

Survival Analysis of Post-Myocardial Infarction Patients

Research: Alvein, Parametric: Orr, Non-Parametric: Pham

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Abstract

Background: The rates of myocardial infarction is becoming an increasing common occurrence in the United States. Rapid development of medical technology and knowledge have led to a decline in myocardial infarction fatalities (Gu, et al 1999). However, there is much to be learned regarding the survival probabilities of patients following an infarction episode. Some studies have already examined the effects of externalities on the survival rates of these patients (Rimm, et al. 1995).

Objective: Our goal is to provide detailed survival statistics of patients during a post-myocardial infarction time period with specific concern addressed to age, ventricular activity, and physiological cardiac state. Using these variables, we aim to provide succinct information on the current state of the dataset as well as provide robust predictors for the future estimates of survival for future patients.

We aim to fit non-parametric (Kaplan-Meier) and parametric curves to describe the data as well as choose a regression model to be used for predictive survivability.

Methods: Data from 133 post-myocardial infarction patients measure the time in months until death in a one year monitoring period of follow-up. We use a combination of non-parametric (Kaplan-Meier) and parametric methods (Weibull, Log-Normal, Log-Logistic, Cox PH) to determine estimates of survival among gender and physiological cardiac state (contraction depth, muscular activity, anatomical status). We fit multiple distributions over the dataset to provide current-state information of the patient dataset. Then, we regress multiple models and use combination of Akaike Information (AIC) statistics, logistic ratio tests, and residual analysis to determine model adequacy.

Results: Initial non-parametric Kaplan-Meier curve shows a median survival time of ~30 months for all age groups. We choose a Weibull regression fit (tentative) for predictive model as we have favorable AIC, ratio, and residual indicators out of all of our model.

Conclusion: Thus, for predictive model we found the Weibull regression fit to be the most ideal candidate for modeling survivability for patient groups. Additionally, when examining the survival times for the Kaplan-Meier step curve, we see that the younger age groups do survive as well as their older counterparts. Given our limited sample size for that population, we recommend continued studies into external effects of the post-myocardial episode survival.

Introduction

Heart disease has become the leading cause of US deaths among all racial and ethnic groups (Heron 2019, Fryar 2012). In 2009 cardiovascular disease represented nearly 64% of all cardiac related deaths (Dalen, et. Al. 2014). These myocardial infarction – commonly known as heart attacks - are becoming largely common among all U.S. demographic populations. As such, researchers are looking to understand the underlying causes of these episodes. Specifically, increases in cardiovascular disease (CVD) cases have been largely

attributed to many risk factors such as high levels of low-density lipoproteins (LPL), high blood pressure, and smoking (Fryar 2012).

These variables are often the results of lifestyle choices and effects of poverty. The prevalence of the disease has closely been followed a large body of conducted researchers aiming to reduce either the number of these cases or reduce the mortality of the specific myocardial infarction rates. Between 1980 and 2002, mortality rates saw a decrease of approximately 49% (Wilmot 2015). Decreases in mortality was common through the world better medical intervention techniques and increase awareness of healthier lifestyle choices became more prevalent (Goldman, et al. 1984).

Unsurprisingly, as more patients survive CVD related infarction episodes, more detail has been paid to understand the survivability the time period following an episode. Wall motion score (a measure of heart contractility during cycling) was significantly higher in those that survived versus those that died (Kan, et. Al. 1986). We hope to examine several factors that determine survivability among these patients. In addition to wall motion score, we hope to stratify and understand the relationships between time to event (death) measurements compared to general heart health and age. Our goal is to describe the survivability of our dataset and provide a model to predict the factors that determine survivability in the one-year period following a myocardial infarction episode.

Dataset

Our data was obtained our data set from Kaggle via the Reed Institute. The data set contains 133 total patient observations across 8 variables: status at the end of the survival period, age, presence of pericardial effusion, fractal shortening, EPSS, wall motion score, wall motion index, and alive at the end of one year. Three patient survival times were not given; thus, we elected to remove those values to develop the most accurate portrayal of survival times.

Since the time of myocardial infarction varies (depending if a patient joined the study prior to the start), some patients were followed for less than a year. This provides a clear censoring and truncation. We discuss the nature of censoring in the following section.

At this point, 40 points of data were missing from the total dataset. A random forest algorithm (see: missForest package) was employed to iteratively impute values. With this in mind, our predictive and summary models will have less than ideal accuracy.

We then classify continuous variables into groups for stratification.

Age is divided into three groups with 0 denoting younger than 55 years, 1 denoting 55 to 70 years, and 2 denoting older than 70 years. Pericardial effusion is already grouped into binary values with 0 denoting the absence of fluid while 1 denotes the presence of the effusion. Finally, wall motion score is divided into three groups: 0 denoting scores less than 12, 1 denoting scores of 12 - 17, and 2 denoting scores greater than 14. These groups are summarized in the table below:

Table 1: Stratification Groupings

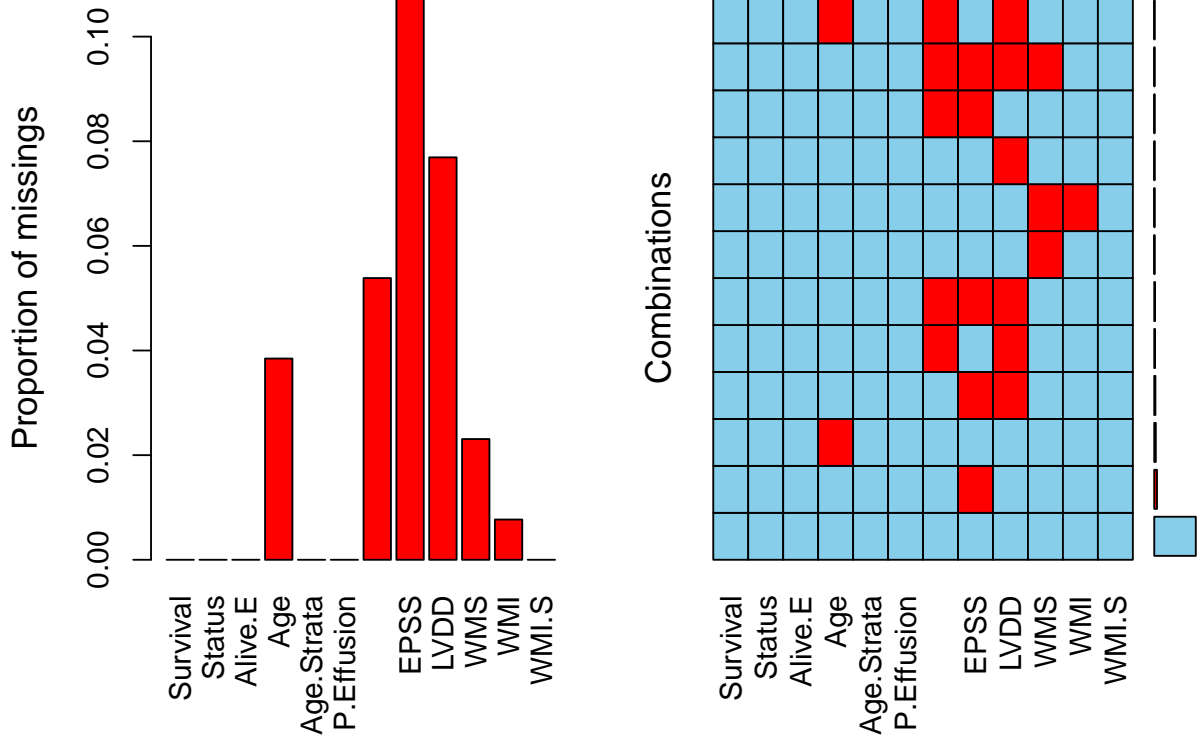
Indicator	Age	Effusion	WMS	FS
0	< 63 Years	Fluid is absent	< 11	< 0.2
1	= 63 Years	Fluid is present	= 11	= 0.2

The reader may find a summary of tables and original dataset in the appendix of this paper.

Imputation

In addition to the two rows that we removed, we further modified the dataset. The provided data contains 40 missing values that we chose to impute using the random forest algorithm methods in the missForest R

package. Below is a summary of the missing data:



missForest iteration 1 in progress... done! missForest iteration 2 in progress... done! missForest iteration 3 in progress... done!

We leverage the missForest package that uses algorithmic process used here uses a modified k-nearest neighbor (KNN) approach. Using a training data set, the routines of the algorithm predicts the missing values trained on the observed parts of the dataset (Stekhoven 2012). The process checks each iteration for an acceptable amount of error. If an iteration produces an error that is smallest than that last iteration, then the algorithm continues to function. This progress stops when an error is larger than the previous iteration. Refer to Stekhoven, et. al 2012 for more detail.

We used the missFortune package to run up to 500 iterations. Each iteration was allotted 1000 trees for the random forest algorithmic approach.

Following imputation, we verify the imputation accuracy using the normalized root mean squared error as an indicator of accuracy (NRMSE, Oba et al. (2003)). The general performance of our imputed dataset can be expressed by:

$$NRMSE = \sqrt{\frac{mean((X^{true} - X^{imp})^2)}{var(X^{true})}}$$

Where X is a matrix of our dataset. Being a random forest iterative process, each imputed dataset will be different from each other. For our particular seed and iterations, we obtained a NRMSE value of 0.1442 - that is our inputted values have an estimate 14.42% deviation from estimated true accuracy.

The full imputed dataset may be found in the appendix of this paper. As well as references to the authors who created the algorithm.

Censoring

Our dataset has numerous censored valued - that is, valued that cannot be recorded due the constraint of the study design. In our data set, we are examining the survival after a heart attack, that is, the event of interest is death given that a patient has had already survived a heart attack (left truncation).

We have fixed start and end dates for when the data was collection. Some patients joined when the study began. Others joined later after the start date. Because of this, we cannot accurately determine how long a patient survived after our observation period is over. In addition, there are some patients that have been lost to follow up or may have died due to the onset of other unrelated factors. These data present themselves as being randomly right censored.

Methodology

Here, we briefly review the methodology and theory behind our analysis techniques for context.

Non-Parametric: Kaplan Meier

We use Kaplan-Meier (KM) survival estimators to model a step curve for the survival of our censored dataset. The KM estimator is an adjustment of an empirical survival function to reflect the presence of right-censored observations (Tableman & Kim, 2004). The estimator can be described in the following equation:

$$\hat{S}(t) = \prod_{y_{(i)} \leq t} p_i = \prod_{i=1}^k \left(\frac{n_i - d_i}{n_i} \right)$$

Where n_i is the number alive before time y_i and d_i is the number of events during during that interval. In our case, y_i is the specific patient being observed, n_i is the number of patients alive at time y_i . With $k = 131$, our KM equation is:

$$\hat{S}(t) = \prod_{i=1}^{131} \left(\frac{n_i - d_i}{n_i} \right)$$

We use this equation to estimate the survival at each time interval. We conduct this analysis for the whole data set and then choose to stratify on age, pericardial effusion presence, and wall motion score. We also include cumulative hazard estimators based on the KM fit.

Cumulative Hazard Estimator

We calculate the hazard of our Kaplan-Survivor function by observing standard cumulative hazard estimate (shown below):

$$\hat{H}(t) = -\log S(t) = -\log \prod_{y_{(i)} \leq t} \frac{d_i - n_i}{n_i}$$

Intuitively, the relationship of the observed hazard is the negative log of the survival function at each interval. We can clearly see a graphical relationship between our survival by examining our hazard plots in the results section. There was the possibility of using Nelson-Aalen's approximation for hazard, but we find that the computation is trivial.

Parametric Modeling of Survival Data

Another technique for characterizing the survival function is to assume a distributional model for the data. Compared with the Kaplan-Meier approach, this method has certain advantages that include a continuous survival curve and simplicity of estimation and prediction. If the selected model accurately describes the data, it may also lend insight into the underlying mechanism for the survival behavior. This method is only applicable if a distributional model can be identified that fits the survival data adequately.

For the post-myocardial infarction dataset, We fit three commonly employed distributional models to the survival data and evaluating goodness of fit of the three models. This is accomplished by comparing the modeled survival curves to the Kaplan-Meier curve and by comparing point estimates for each model.

The three models chosen for comparison are the well-known Weibull, Lognormal, and Loglogistic distributions.

The Weibull hazard function is given below, where λ and α are the scale and shape parameters. Weibull hazard is rising if $\alpha > 1$, constant if $\alpha = 1$, and declining if $\alpha < 1$.

$$h(t) = \lambda^{-1}(-\log(1 - p))^{1/\alpha}$$

The log-normal distribution can be defined relative to the standard normal distribution; a random variable Y may be said to have the log-normal distribution if for some random variable T that has standard normal distribution:

$$\log(Y) = \alpha + \sigma T$$

The hazard function of the log-normal distribution increases with time from 0 until it reaches a maximum and then decreases, approaching 0 as time approaches infinity.

