



**KARATINA UNIVERSITY**

**UNIVERSITY EXAMINATIONS  
2024/2025 ACADEMIC YEAR**

**FOURTH YEAR FIRST SEMESTER  
EXAMINATION**

**FOR THE DEGREE OF**

**BACHELOR OF SCIENCE IN ACTUARIAL SCIENCE**

**COURSE CODE: ACS 412**

**COURSE TITLE: SURVIVAL MODELS**

**DATE: 9<sup>TH</sup> DECEMBER 2024**

**TIME: 12 – 2 P.M.**

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**INSTRUCTION TO CANDIDATES**

- ANSWER ALL QUESTIONS IN SECTION A
- ANSWER ANY TWO QUESTIONS FROM SECTION B

## SECTION A

**Answer ALL QUESTIONS in this section**

### QUESTION ONE (30 marks)

- (a) Define the following terms as used in the analysis of survival data:
- (i) Censoring [2 marks]
  - (ii) Hazard function [2 marks]
  - (iii) Median Survival [2 marks]
- (b) Differentiate between the following terms:
- (i) Informative and non - informative censoring [2 marks]
  - (ii) Right and left censoring [2 marks]
- (c) The Weibull survival distribution can be characterized by hazard function
- $$\lambda(t) = \lambda \gamma t^{\gamma-1}.$$
- Determine the:
- (i) Survival function for the Weibull distribution. [3 marks]
  - (ii) Probability density function for the Weibull distribution. [2 marks]
- (d) A clinical trial to evaluate the efficacy of maintenance chemotherapy for Acute Myelogenous Leukemia (AML) was conducted. The following data shows times (weeks) of remission (i.e. freedom from symptoms in a precisely defined sense) of AML patients received chemotherapy:
- 9, 13, 13+, 18, 23, 28+, 31, 34, 45+, 48, 161+.
- Observations with plus (+) sign are right censored.
- (i) Calculate the Kaplan - Meier estimator of the survival function for this treatment group. [6 marks]
  - (ii) Calculate the 95% confidence interval for the survival function for  $\hat{S}(23)$  using the Greenwood's formula. [7 marks]
  - (iii) Compute the median survival time. [2 marks]

## SECTION B

ANSWER ANY TWO QUESTIONS FROM THIS SECTION.

### QUESTION TWO (20 marks)

The following life table shows data on time to HIV development for a sample of 100 individuals with STD but free of HIV at time 0.

Year Interval	Number of HIV positive	Number Lost to follow-up
0 – 2	1	1
2 – 4	2	1
4 – 6	8	4
6 – 8	5	8
8 – 10	5	18
10 – 12	3	20
12 – 14	8	16

Compute the life table for this data.

[20 marks]

### QUESTION THREE (20 marks)

Consider the following data set having two treatment groups (0 and 1).

Death Time	Treatment Group
1	0
2	0
3	0
4	0
5+	0
6	0
1+	1
2	1
3+	1
14	1
15	1
16	1

## ACS 412: SURVIVAL MODELS AND ANALYSIS

- (a) Compute the logrank CMH – type variance and the logrank CMH – Statistic for the data. [10 marks]
- (b) Compute the linear rank – type variance and the linear rank type – Statistic for the data. [10 marks]

### QUESTION FOUR (20 marks)

- (a) A study was done to assess the effects of treatment on the duration of steroid – induced remission for 42 acute leukaemia patients.

The variable definitions are given below:

Variable name	Description
sex	Gender (1 = male, 2 = female)
logwbc	log (base 10) white blood cell count
trt	Treatment (1 = placebo, 0 = treatment)
survtime	Time to relapse (weeks)
status	Status (0 = censored, 1 = relapse)

The following SAS code was used to fit several models to the data.

Model 1:     proc lifetest;  
              strata sex;  
              time survtime\*status (0);  
              run;

Model 2:     proc phreg;  
              model survtime\*status (0) = sex trt logwbc ;  
              run;

Model 3:     proc phreg;  
              strata sex;  
              model survtime\*status (0) = trt logwbc ;  
              run;

Model 4:     proc phreg;  
              model survtime\*status (0) = sex trt logwbc sextime;  
              sextime = log (survtime) \* sex;  
              run;

## ACS 412: SURVIVAL MODELS AND ANALYSIS

- (i) Write out the form of the Cox proportional hazards model for Model 2. [4 marks]
- (ii) Using this model, demonstrate that the hazard ratio for males versus females satisfies the assumption of “proportionality”. [3 marks]
- (iii) Which of the models 1 – 4 would *not* provide an estimate of the log hazard ratio for the gender effect? [2 marks]
- (iv) Which of the models 1 – 4 would *not* provide a test statistic (Score, Wald or LR) for the gender effect? [1 mark]
- (v) Which of the models 1 – 4 could be used as a basis for *graphically* assessing the proportional hazards assumption for sex, while controlling for the other covariates? [1 mark]
- (vi) Describe how this assessment would be done. [3 marks]
- (b) Let the event of interest be contracting lung cancer. The lung cancer hazard rate for an  $x$  – old male smoker is given by:
- $$\lambda(t) = 0.027 + 0.00025(t - 40)^2, \quad t \geq 40$$
- Assuming that a 40 – year – old male smoker is still alive at his age 50, determine the probability that he survives to age 50 without contracting lung cancer. [6 marks]

### QUESTION FIVE (20 marks)

- (a) (i) Describe the Cox regression model. [2 marks]
- (ii) Why is this a semi parametric model? [2 marks]
- (iii) Why is the Cox model often called a proportional hazards model? Explain it carefully. [2 marks]
- (b) Consider the lengths of remission (in weeks) of one group of patients who are administered with 6-MP drug in the treatment of patients with melanoma cancer. [Survival times are in weeks]

Group	Lengths of Remission (in weeks)
6-MP Drug	6, 6, 6, 6+, 7, 9+, 10, 10+, 11+, 13, 16, 17+, 19+, 20+, 22, 23, 25+, 32+, 32+, 34+, 35+

The plus observations are censored times.

## ACS 412: SURVIVAL MODELS AND ANALYSIS

- (i) Ignore the censoring indicators for treatment group (6-MP) for the moment (that is, treat all times as failure times). Compute the Kaplan-Meier estimator of the survival function for this treatment group. [5 marks]
- (ii) Plot the Kaplan-Meier estimator of the survival functions for this treatment group. [2 marks]
- (c) Given the survivorship function  $S(t) = \exp(-t^\gamma)$ ; derive the:
  - (i) Probability density function, [2 marks]
  - (ii) Hazard function, [2 marks]
  - (iii) Cumulative hazard function. [3 marks]