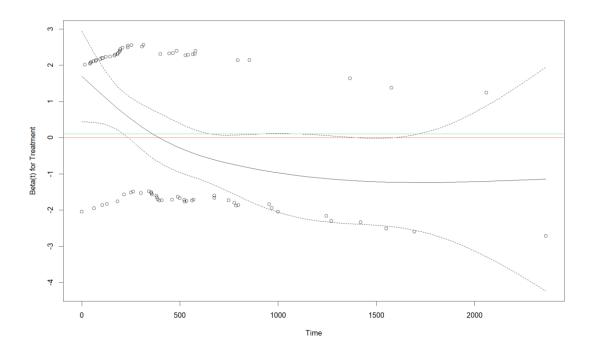
Time

500

2000

1500

5. Use Schoenfeld residuals for testing time-varying effect, and draw time-varying effect $\beta(t)$.



```
> zph.cox
chisq df p
Treatment 11 1 0.00093
GLOBAL 11 1 0.00093
```

Treatment:

 $H_0: \beta(t) = \beta$ (The hazard ratio for Treatment is constant over time, proportional hazards holds) $H_1: \beta(t) \neq \beta$ (The hazard ratio for Treatment changes over time, proportional hazards does not hold)

GLOBAL:

 H_0 : Proportional hazards assumption holds for all covariates

 H_1 : Proportional hazards assumption is violated for at least one covariate

Schoenfeld 殘差檢定顯示 Treatment 的比例風險假設不成立 (p < 0.001),表示其危險比隨時間變化;同時也違反整體模型 PH 的假設。

程式碼網址:

https://github.com/Lai-jun-

 $\underline{yan/Survival_data_analysis/tree/master/\%E4\%BD\%9C\%E6\%A5\%AD3}$