The log-logistic distribution can be defined relative to the standard logistic distribution; a random variable X may be said to have the log-logistic distribution if for some random variable S that has standard logistic distribution:

$$\log(X) = \alpha + \sigma S$$

the hazard function of the log-logistic distribution decreases with time from ∞ if $\alpha < 1$, decreases from λ if $\alpha = 1$, and if $\alpha > 1$ resembles the log-normal distribution.

Semi-Parametric Modeling of Survival Data

Where fully parametric models offer flexibility and the efficient, relatively simple estimation of overall survival function parameters, semi-parametric models offer the advantage of being well-suited to the estimation of covariate effects. Semi-parametric models decompose risk into a baseline hazard component and a relative risk component that is dependent on the covariates.

The semi-parametric Cox proportional hazards model is employed here to explore the relationship between predictor variables and survival behavior.

The Cox PH hazard function is defined as follows:

$$h(t) = h_0(t)\text{Öexp}(b_1x_1 + b_2x_2 + \dots + b_nx_n)$$

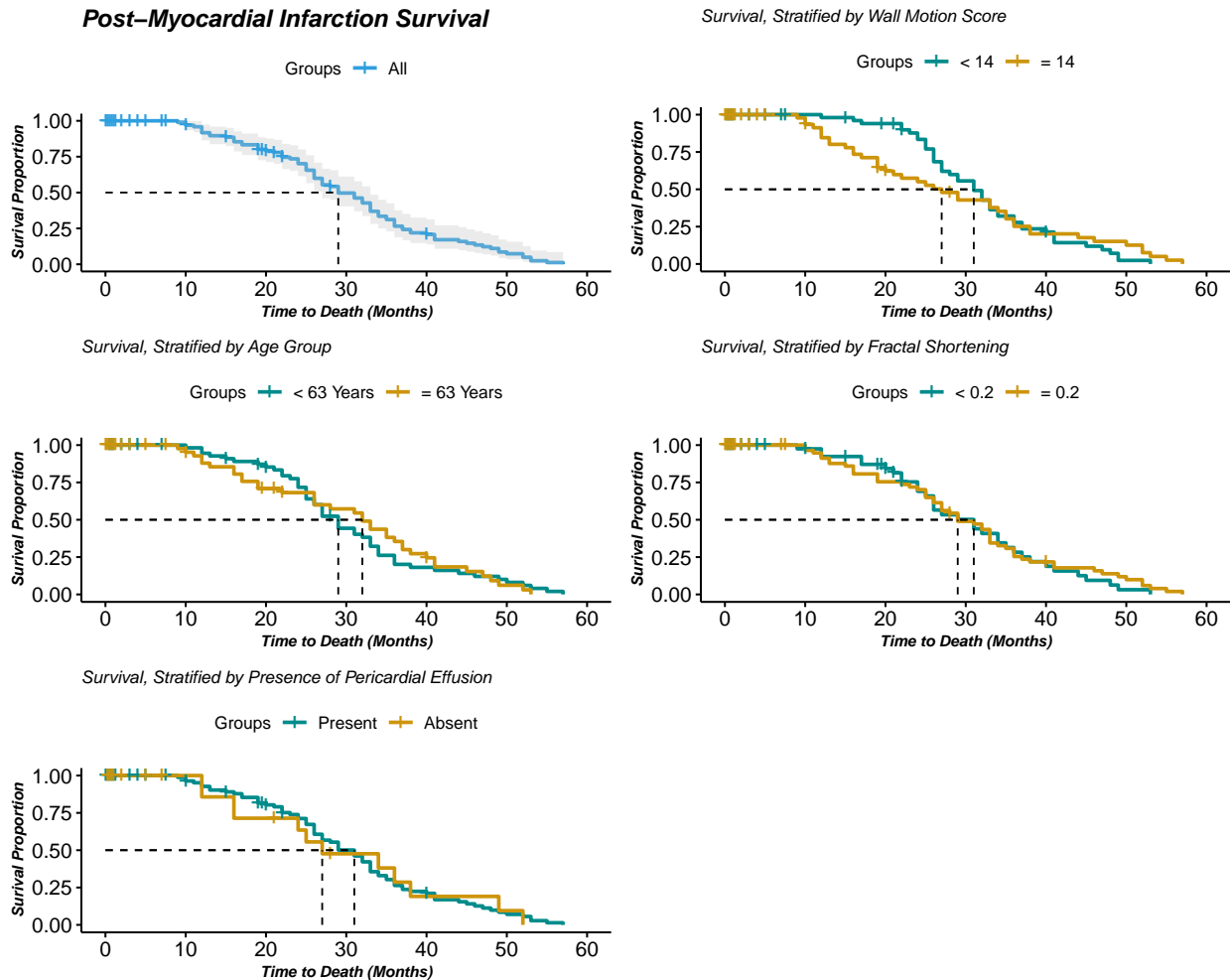
where t represents the survival time, x_1, x_2, \dots, x_n are the set of prognostic factors or covariates, and the coefficients b_1, b_2, \dots, b_n measure the effect of the covariates on survival time. The baseline hazard $h_0(t)$ corresponds to the value of the hazard if all covariates are equal to zero.

Use of the Cox PH model requires that the baseline hazard is not dependent on the covariates, and that the covariate terms do not depend on time - that is, that the slope coefficients are constant. Hazard functions stratified on covariate group may be used to assess the proportional hazard assumption. If the hazards functions for covariate groups cross over time, the proportionality assumption is not met and alternate analysis methods should be employed. One such method involves sub-setting the survival data and covariates based on hazard cross-over time. In this approach, a separate model is fit to each subset where it has been determined that the proportionality assumption holds. Alternately one may apply alternate modeling techniques that are suitable for time-varying effects, or simply investigate the impact of covariate by inspection of stratified Kaplan-Meier survival curves (Kim, 2000).

Results

Non-Parametric: Kaplan-Meier Survival Estimates

Output from our Kaplan-Meier estimators give us the following curve (full KM estimator table can be found in the appendix).

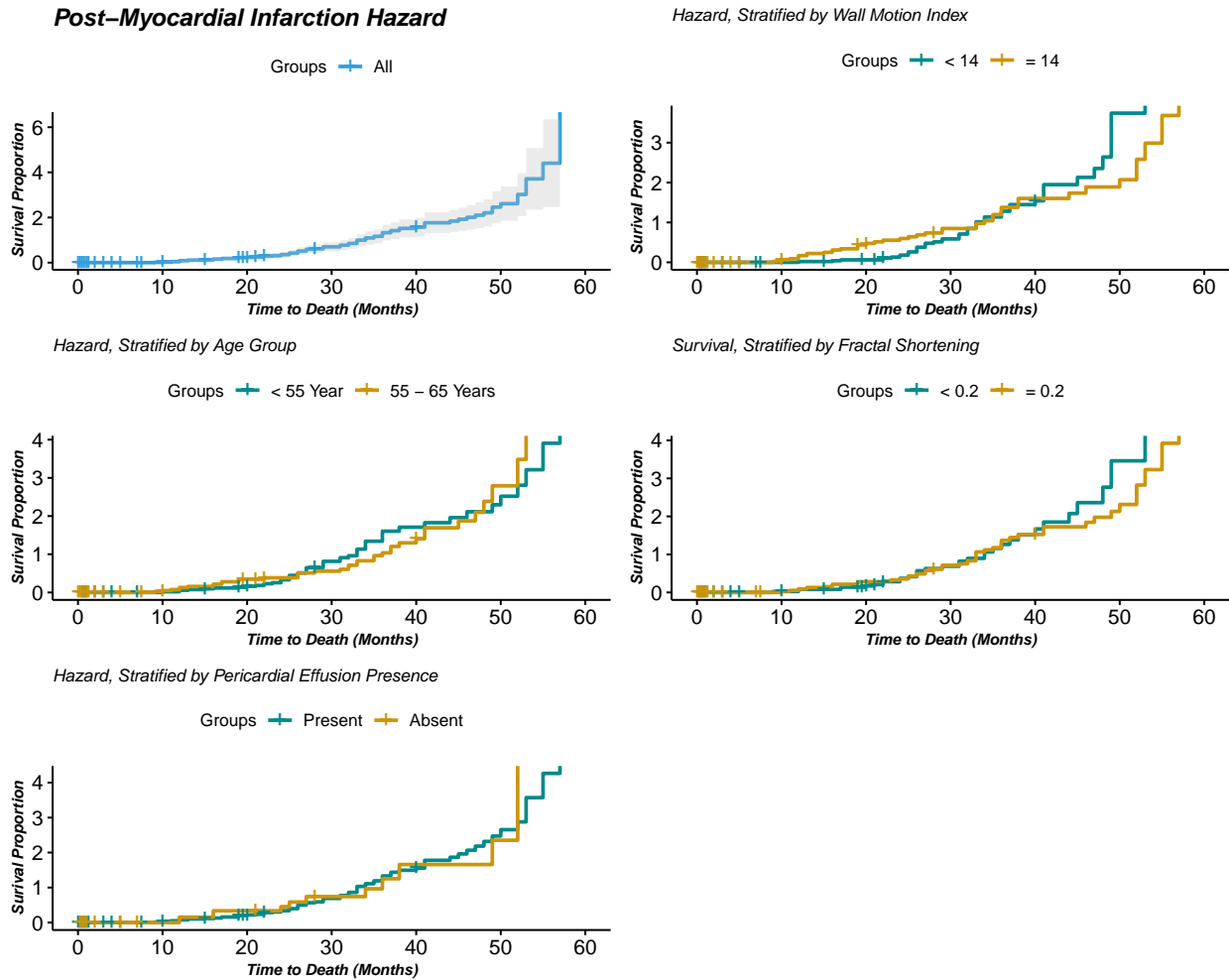


Here is the survival curve for all groups within our dataset. With time spanning to a maximum of 57 months, we have a mean survival time of approximately 30.5 months. The median survival time is approximately 29 months a 95% lower and upper confidence limits of 27 and 33 months, respectively. We see that a majority of our censored values have very short survival times. This is an intuitive finding as patients who joined the study late have relatively unknown survival (hence, their right censored nature. A summary of the results from our KM fit is below:

Table 2: Kaplan-Meier Results

	Records	Events	Mean	Median	Median 0.95 LCL	Median 0.95 UCL
All	130	88	30.53	29	27	33
Age:<55	66	51	30.47	29	26	33
Age: 55-65	64	37	30.60	32	26	37
P.Eff: Absent	106	76	30.63	31	27	33
P.Eff: Present	24	12	29.94	27	24	NA
WMS: < 14	62	46	32.17	31	27	34
WMS: = 14	68	42	28.61	27	20	35
Fractal Shortening: < 0.2	61	33	30.45	31	26	36
Fractal Shortening: = 0.2	69	55	30.44	29	26	33

Cumulative Hazard Estimates

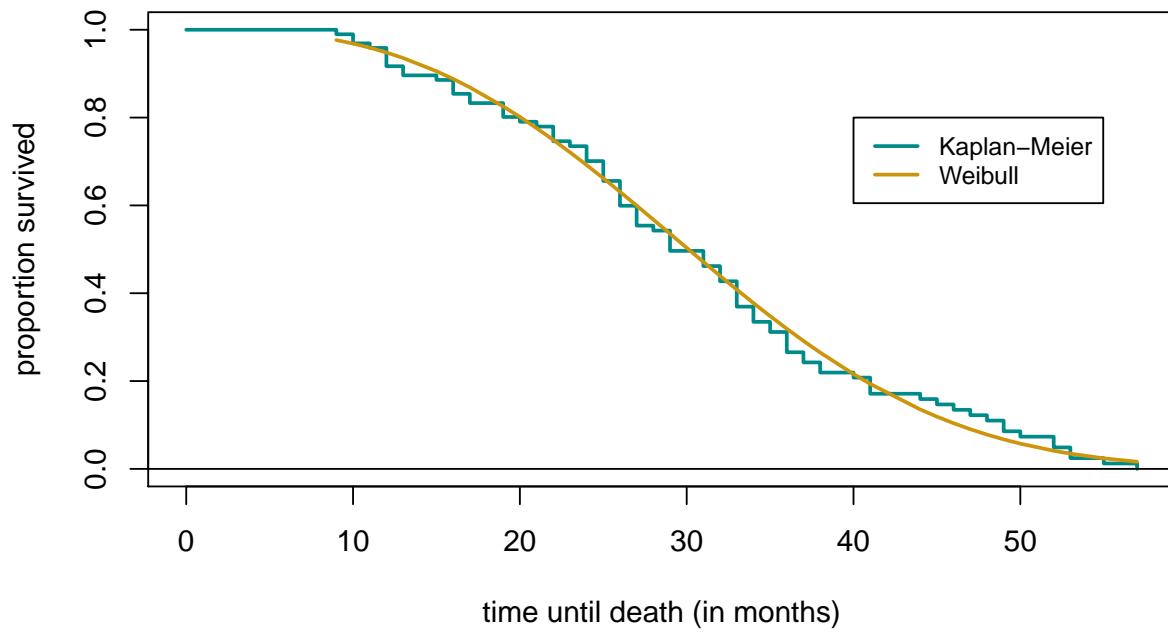


Hazard stratified by pericardial effusion shows relatively similar results between both groups. Age stratified risk shows more risk for the younger groups. Wall motion scores stratification shows a slight increase in risk past the median for higher scores.

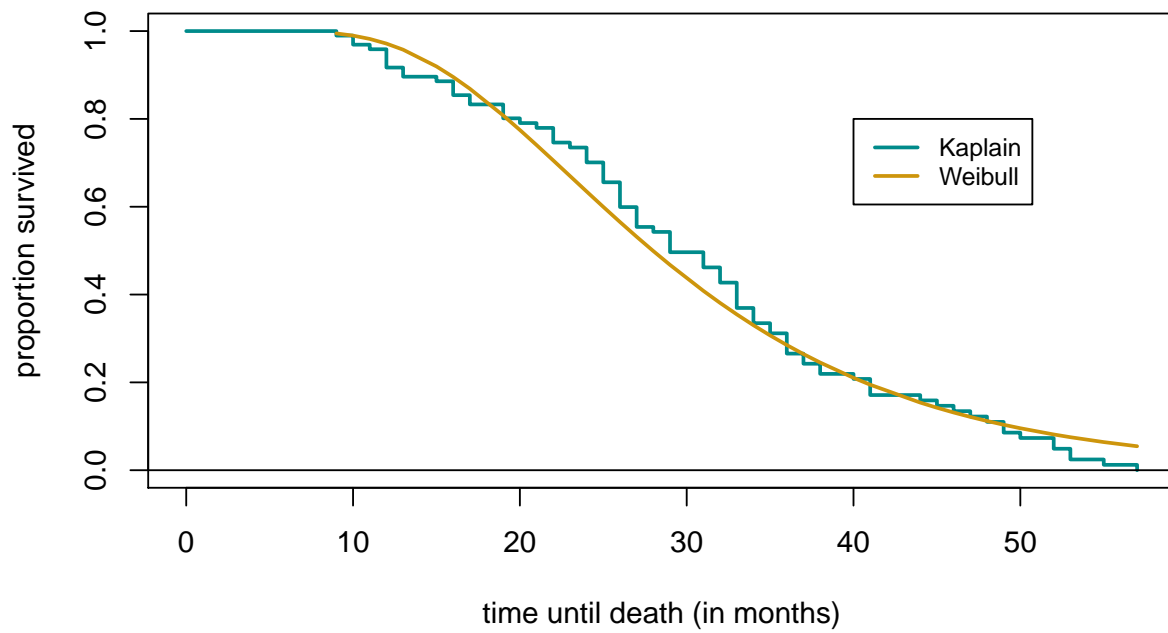
Parameter Estimation

The estimated distributional model curves are overlaid on the K-M curve for the post-myocardial infarction data in the figures below.

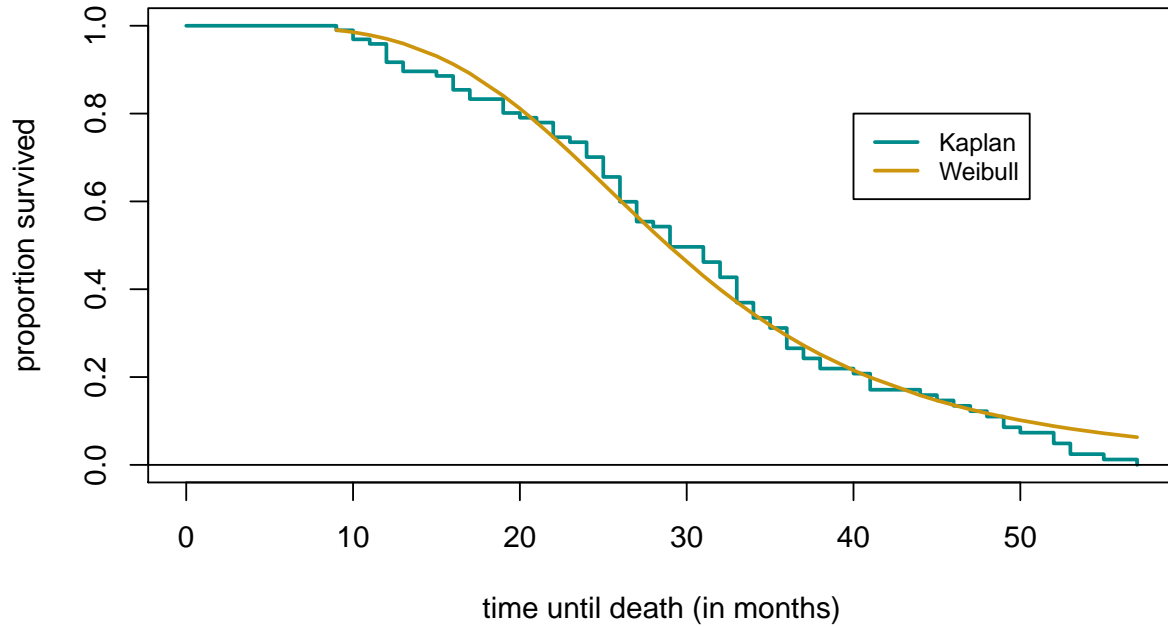
Survival Curves – Weibull and Kaplan–Meier



Survival Curves – Log-normal and Kaplan–Meier



Survival Curves – Log-logistic and Kaplan–Meier



The table below summarizes the parameter point estimates and corresponding 95% confidence intervals.

Model	Quantile	Point Estimate	95% LCL	95% UCL	Interval Length
Weibull	0.25	22.42	19.96	25.19	5.23
NA	0.50	30.32	27.86	33.01	5.15
NA	0.75	38.47	35.71	41.44	5.72
Log-normal	0.25	20.73	18.77	22.89	4.11
NA	0.50	27.97	25.54	30.63	5.09
NA	0.75	37.74	34.06	41.83	7.77
Log-logistic	0.25	21.90	19.77	24.27	4.50
NA	0.50	28.88	26.41	31.59	5.18
NA	0.75	38.09	34.45	42.12	7.67

Q-Q plots were also prepared for each distribution, and are provided in the Appendix. Based on inspection of the Q-Q plots and the K-M overlay plots, we found that the Weibull model appears to provide the best fit. In addition, it was noted that the confidence intervals for point estimation were overall more narrow for the Weibull model compared to log-normal and log-logistic. It was also observed that the point estimate for median is close to that identified by the Kaplan-Meier approach (29 months in K-M estimation, 30 months when estimating with Weibull fit). We propose the Weibull model as descriptive of the post-myocardial infarction survival data.

Regression Analysis - Cox Proportional Hazard Modeling

Regression analysis was conducted to identify the relationship between potential predictor variables and survival. As a first step, the covariate pool was screened for potential multicollinearity (see Appendix for plots of the covariates). After eliminating colinear variables, the covariate pool comprised the following factors:

- Age
- Pericardial Effusion
- Wall Motion Score
- Fractional Shortening

The proportional hazard assumption was evaluated for this model. Survival curves stratified by group were assessed for each of the four selected covariates. It was found that the proportional hazard assumption does not hold, as hazards for each group cross over with time for each covariate.

This limitation was addressed through the identification of survival time subsets in which covariates met the proportional hazards assumption, and could thus be employed as predictor variables. These subsets were identified by inspecting hazard function data stratified on all covariates and identifying regions of crossover. (See Appendix for plots of stratified hazard functions with the approximate crossover regions marked).

The selected time subsets were obtained as follows:

Subset 1 : $t \leq 26$ months Subset 2 : $26 \text{ months} < t < 46$ months Subset 3 : $46 \text{ months} \leq t$

Within each time subset, the proportional hazard assumption was tested for each of the four covariates and found to hold. A Cox PH model was fitted to each subset of the survival data, reduced as far as possible using a Step AIC procedure with Likelihood Ratio Test selection, and evaluated for adequacy using standard diagnostic techniques.

The three models are summarized below:

Characteristic	**HR**	**95% CI**	**p-value**	**HR**	**95% CI**	**p-value**
Wall_Motion_Score	0.85	0.69, 1.05	0.13	0.90	0.79, 1.02	0.11
Fractional_Shortening	0.00	0.00, 0.01	0.009			
Wall_Motion_Score * Fractional_Shortening	4.31	1.51, 12.4	0.006			
Age						

For Model 1, the significant predictors of survival time are identified as Fractional Shortening and its interaction with Wall Motion Score. Although Wall Motion Score does not have a significant effect on survival as a main effect, it is retained due to its presence in the interaction term.

For Model 2, the significant predictor of survival time is identified as patient age. Age is found to have a very weak effect on survival; none of the echocardiographic covariates had an impact on survival in this time region. Both these findings are supported by inspection of the stratified hazard function plots.

For subset 3, the sample size of events in the survival data subset was small (n=11) and a significant model was not obtained. The model is presented for context but is poorly descriptive of the relationship between prognostic factors and survival.

Model residual diagnostics were assessed for all three models. Models 1 and 2 showed overall good fit to the data and demonstrated proportional hazard assumption compliance. As expected given the discussion above, Model 3 did not result in a good overall fit, although proportional hazard assumption was met. Model diagnostic residual plots and further discussion may be found in the Appendix.

Discussion

[Add F-shortening discussion]

Explanation of results

In our non-parametric analysis, median survival times for nearly all of the stratification elements show similar results. The exception of these results are patients with pericardial effusion present and high wall motion scores. Well functioning hearts generally do not have this effusion present as [insert additional explanation of the pathology]. Wall motion scores reflecting heart muscle activity outside of the 12-14 range indicate abnormal heart function. This too, may be hinting at the survivability of post-infarction patients.

An interesting observation of non-parametric analysis when stratified on age shows a lower survivability in the younger age groups. Intuitively, we would expect higher survivability given that younger patients have more robust bodies. An explanation could be the externalities attached to having a heart attack at such young ages. For example, if a patient is young and at risk for heart attacks, then it's likely that that patient is already at risk for other factors, increasing the relative risk for that group.

In the parametric analysis, it was found that the survivor data is well-described by the Weibull distribution. This finding enables the construction of a continuous survivor function and provides the opportunity for flexible point estimation as well as estimation of survival probability over a given input time. Semi-parametric modeling was also performed. Three Cox Proportional Hazards models were developed that described the relationship between prognostic factors and survival; one for short survival times; one for intermediate survival times; and another for long survival times. Given a small sample size for long survival times, only the first two models were significant. A key finding from this analysis is that when survival time is shorter, abnormalities in the echocardiographic profile of the patient's heart are what predict survival, rather than age. Conversely, when the survival times are longer, it is the age of the patient that predicts survival (albeit weakly), rather than any of the echocardiographic properties do not. This is consistent with the behavior of the stratified hazard curves for these variables over time. This result seems to imply that in the acute recovery period following a myocardial infarction, patient prognosis is closely tied to heart health; but as time progresses, patient age becomes the key driver of prognosis. This finding is intuitive, since as treatment and recovery progress over time, risk related to the myocardial event would decrease and other pre-existing risks would play a larger role in survival.

Summary of Limitations

Very clearly, our data is smaller than we hoped for, both in the number of observations and in the availability of meaningful prognostic factors. Development of a single large regression fit that reliably predicts covariates would require a much larger dataset. In particular, the lack of gender as an available covariate is a potential limitation of our data, as the effect of gender on heart-related survival is well established in the literature. Longer collection period over a greater number of patients would yield stronger regression model development. As-is, there is not enough data to produce a significant model across the entire survival time window.

A particular limitation related to subgroup dimension was identified through the stratification analysis. Examination of stratified data showed relative unequal distributions among groups. For example, when examining pericardial effusion, we compare 106 records of with effusion absent to 24 records of effusion present. Such comparisons are unequal comparisons.

Conclusion

This study identified a variety of properties of the post-myocardial infarction survival data. The overall survival function was characterized by both non-parametric Kaplan-Meier and Weibull models, and point estimates determined. Analysis of the effect of echocardiographic data and of age on survival was performed both by inspection of stratified survival curves and by regression analysis. Findings included that age group has an impact on survival, but that this effect potentially varies over time, and that certain heart-related prognostic factors (in particular wall motion score and its interaction with fractional shortening) also have an impact on survival that varies over time. We conclude that future studies should expand on collection of variables that could potentially influence survivability, as well as the cohort size and length of follow-up

period. Interesting effects to include may include poverty, diet, ethnicity, race, sex, and even work place stress/effects. Finally, further investigation with a larger dataset on the relationship between acute survival time and echocardiographic data profile and between the relationship of age group on survival data may yield useful insights into specific risk profiles for post-myocardial infarction patients.

Appendix

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Dataset Variable Summary

Table 3: Summary of Dataset Covariates

Variable	Label	Definition
Survival	Survival	The number of months the patints survived, post-myocardial infarction.
Status	Status	Censorship status. 0 denotes that a patient is a censored while 1 denotes that a patient is uncensored.
Alive at the end of Survival Period	Alive.E	Binary variable. 0 denotes that patient is alive at the end of the survival period while 1 indicates that a patient is still alive.
Patient Age	Age	The age in years when a myocardial infarction occurs.
Age Group	Age.S	0 denotes younger than 55 years . 1 denotes 55 - 70 years. 2 denotes older than 70
Pericardial Effusion	P.Effusion	Binary variable. Pericardial effusion is excess fluid surrounding the heart. Though excess is not harmful, it is sometimes indicates a porly functioning heart. 0 denotes that pericardial effusion is absent while 1 denotes that fluid is present.
Fractional Shortening	F.Shortening	Fractional shortening is a measure of contractility around the heart. Generally, lower numbers are considered to be abnormal.
E-Point Septal Separation	EPSS	E-point septal separation is an addition measure of heart contractivity. Larger numbers are considered to be abnormal.
Left Ventricular End-Diastolic Dimension	LVDD	Left ventricular end-diastolic dimension is the measure of the heart at the end of disatole. The larger this value is indicates a larger heart. Larger hearts are generally in poor health.
Wall Motion Score	WMS	Wall motion score is a measure of how the segments of the left ventricle are moving during systol.

Table 3: Summary of Dataset Covariates (*continued*)

Variable	Label	Definition
Wall Motion Index	WMI	Wall motion index is the wall motion score divided by the number of segments that are moving. Normally, 12-13 segments can be seen in an echocardiogram.
Wall Motion Strata	WMS.S	0 denotes score less than 11, 1 denotes score 12-14, 2 denotes score greater than 14

Original Dataset

Table 4: Original Dataset

Survival	Status	Alive.E	Age	Age.Strata	LA.Effusion	L.Shortening	EF.SS	LVDD	WMS	WMI	WMI.S
11.00	1	0	71.00	2	0	0.260	9.000	4.600	14.00	1.000	0
19.00	1	0	72.00	2	0	0.380	6.000	4.100	14.00	1.700	1
16.00	1	0	55.00	1	0	0.260	4.000	3.420	14.00	1.000	0
57.00	1	0	60.00	1	0	0.253	12.062	4.603	16.00	1.450	1
19.00	0	1	57.00	1	0	0.160	22.000	5.750	18.00	2.250	1
26.00	1	0	68.00	2	0	0.260	5.000	4.310	12.00	1.000	0
13.00	1	0	62.00	1	0	0.230	31.000	5.430	22.50	1.875	1
50.00	1	0	60.00	1	0	0.330	8.000	5.250	14.00	1.000	0
19.00	1	0	46.00	0	0	0.340	0.000	5.090	16.00	1.140	0
25.00	1	0	54.00	1	0	0.140	13.000	4.490	15.50	1.190	0
10.00	0	1	77.00	2	0	0.130	16.000	4.230	18.00	1.800	1
52.00	1	0	62.00	1	1	0.450	9.000	3.600	16.00	1.140	0
52.00	1	0	73.00	2	0	0.330	6.000	4.000	14.00	1.000	0
44.00	1	0	60.00	1	0	0.150	10.000	3.730	14.00	1.000	0
0.50	0	1	62.00	1	0	0.120	23.000	5.800	11.67	2.330	1
24.00	1	0	55.00	1	1	0.250	12.063	4.290	14.00	1.000	0
0.50	0	1	69.00	2	1	0.260	11.000	4.650	18.00	1.640	1
0.50	0	1	62.53	1	1	0.070	20.000	5.200	24.00	2.000	1
22.00	0	1	66.00	2	0	0.090	17.000	5.819	8.00	1.333	1
1.00	0	1	66.00	2	1	0.220	15.000	5.400	27.00	2.250	1
0.75	0	1	69.00	2	0	0.150	12.000	5.390	19.50	1.625	1
0.75	0	1	85.00	2	1	0.180	19.000	5.460	13.83	1.380	1
0.50	0	1	73.00	2	0	0.230	12.733	6.060	7.50	1.500	1
5.00	0	1	71.00	2	0	0.170	0.000	4.650	8.00	1.000	0
48.00	1	0	64.00	1	0	0.190	5.900	3.480	10.00	1.110	0
29.00	1	0	54.00	1	0	0.300	7.000	3.850	10.00	1.667	1
29.00	1	0	35.00	0	0	0.300	5.000	4.170	14.00	1.000	0
29.00	1	0	55.00	1	0	NA	7.000	NA	2.00	1.000	0
0.25	0	1	75.00	2	0	NA	NA	NA	NA	1.000	0
36.00	1	0	55.00	1	1	0.210	4.200	4.160	14.00	1.560	1

Table 4: Original Dataset (*continued*)

Survival	Status	Alive.E	Age	Age.Strata	AEffusion	F.Shortening	FBSS	LVDD	WMS	WMI	WMI.S
1.00	0	1	65.00	2	0	0.150	NA	5.050	10.00	1.000	0
1.00	0	1	52.00	1	1	0.170	17.200	5.320	14.00	1.170	0
3.00	0	1	NA	2	0	NA	12.000	NA	6.00	3.000	1
27.00	1	0	47.00	0	0	0.400	5.120	3.100	12.00	1.000	0
35.00	1	0	63.00	1	0	NA	10.000	NA	14.00	1.170	0
26.00	1	0	61.00	1	0	0.610	13.100	4.070	13.00	1.625	1
16.00	1	0	63.00	1	1	NA	NA	5.310	5.00	1.000	0
1.00	0	1	65.00	2	0	0.060	23.600	NA	21.50	2.150	1
19.00	1	0	68.00	2	0	0.510	NA	3.880	15.00	1.670	1
31.00	1	0	80.00	2	0	0.410	5.400	4.360	NA	1.000	0
32.00	1	0	54.00	1	0	0.350	9.300	3.630	11.00	1.222	0
16.00	1	0	70.00	2	1	0.270	4.700	4.490	22.00	2.000	1
40.00	1	0	79.00	2	0	0.150	17.500	4.270	13.00	1.300	1
46.00	1	0	56.00	1	0	0.330	NA	3.590	14.00	1.000	0
2.00	0	1	67.00	2	1	0.440	9.000	3.960	17.50	1.450	1
37.00	1	0	64.00	1	0	0.090	NA	NA	12.00	2.000	1
19.50	0	1	81.00	2	0	0.120	NA	NA	9.00	1.250	0
20.00	0	1	59.00	1	0	0.030	21.300	6.290	17.00	1.310	1
0.25	0	1	63.00	1	1	NA	NA	NA	23.00	2.300	1
2.00	0	1	56.00	1	1	0.040	14.000	5.000	NA	NA	1
7.00	0	1	61.00	1	1	0.270	NA	NA	9.00	1.500	1
10.00	1	0	57.00	1	0	0.240	14.800	5.260	18.00	1.380	1
12.00	1	0	58.00	1	0	0.300	9.400	3.490	14.00	1.000	0
1.00	0	1	60.00	1	0	0.010	24.600	5.650	39.00	3.000	1
10.00	1	0	66.00	2	0	0.290	15.600	6.150	14.00	1.000	0
45.00	1	0	63.00	1	0	0.150	13.000	4.570	13.00	1.080	0
22.00	1	0	57.00	1	0	0.130	18.600	4.370	12.33	1.370	1
53.00	1	0	70.00	2	0	0.100	9.800	5.300	23.00	2.300	1
38.00	1	0	68.00	2	0	0.290	NA	4.410	14.00	1.167	0
26.00	1	0	79.00	2	0	0.170	11.900	5.150	10.50	1.050	0
9.00	1	0	73.00	2	0	0.120	NA	6.780	16.67	1.390	1
26.00	1	0	72.00	2	0	0.187	12.000	5.020	13.00	1.180	0
0.50	0	1	59.00	1	0	0.130	16.400	4.960	17.83	1.370	1
12.00	1	0	67.00	2	1	0.110	10.300	4.680	11.00	1.000	0
49.00	1	0	51.00	1	0	0.160	13.200	5.260	11.00	1.000	0
0.75	0	1	50.00	1	0	0.140	11.400	4.750	10.00	2.500	1
49.00	1	0	70.00	2	1	0.250	9.700	5.570	5.50	1.100	0
47.00	1	0	65.00	2	0	0.360	8.800	5.780	12.00	1.000	0
41.00	1	0	78.00	2	0	0.060	16.100	5.620	13.67	1.367	1
0.25	0	1	86.00	2	0	0.225	12.200	5.200	24.00	2.180	1
33.00	1	0	56.00	1	0	0.250	11.000	4.720	11.00	1.000	0
29.00	1	0	60.00	1	0	0.120	10.200	4.310	15.00	1.670	1
41.00	1	0	59.00	1	0	0.290	7.500	4.750	13.00	1.080	0
26.00	1	0	50.00	1	0	0.060	30.100	5.950	21.50	2.390	1
15.00	1	0	54.00	1	0	0.217	17.900	4.540	16.50	1.180	0
0.25	0	1	68.00	2	0	0.220	21.700	4.850	15.00	1.150	0

Table 4: Original Dataset (*continued*)

Survival	Status	Alive.E	Age	Age.Strata	AEffusion	F.Shortening	FBSS	LVDD	WMS	WMI	WMI.S
0.03	0	1	NA	2	0	0.260	19.400	4.770	21.00	2.100	1
12.00	1	0	64.00	1	0	0.200	7.100	4.580	14.00	1.000	0
32.00	1	0	63.00	1	0	0.200	5.000	5.200	8.00	1.000	0
32.00	1	0	65.00	2	0	0.060	23.600	6.740	12.00	1.090	0
27.00	1	0	54.00	1	1	0.070	16.800	4.160	18.00	1.500	1
23.00	1	0	62.00	1	0	0.250	6.000	4.480	11.00	1.000	0
0.75	0	1	78.00	2	0	0.050	10.000	4.440	15.00	1.360	1
0.75	0	1	61.00	1	0	NA	NA	NA	28.00	2.330	1
34.00	1	0	52.00	1	0	0.140	25.000	6.210	11.50	1.150	0
1.00	0	1	73.00	2	0	0.050	14.800	4.140	15.50	1.410	1
21.00	0	1	70.00	2	1	0.160	19.200	5.250	11.00	1.000	0
55.00	1	0	55.00	1	0	0.280	5.500	4.480	22.00	1.830	1
15.00	0	1	60.00	1	0	0.180	8.700	4.560	13.50	1.040	0
0.50	0	1	67.00	2	0	0.155	11.300	5.160	13.00	1.000	0
35.00	1	0	64.00	1	0	0.300	6.600	4.360	14.00	1.270	0
53.00	1	0	59.00	1	0	0.344	9.100	4.040	9.00	1.000	0
33.00	1	0	46.00	0	0	0.272	16.500	5.360	12.67	1.060	0
33.00	1	0	63.00	1	0	0.250	5.600	3.870	18.00	1.500	1
40.00	0	1	74.00	2	0	0.200	4.800	4.560	12.50	1.040	0
33.00	1	0	59.00	1	0	0.500	9.100	3.420	18.00	1.500	1
5.00	0	1	65.00	2	1	0.160	8.500	5.470	16.00	1.450	1
4.00	0	1	58.00	1	0	0.170	28.900	6.730	26.08	2.010	1
31.00	1	0	53.00	1	0	0.170	NA	4.690	10.00	1.000	0
33.00	1	0	66.00	2	0	0.200	NA	4.230	12.00	1.000	0
22.00	1	0	70.00	2	0	0.380	0.000	4.550	10.00	1.000	0
25.00	1	0	62.00	1	0	0.258	11.800	4.870	11.00	1.000	0
1.25	0	1	63.00	1	0	0.300	6.900	3.520	18.16	1.510	1
24.00	1	0	59.00	1	0	0.170	14.300	5.490	13.50	1.500	1
25.00	1	0	57.00	1	0	0.228	9.700	4.290	11.00	1.000	0
24.00	1	0	57.00	1	0	0.036	7.000	4.120	13.50	1.230	0
0.75	0	1	78.00	2	0	0.230	40.000	6.230	14.00	1.400	1
3.00	0	1	62.00	1	0	0.260	7.600	4.420	14.00	1.000	0
27.00	1	0	62.00	1	0	0.220	12.100	3.920	11.00	1.000	0
13.00	1	0	66.00	2	0	0.240	13.600	4.380	22.00	2.200	1
36.00	1	0	61.00	1	0	0.270	9.000	4.060	12.00	1.000	0
25.00	1	0	59.00	1	1	0.400	9.200	5.360	12.00	1.000	0
27.00	1	0	57.00	1	0	0.290	9.400	4.770	9.00	1.000	0
34.00	1	0	62.00	1	1	0.190	28.900	6.630	19.50	1.950	1
37.00	1	0	NA	2	0	0.260	0.000	4.380	9.00	1.000	0
34.00	1	0	54.00	1	0	0.430	9.300	4.790	10.00	1.000	0
28.00	0	1	62.00	1	1	0.240	28.600	5.860	21.50	1.950	1
28.00	1	0	NA	2	0	0.230	19.100	5.490	12.00	1.200	0
17.00	1	0	64.00	1	0	0.150	6.600	4.170	14.00	1.270	0
38.00	1	0	57.00	1	1	0.120	0.000	2.320	16.50	1.375	1
31.00	1	0	61.00	1	0	0.180	0.000	4.480	11.00	1.375	1
12.00	1	0	61.00	1	1	0.190	13.200	5.040	19.00	1.730	1

Table 4: Original Dataset (*continued*)

Survival	Status	Alive.E	Age	Age.Strata	Reffusion	F.Shortening	FPSS	LVDD	WMS	WMI	WMI.S
36.00	1	0	48.00	0	0	0.150	12.000	3.660	10.00	1.000	0
17.00	1	0	NA	2	0	0.090	6.800	4.960	13.00	1.080	0
21.00	1	0	61.00	1	0	0.140	25.500	5.160	14.00	1.270	0
7.50	0	1	64.00	1	0	0.240	12.900	4.720	12.00	1.000	0
41.00	1	0	64.00	1	0	0.280	5.400	5.470	11.00	1.100	0
36.00	1	0	69.00	2	0	0.200	7.000	5.050	14.50	1.210	0
22.00	1	0	57.00	1	0	0.140	16.100	4.360	15.00	1.360	1
20.00	1	0	62.00	1	0	0.150	0.000	4.510	15.50	1.409	1

Imputed Dataset

Table 5: Imputed Dataset

Survival	Status	Alive	E Age	P.Effusion	F.Shortening	EPSS	LVDD	WMS	WMI	Age.s	WMS.s	F.Short	LVDD.s	EPSS.s
11.00	1	0	71.00	0	0.26	9.00	4.60	14.00	1.00	1	1	1	0	0
19.00	1	0	72.00	0	0.38	6.00	4.10	14.00	1.70	1	1	1	0	0
16.00	1	0	55.00	0	0.26	4.00	3.42	14.00	1.00	0	1	1	0	0
57.00	1	0	60.00	0	0.25	12.06	4.60	16.00	1.45	0	1	1	0	1
19.00	0	1	57.00	0	0.16	22.00	5.75	18.00	2.25	0	1	0	1	1
26.00	1	0	68.00	0	0.26	5.00	4.31	12.00	1.00	1	0	1	0	0
13.00	1	0	62.00	0	0.23	31.00	5.43	22.50	1.88	0	1	1	1	1
50.00	1	0	60.00	0	0.33	8.00	5.25	14.00	1.00	0	1	1	1	0
19.00	1	0	46.00	0	0.34	0.00	5.09	16.00	1.14	0	1	1	1	0
25.00	1	0	54.00	0	0.14	13.00	4.49	15.50	1.19	0	1	0	0	1
10.00	0	1	77.00	0	0.13	16.00	4.23	18.00	1.80	1	1	0	0	1
52.00	1	0	62.00	1	0.45	9.00	3.60	16.00	1.14	0	1	1	0	0
52.00	1	0	73.00	0	0.33	6.00	4.00	14.00	1.00	1	1	1	0	0
44.00	1	0	60.00	0	0.15	10.00	3.73	14.00	1.00	0	1	0	0	0
0.50	0	1	62.00	0	0.12	23.00	5.80	11.67	2.33	0	0	0	1	1
24.00	1	0	55.00	1	0.25	12.06	4.29	14.00	1.00	0	1	1	0	1
0.50	0	1	69.00	1	0.26	11.00	4.65	18.00	1.64	1	1	1	0	0
0.50	0	1	62.53	1	0.07	20.00	5.20	24.00	2.00	0	1	0	1	1
22.00	0	1	66.00	0	0.09	17.00	5.82	8.00	1.33	1	0	0	1	1
1.00	0	1	66.00	1	0.22	15.00	5.40	27.00	2.25	1	1	1	1	1
0.75	0	1	69.00	0	0.15	12.00	5.39	19.50	1.62	1	1	0	1	1
0.75	0	1	85.00	1	0.18	19.00	5.46	13.83	1.38	1	0	0	1	1
0.50	0	1	73.00	0	0.23	12.73	6.06	7.50	1.50	1	0	1	1	1
5.00	0	1	71.00	0	0.17	0.00	4.65	8.00	1.00	1	0	0	0	0
48.00	1	0	64.00	0	0.19	5.90	3.48	10.00	1.11	1	0	0	0	0
29.00	1	0	54.00	0	0.30	7.00	3.85	10.00	1.67	0	0	1	0	0
29.00	1	0	35.00	0	0.30	5.00	4.17	14.00	1.00	0	1	1	0	0
29.00	1	0	55.00	0	0.27	7.00	4.57	2.00	1.00	0	0	1	0	0
0.25	0	1	75.00	0	0.18	12.09	5.04	12.27	1.00	1	0	0	1	1
36.00	1	0	55.00	1	0.21	4.20	4.16	14.00	1.56	0	1	1	0	0
1.00	0	1	65.00	0	0.15	11.61	5.05	10.00	1.00	1	0	0	1	1
1.00	0	1	52.00	1	0.17	17.20	5.32	14.00	1.17	0	1	0	1	1
3.00	0	1	68.34	0	0.15	12.00	5.27	6.00	3.00	1	0	0	1	1
27.00	1	0	47.00	0	0.40	5.12	3.10	12.00	1.00	0	0	1	0	0
35.00	1	0	63.00	0	0.19	10.00	4.43	14.00	1.17	1	1	0	0	0
26.00	1	0	61.00	0	0.61	13.10	4.07	13.00	1.62	0	0	1	0	1
16.00	1	0	63.00	1	0.20	9.83	5.31	5.00	1.00	1	0	1	1	0
1.00	0	1	65.00	0	0.06	23.60	5.66	21.50	2.15	1	1	0	1	1
19.00	1	0	68.00	0	0.51	7.44	3.88	15.00	1.67	1	1	1	0	0
31.00	1	0	80.00	0	0.41	5.40	4.36	11.68	1.00	1	0	1	0	0
32.00	1	0	54.00	0	0.35	9.30	3.63	11.00	1.22	0	0	1	0	0
16.00	1	0	70.00	1	0.27	4.70	4.49	22.00	2.00	1	1	1	0	0
40.00	1	0	79.00	0	0.15	17.50	4.27	13.00	1.30	1	0	0	0	1
46.00	1	0	56.00	0	0.33	8.05	3.59	14.00	1.00	0	1	1	0	0

Table 5: Imputed Dataset (*continued*)

Survival	Status	Alive	E Age	P.Effusion	F.Shortening	EPSS	LVDD	WMS	WMI	Age.s	WMS.s	F.Shortening	LVDD	EPSS.s
2.00	0	1	67.00	1	0.44	9.00	3.96	17.50	1.45	1	1	1	0	0
37.00	1	0	64.00	0	0.09	12.48	4.75	12.00	2.00	1	0	0	1	1
19.50	0	1	81.00	0	0.12	12.39	5.03	9.00	1.25	1	0	0	1	1
20.00	0	1	59.00	0	0.03	21.30	6.29	17.00	1.31	0	1	0	1	1
0.25	0	1	63.00	1	0.15	17.62	5.23	23.00	2.30	1	1	0	1	1
2.00	0	1	56.00	1	0.04	14.00	5.00	17.30	1.65	0	1	0	1	1
7.00	0	1	61.00	1	0.27	11.22	4.86	9.00	1.50	0	0	1	1	1
10.00	1	0	57.00	0	0.24	14.80	5.26	18.00	1.38	0	1	1	1	1
12.00	1	0	58.00	0	0.30	9.40	3.49	14.00	1.00	0	1	1	0	0
1.00	0	1	60.00	0	0.01	24.60	5.65	39.00	3.00	0	1	0	1	1
10.00	1	0	66.00	0	0.29	15.60	6.15	14.00	1.00	1	1	1	1	1
45.00	1	0	63.00	0	0.15	13.00	4.57	13.00	1.08	1	0	0	0	1
22.00	1	0	57.00	0	0.13	18.60	4.37	12.33	1.37	0	0	0	0	1
53.00	1	0	70.00	0	0.10	9.80	5.30	23.00	2.30	1	1	0	1	0
38.00	1	0	68.00	0	0.29	6.89	4.41	14.00	1.17	1	1	1	0	0
26.00	1	0	79.00	0	0.17	11.90	5.15	10.50	1.05	1	0	0	1	1
9.00	1	0	73.00	0	0.12	21.14	6.78	16.67	1.39	1	1	0	1	1
26.00	1	0	72.00	0	0.19	12.00	5.02	13.00	1.18	1	0	0	1	1
0.50	0	1	59.00	0	0.13	16.40	4.96	17.83	1.37	0	1	0	1	1
12.00	1	0	67.00	1	0.11	10.30	4.68	11.00	1.00	1	0	0	0	0
49.00	1	0	51.00	0	0.16	13.20	5.26	11.00	1.00	0	0	0	1	1
0.75	0	1	50.00	0	0.14	11.40	4.75	10.00	2.50	0	0	0	1	1
49.00	1	0	70.00	1	0.25	9.70	5.57	5.50	1.10	1	0	1	1	0
47.00	1	0	65.00	0	0.36	8.80	5.78	12.00	1.00	1	0	1	1	0
41.00	1	0	78.00	0	0.06	16.10	5.62	13.67	1.37	1	0	0	1	1
0.25	0	1	86.00	0	0.22	12.20	5.20	24.00	2.18	1	1	1	1	1
33.00	1	0	56.00	0	0.25	11.00	4.72	11.00	1.00	0	0	1	0	0
29.00	1	0	60.00	0	0.12	10.20	4.31	15.00	1.67	0	1	0	0	0
41.00	1	0	59.00	0	0.29	7.50	4.75	13.00	1.08	0	0	1	1	0
26.00	1	0	50.00	0	0.06	30.10	5.95	21.50	2.39	0	1	0	1	1
15.00	1	0	54.00	0	0.22	17.90	4.54	16.50	1.18	0	1	1	0	1
0.25	0	1	68.00	0	0.22	21.70	4.85	15.00	1.15	1	1	1	1	1
0.03	0	1	72.91	0	0.26	19.40	4.77	21.00	2.10	1	1	1	1	1
12.00	1	0	64.00	0	0.20	7.10	4.58	14.00	1.00	1	1	1	0	0
32.00	1	0	63.00	0	0.20	5.00	5.20	8.00	1.00	1	0	1	1	0
32.00	1	0	65.00	0	0.06	23.60	6.74	12.00	1.09	1	0	0	1	1
27.00	1	0	54.00	1	0.07	16.80	4.16	18.00	1.50	0	1	0	0	1
23.00	1	0	62.00	0	0.25	6.00	4.48	11.00	1.00	0	0	1	0	0
0.75	0	1	78.00	0	0.05	10.00	4.44	15.00	1.36	1	1	0	0	0
0.75	0	1	61.00	0	0.13	18.21	5.31	28.00	2.33	0	1	0	1	1
34.00	1	0	52.00	0	0.14	25.00	6.21	11.50	1.15	0	0	0	1	1
1.00	0	1	73.00	0	0.05	14.80	4.14	15.50	1.41	1	1	0	0	1
21.00	0	1	70.00	1	0.16	19.20	5.25	11.00	1.00	1	0	0	1	1
55.00	1	0	55.00	0	0.28	5.50	4.48	22.00	1.83	0	1	1	0	0
15.00	0	1	60.00	0	0.18	8.70	4.56	13.50	1.04	0	0	0	0	0
0.50	0	1	67.00	0	0.16	11.30	5.16	13.00	1.00	1	0	0	1	1

Table 5: Imputed Dataset (*continued*)

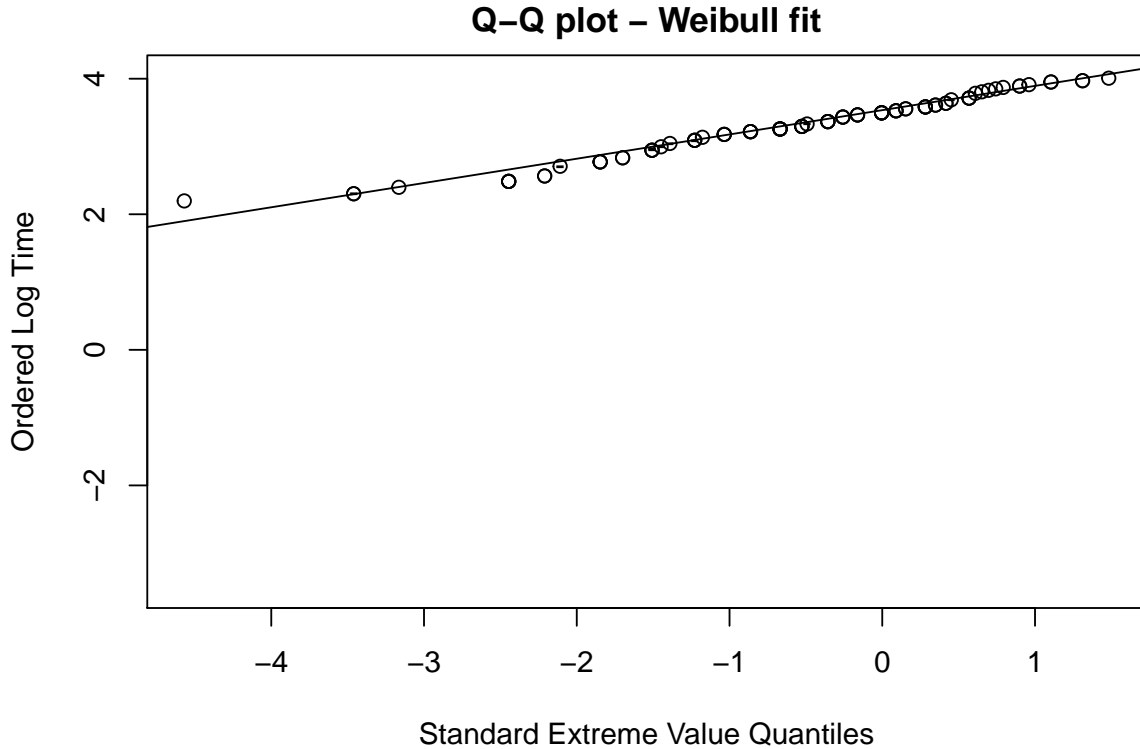
Survival	Status	Alive	E Age	P.Effusion	F.Shortening	EPSS	LVDD	WMS	WMI	Age.s	WMS.s	F.Shortening	LVDD.s	EPSS.s
35.00	1	0	64.00	0	0.30	6.60	4.36	14.00	1.27	1	1	1	0	0
53.00	1	0	59.00	0	0.34	9.10	4.04	9.00	1.00	0	0	1	0	0
33.00	1	0	46.00	0	0.27	16.50	5.36	12.67	1.06	0	0	1	1	1
33.00	1	0	63.00	0	0.25	5.60	3.87	18.00	1.50	1	1	1	0	0
40.00	0	1	74.00	0	0.20	4.80	4.56	12.50	1.04	1	0	1	0	0
33.00	1	0	59.00	0	0.50	9.10	3.42	18.00	1.50	0	1	1	0	0
5.00	0	1	65.00	1	0.16	8.50	5.47	16.00	1.45	1	1	0	1	0
4.00	0	1	58.00	0	0.17	28.90	6.73	26.08	2.01	0	1	0	1	1
31.00	1	0	53.00	0	0.17	10.30	4.69	10.00	1.00	0	0	0	0	0
33.00	1	0	66.00	0	0.20	8.12	4.23	12.00	1.00	1	0	1	0	0
22.00	1	0	70.00	0	0.38	0.00	4.55	10.00	1.00	1	0	1	0	0
25.00	1	0	62.00	0	0.26	11.80	4.87	11.00	1.00	0	0	1	1	1
1.25	0	1	63.00	0	0.30	6.90	3.52	18.16	1.51	1	1	1	0	0
24.00	1	0	59.00	0	0.17	14.30	5.49	13.50	1.50	0	0	0	1	1
25.00	1	0	57.00	0	0.23	9.70	4.29	11.00	1.00	0	0	1	0	0
24.00	1	0	57.00	0	0.04	7.00	4.12	13.50	1.23	0	0	0	0	0
0.75	0	1	78.00	0	0.23	40.00	6.23	14.00	1.40	1	1	1	1	1
3.00	0	1	62.00	0	0.26	7.60	4.42	14.00	1.00	0	1	1	0	0
27.00	1	0	62.00	0	0.22	12.10	3.92	11.00	1.00	0	0	1	0	1
13.00	1	0	66.00	0	0.24	13.60	4.38	22.00	2.20	1	1	1	0	1
36.00	1	0	61.00	0	0.27	9.00	4.06	12.00	1.00	0	0	1	0	0
25.00	1	0	59.00	1	0.40	9.20	5.36	12.00	1.00	0	0	1	1	0
27.00	1	0	57.00	0	0.29	9.40	4.77	9.00	1.00	0	0	1	1	0
34.00	1	0	62.00	1	0.19	28.90	6.63	19.50	1.95	0	1	0	1	1
37.00	1	0	69.34	0	0.26	0.00	4.38	9.00	1.00	1	0	1	0	0
34.00	1	0	54.00	0	0.43	9.30	4.79	10.00	1.00	0	0	1	1	0
28.00	0	1	62.00	1	0.24	28.60	5.86	21.50	1.95	0	1	1	1	1
28.00	1	0	69.19	0	0.23	19.10	5.49	12.00	1.20	1	0	1	1	1
17.00	1	0	64.00	0	0.15	6.60	4.17	14.00	1.27	1	1	0	0	0
38.00	1	0	57.00	1	0.12	0.00	2.32	16.50	1.38	0	1	0	0	0
31.00	1	0	61.00	0	0.18	0.00	4.48	11.00	1.38	0	0	0	0	0
12.00	1	0	61.00	1	0.19	13.20	5.04	19.00	1.73	0	1	0	1	1
36.00	1	0	48.00	0	0.15	12.00	3.66	10.00	1.00	0	0	0	0	1
17.00	1	0	69.67	0	0.09	6.80	4.96	13.00	1.08	1	0	0	1	0
21.00	1	0	61.00	0	0.14	25.50	5.16	14.00	1.27	0	1	0	1	1
7.50	0	1	64.00	0	0.24	12.90	4.72	12.00	1.00	1	0	1	0	1
41.00	1	0	64.00	0	0.28	5.40	5.47	11.00	1.10	1	0	1	1	0
36.00	1	0	69.00	0	0.20	7.00	5.05	14.50	1.21	1	1	1	1	0
22.00	1	0	57.00	0	0.14	16.10	4.36	15.00	1.36	0	1	0	0	1
20.00	1	0	62.00	0	0.15	0.00	4.51	15.50	1.41	0	1	0	0	0

Table of Kaplan-Meier Estimators

Parametric Model Q-Q Plots

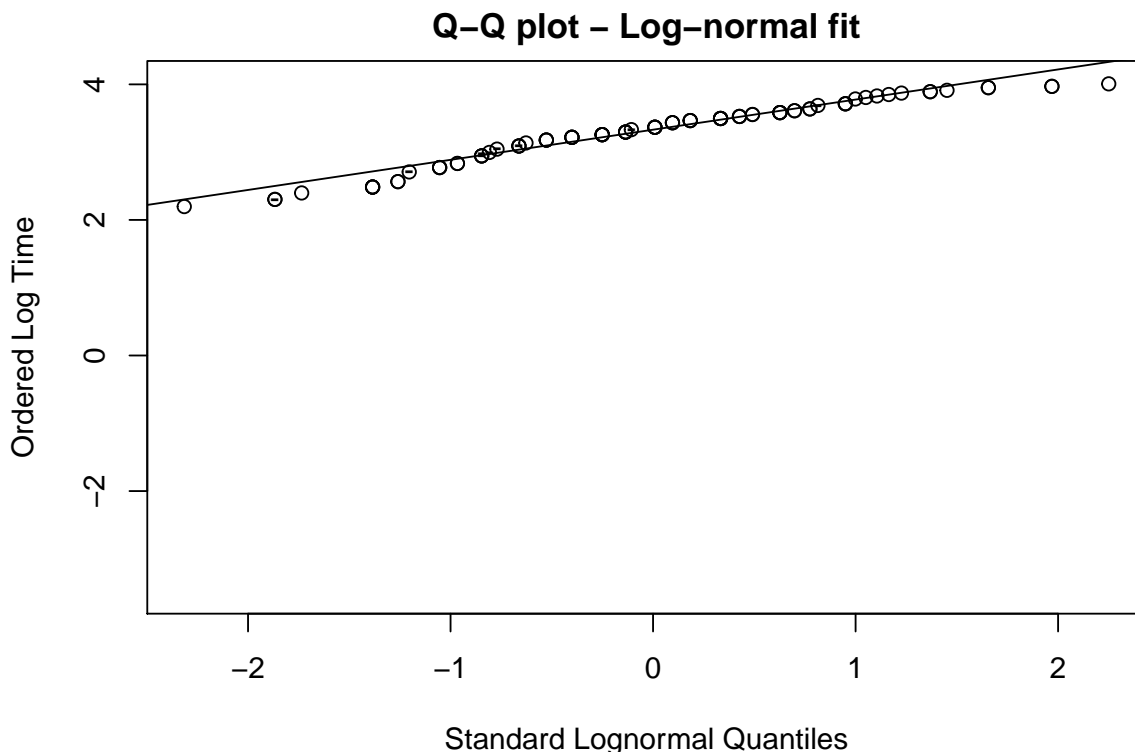
Q-Q plots for each of the three assessed parametric models are provided below. We see that the Weibull model has the best fit, with no large departures from the straight line.

```
[1] "Smallest observation is censored!" logtime sevrq 1 -3.5065579 -Inf 2 -1.3862944 -Inf 3 -1.3862944 -Inf
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10 -0.6931472 -Inf 11 -0.6931472 -Inf 12 -0.2876821 -Inf 13 -0.2876821 -Inf 14 -0.2876821 -Inf 15 -0.2876821
-Inf 16 -0.2876821 -Inf 17 -0.2876821 -Inf 18 0.0000000 -Inf 19 0.0000000 -Inf 20 0.0000000 -Inf 21 0.0000000
-Inf 22 0.0000000 -Inf 23 0.0000000 -Inf 24 0.2231436 -Inf 25 0.6931472 -Inf 26 0.6931472 -Inf 27 1.0986123
-Inf 28 1.0986123 -Inf 29 1.3862944 -Inf 30 1.6094379 -Inf 31 1.6094379 -Inf 32 1.9459101 -Inf 33 2.0149030
-Inf 34 2.1972246 -4.5695341 35 2.3025851 -3.4604317 36 2.3025851 -3.4604317 37 2.3025851 -3.4604317
38 2.3978953 -3.1646928 39 2.4849066 -2.4455451 40 2.4849066 -2.4455451 41 2.4849066 -2.4455451
42 2.4849066 -2.4455451 43 2.5649494 -2.2102941 44 2.5649494 -2.2102941 45 2.7080502 -2.1089574
46 2.7080502 -2.1089574 47 2.7725887 -1.8468574 48 2.7725887 -1.8468574 49 2.7725887 -1.8468574
50 2.8332133 -1.6997271 51 2.8332133 -1.6997271 52 2.9444390 -1.5075680 53 2.9444390 -1.5075680
54 2.9444390 -1.5075680 55 2.9444390 -1.5075680 56 2.9704145 -1.5075680 57 2.9957323 -1.4479419
58 2.9957323 -1.4479419 59 3.0445224 -1.3901426 60 3.0445224 -1.3901426 61 3.0910425 -1.2281207
62 3.0910425 -1.2281207 63 3.0910425 -1.2281207 64 3.0910425 -1.2281207 65 3.1354942 -1.1772988
66 3.1780538 -1.0346161 67 3.1780538 -1.0346161 68 3.1780538 -1.0346161 69 3.2188758 -0.8626217
70 3.2188758 -0.8626217 71 3.2188758 -0.8626217 72 3.2188758 -0.8626217 73 3.2580965 -0.6690299
74 3.2580965 -0.6690299 75 3.2580965 -0.6690299 76 3.2580965 -0.6690299 77 3.2580965 -0.6690299 78
3.2958369 -0.5264862 79 3.2958369 -0.5264862 80 3.2958369 -0.5264862 81 3.2958369 -0.5264862 82 3.3322045
-0.4921738 83 3.3322045 -0.4921738 84 3.3672958 -0.3563270 85 3.3672958 -0.3563270 86 3.3672958 -0.3563270
87 3.3672958 -0.3563270 88 3.4339872 -0.2580401 89 3.4339872 -0.2580401 90 3.4339872 -0.2580401 91
3.4657359 -0.1619005 92 3.4657359 -0.1619005 93 3.4657359 -0.1619005 94 3.4965076 -0.0043014 95 3.4965076
-0.0043014 96 3.4965076 -0.0043014 97 3.4965076 -0.0043014 98 3.4965076 -0.0043014 99 3.5263605 0.0899759
100 3.5263605 0.0899759 101 3.5263605 0.0899759 102 3.5553481 0.1532419 103 3.5553481 0.1532419 104
3.5835189 0.2821288 105 3.5835189 0.2821288 106 3.5835189 0.2821288 107 3.5835189 0.2821288 108
3.6109179 0.3484864 109 3.6109179 0.3484864 110 3.6375862 0.4167378 111 3.6375862 0.4167378 112
3.6888795 0.4517582 113 3.6888795 0.4517582 114 3.7135721 0.5682799 115 3.7135721 0.5682799 116
3.7135721 0.5682799 117 3.7841896 0.6094046 118 3.8066625 0.6520015 119 3.8286414 0.6963371 120
3.8501476 0.7427467 121 3.8712010 0.7916617 122 3.8918203 0.8995006 123 3.8918203 0.8995006 124
3.9120230 0.9603175 125 3.9512437 1.1045914 126 3.9512437 1.1045914 127 3.9702919 1.3113386 128
3.9702919 1.3113386 129 4.0073332 1.4825780 130 4.0430513 Inf
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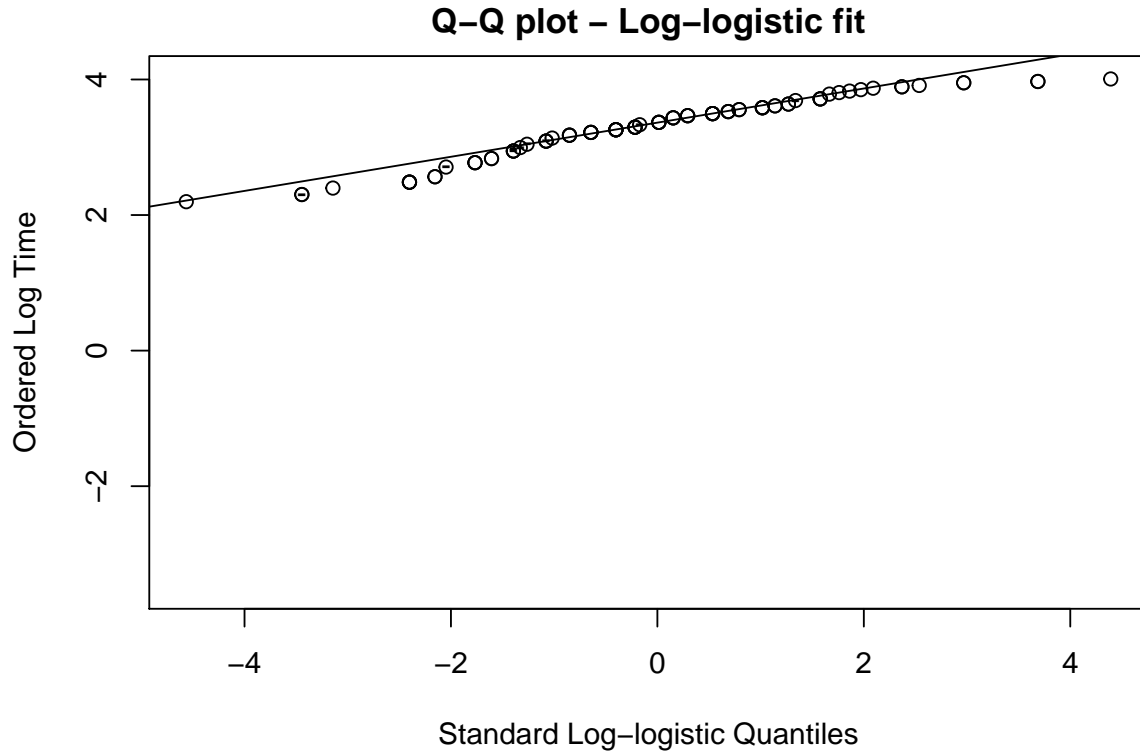
[1] “Q-Q plot for weibull done” [1] “Smallest observation is censored!” logtime sevf -3.5065579 -Inf 2 -1.3862944 -Inf 3 -1.3862944 -Inf 4 -1.3862944 -Inf 5 -1.3862944 -Inf 6 -0.6931472 -Inf 7 -0.6931472 -Inf 8 -0.6931472 -Inf 9 -0.6931472 -Inf 10 -0.6931472 -Inf 11 -0.6931472 -Inf 12 -0.2876821 -Inf 13 -0.2876821 -Inf 14 -0.2876821 -Inf 15 -0.2876821 -Inf 16 -0.2876821 -Inf 17 -0.2876821 -Inf 18 0.0000000 -Inf 19 0.0000000 -Inf 20 0.0000000 -Inf 21 0.0000000 -Inf 22 0.0000000 -Inf 23 0.0000000 -Inf 24 0.2231436 -Inf 25 0.6931472 -Inf 26 0.6931472 -Inf 27 1.0986123 -Inf 28 1.0986123 -Inf 29 1.3862944 -Inf 30 1.6094379 -Inf 31 1.6094379 -Inf 32 1.9459101 -Inf 33 2.0149030 -Inf 34 2.1972246 -2.314897235 35 2.3025851 -1.867328913 36 2.3025851 -1.867328913 37 2.3025851 -1.867328913 38 2.3978953 -1.735253400 39 2.4849066 -1.384985248 40 2.4849066 -1.384985248 41 2.4849066 -1.384985248 42 2.4849066 -1.384985248 43 2.5649494 -1.259811137 44 2.5649494 -1.259811137 45 2.7080502 -1.204030570 46 2.7080502 -1.204030570 47 2.7725887 -1.054087534 48 2.7725887 -1.054087534 49 2.7725887 -1.054087534 50 2.8332133 -0.966049199 51 2.8332133 -0.966049199 52 2.9444390 -0.846480688 53 2.9444390 -0.846480688 54 2.9444390 -0.846480688 55 2.9444390 -0.846480688 56 2.9704145 -0.846480688 57 2.9957323 -0.808256465 58 2.9957323 -0.808256465 59 3.0445224 -0.770670800 60 3.0445224 -0.770670800 61 3.0910425 -0.662391162 62 3.0910425 -0.662391162 63 3.0910425 -0.662391162 64 3.0910425 -0.662391162 65 3.1354942 -0.627501031 66 3.1780538 -0.527045857 67 3.1780538 -0.527045857 68 3.1780538 -0.527045857 69 3.2188758 -0.400751729 70 3.2188758 -0.400751729 71 3.2188758 -0.400751729 72 3.2188758 -0.400751729 73 3.2580965 -0.251206585 74 3.2580965 -0.251206585 75 3.2580965 -0.251206585 76 3.2580965 -0.251206585 77 3.2580965 -0.251206585 78 3.2958369 -0.135653042 79 3.2958369 -0.135653042 80 3.2958369 -0.135653042 81 3.2958369 -0.135653042 82 3.3322045 -0.107104766 83 3.3322045 -0.107104766 84 3.3672958 0.008862707 85 3.3672958 0.008862707 86 3.3672958 0.008862707 87 3.3672958 0.008862707 88 3.4339872 0.095831270 89 3.4339872 0.095831270 90 3.4339872 0.095831270 91 3.4657359 0.183532159 92 3.4657359 0.183532159 93 3.4657359 0.183532159 94 3.4965076 0.333279032 95 3.4965076 0.333279032 96 3.4965076 0.333279032 97 3.4965076 0.333279032 98 3.4965076 0.333279032 99 3.5263605 0.426629000 100 3.5263605 0.426629000 101 3.5263605 0.426629000 102 3.5553481 0.490942887 103 3.5553481 0.490942887 104 3.5835189 0.626325621 105 3.5835189 0.626325621 106 3.5835189 0.626325621 107 3.5835189 0.626325621 108 3.6109179 0.698413469 109 3.6109179 0.698413469 110 3.6375862 0.774329562 111 3.6375862 0.774329562 112 3.6888795 0.814000679 113 3.6888795 0.814000679 114 3.7135721 0.949639380 115 3.7135721 0.949639380 116 3.7135721 0.949639380 117 3.7841896 0.998894416 118 3.8066625 1.050700511 119 3.8286414 1.105491949 120 3.8501476

1.163820405 121 3.8712010 1.226402832 122 3.8918203 1.368525132 123 3.8918203 1.368525132 124 3.9120230
1.451293751 125 3.9512437 1.655620380 126 3.9512437 1.655620380 127 3.9702919 1.969468118 128 3.9702919
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[1] “Q-Q plot for lognormal done” [1] “Smallest observation is censored!” logtime sevf 1 -3.5065579 -Inf 2
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-Inf 26 0.6931472 -Inf 27 1.0986123 -Inf 28 1.0986123 -Inf 29 1.3862944 -Inf 30 1.6094379 -Inf 31 1.6094379
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-1.39480156 53 2.9444390 -1.39480156 54 2.9444390 -1.39480156 55 2.9444390 -1.39480156 56 2.9704145
-1.39480156 57 2.9957323 -1.32811414 58 2.9957323 -1.32811414 59 3.0445224 -1.26303986 60 3.0445224
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0.53452625 97 3.4965076 0.53452625 98 3.4965076 0.53452625 99 3.5263605 0.68644312 100 3.5263605

0.68644312	101	3.5263605	0.68644312	102	3.5553481	0.79202783	103	3.5553481	0.79202783	104	3.5835189
1.01731522	105	3.5835189	1.01731522	106	3.5835189	1.01731522	107	3.5835189	1.01731522	108	3.6109179
1.13924325	109	3.6109179	1.13924325	110	3.6375862	1.26935338	111	3.6375862	1.26935338	112	3.6888795
1.33810246	113	3.6888795	1.33810246	114	3.7135721	1.57751462	115	3.7135721	1.57751462	116	3.7135721
1.57751462	117	3.7841896	1.66626400	118	3.8066625	1.76073683	119	3.8286414	1.86197307	120	3.8501476
1.97130859	121	3.8712010	2.09050046	122	3.8918203	2.36891658	123	3.8918203	2.36891658	124	3.9120230
2.53634755	125	3.9512437	2.96785557	126	3.9512437	2.96785557	127	3.9702919	3.68638461	128	3.9702919
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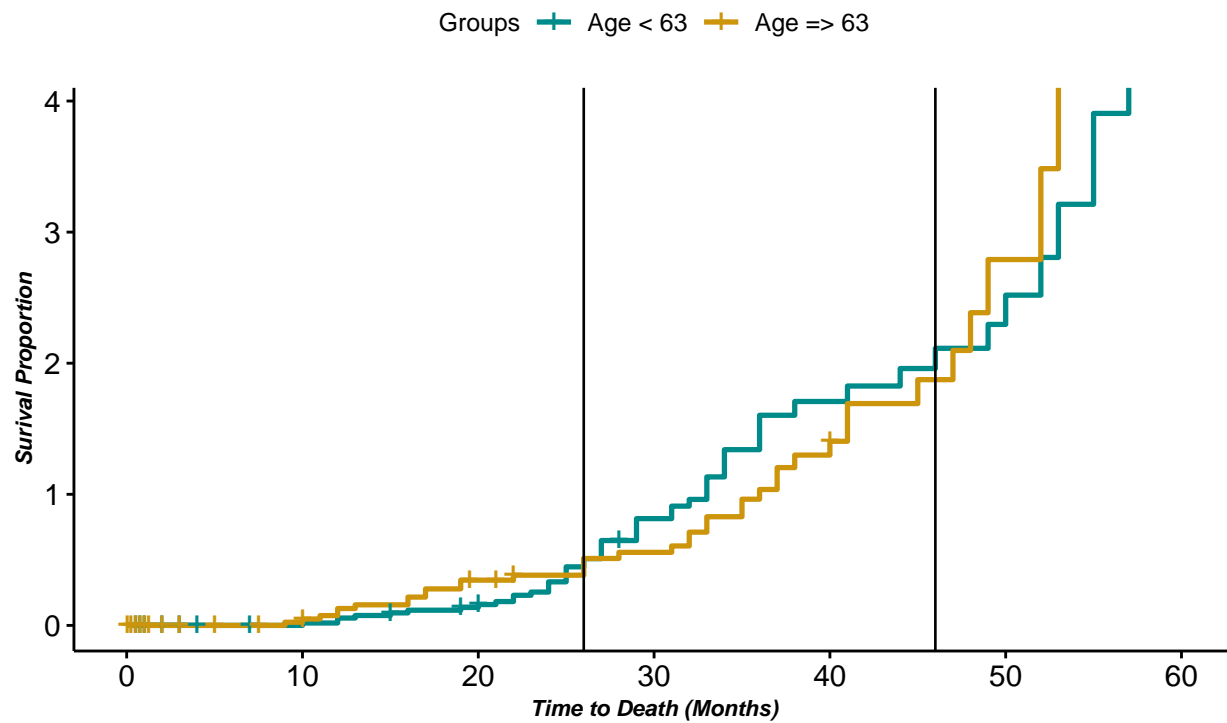


[1] “Q-Q plot for loglogistic done”

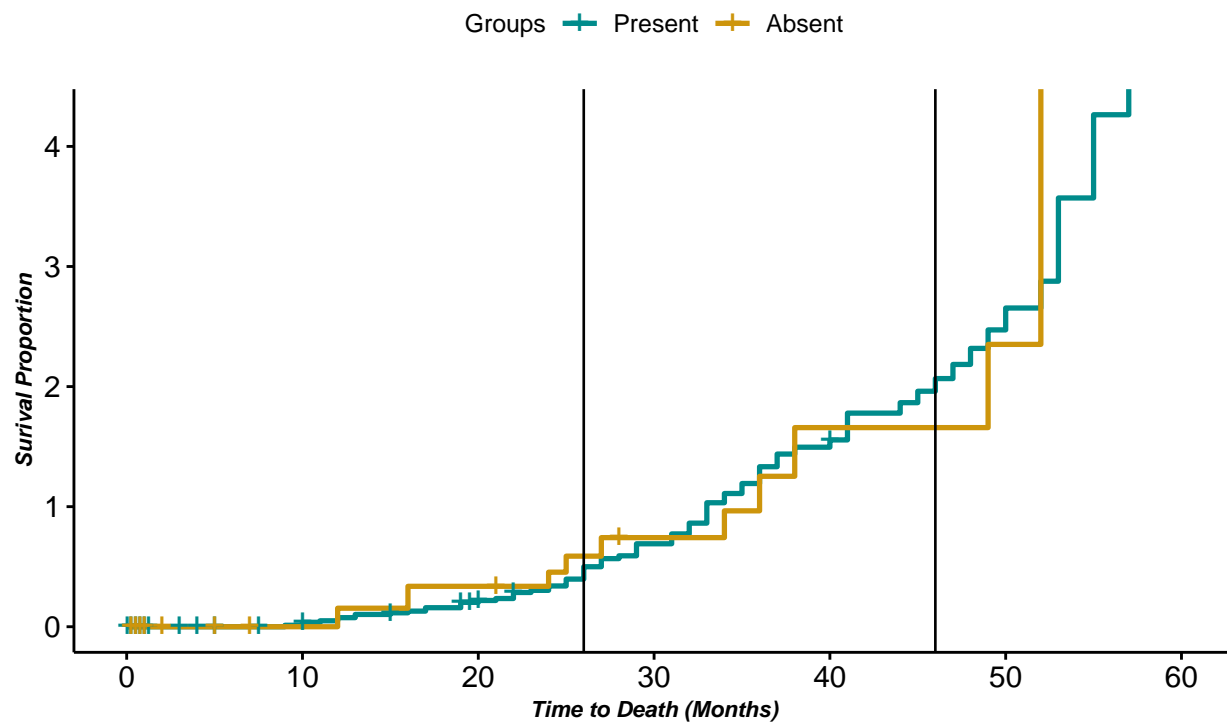
Hazard Plots for Regression Covariates - Regression Time Interval Identification

Hazard curves stratified by group for each regression covariate are provided below, with the three time interval for the cox PH modeling marked on each plot. It can be observed that these time intervals approximately correspond to hazard crossover behavior for the covariates, with the exception of pericardial infusion.

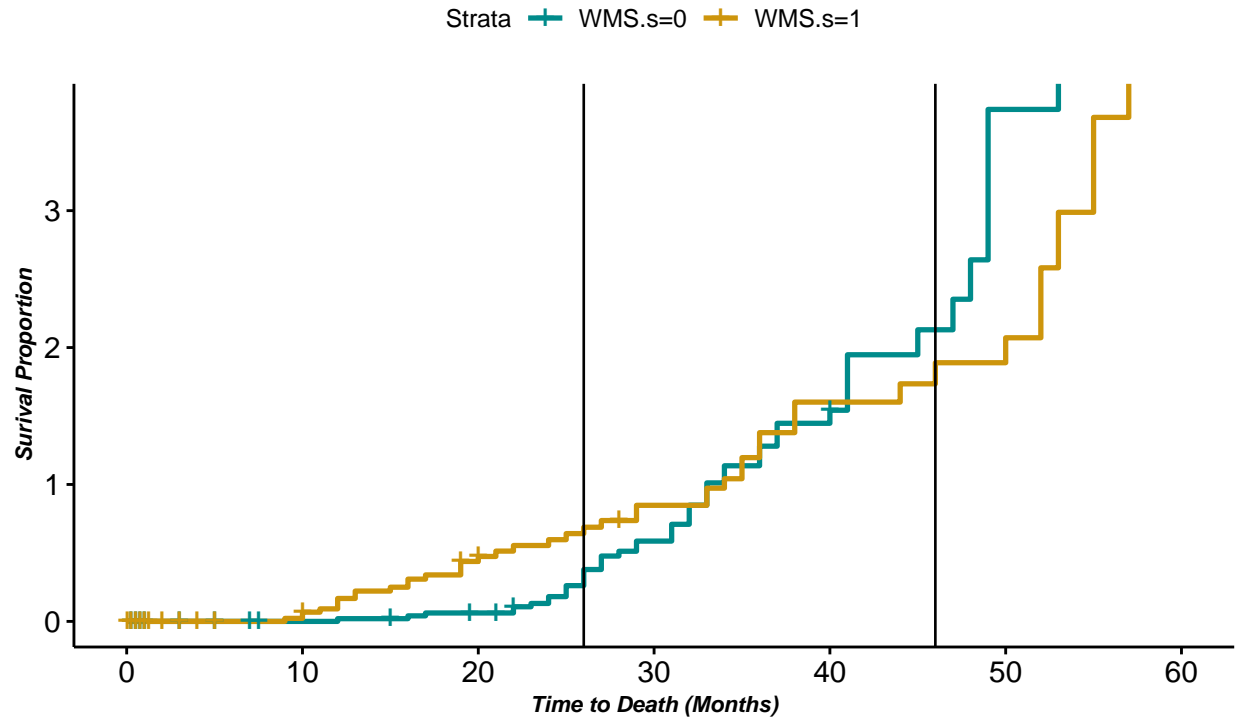
Hazard, Stratified by Age Group



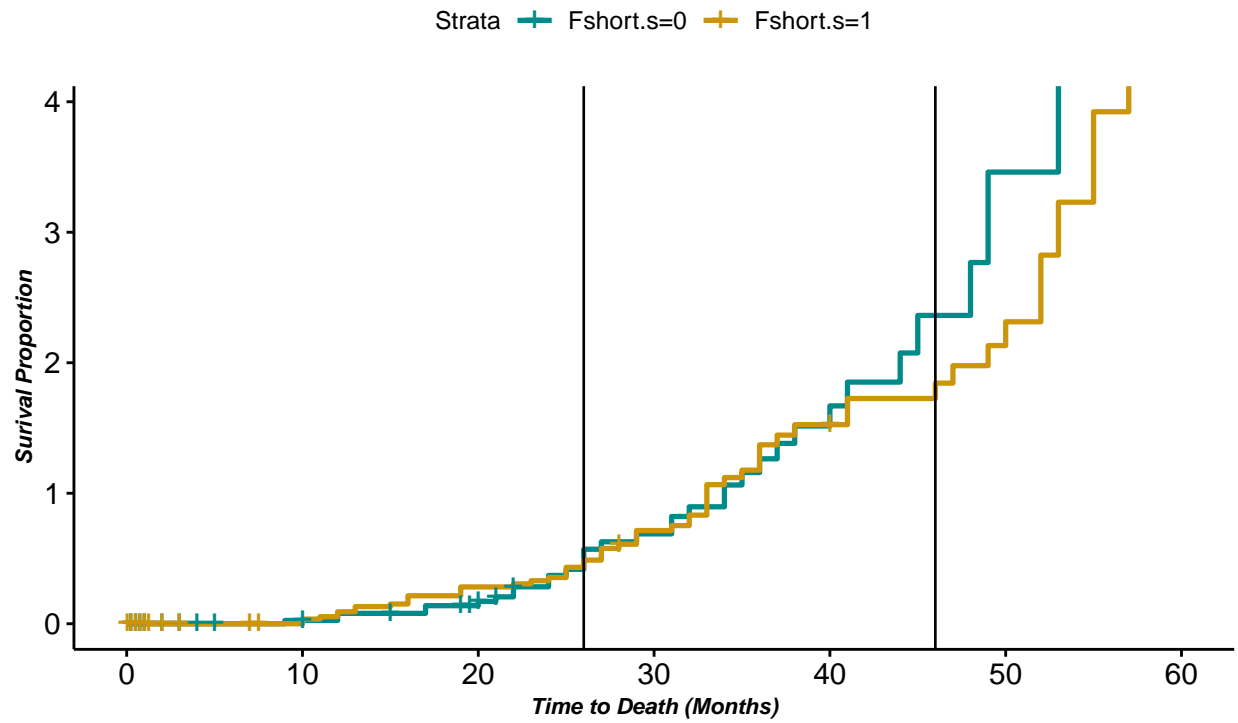
Hazard, Stratified by Pericardial Effusion Presence



Hazard, Stratified by Wall Motion Index



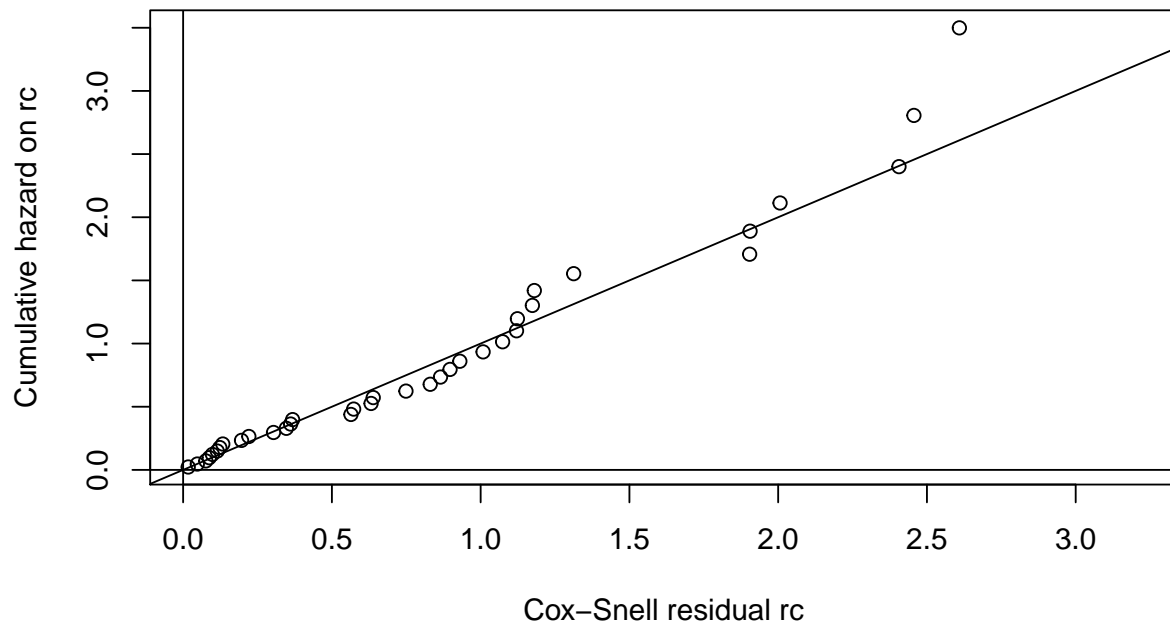
Hazard, Stratified by Fractional Shortening



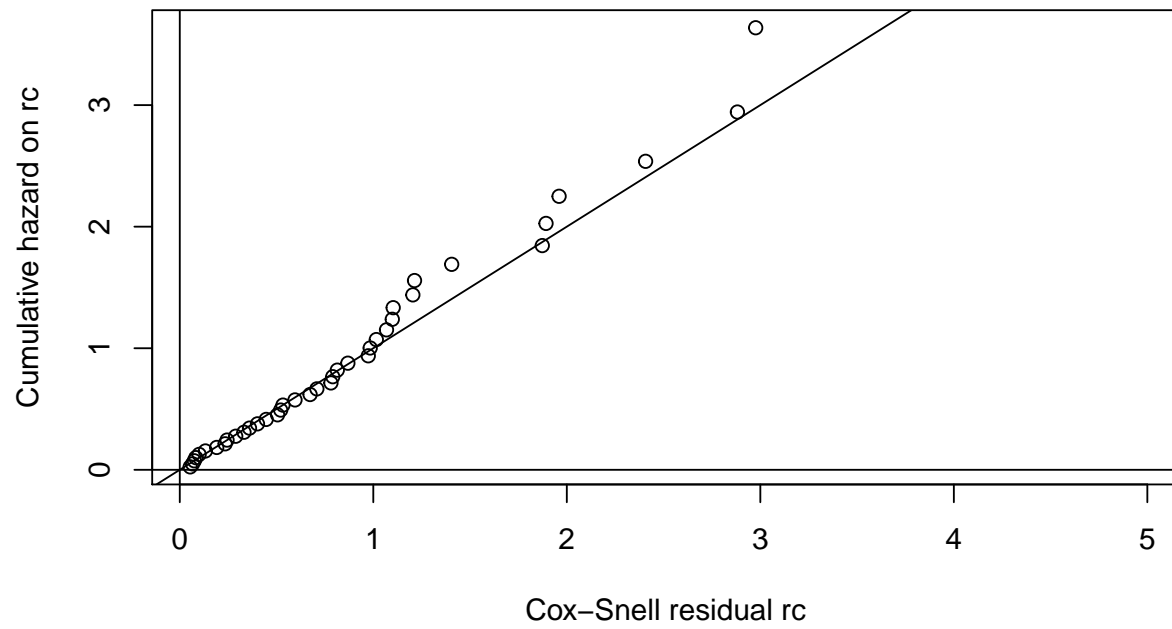
Regression Model Diagnostics

Cox-Snell residual plot for assessment of overall model fit and dfbeta residual plots for evaluation of constant coefficients are presented below for each of the three models. It can be seen that the model fits the data well overall and meets the constant coefficient assumption for Models 1 and 2, given that the Cox-Snell residuals fall closely along the straight line and that Schoenfeld residuals are symmetric about 0 with large p-values. However, these assumptions are not met for Model 3, which is not unexpected due to the small sample size.

Model 1 – Cox–Snell residual model fit evaluation

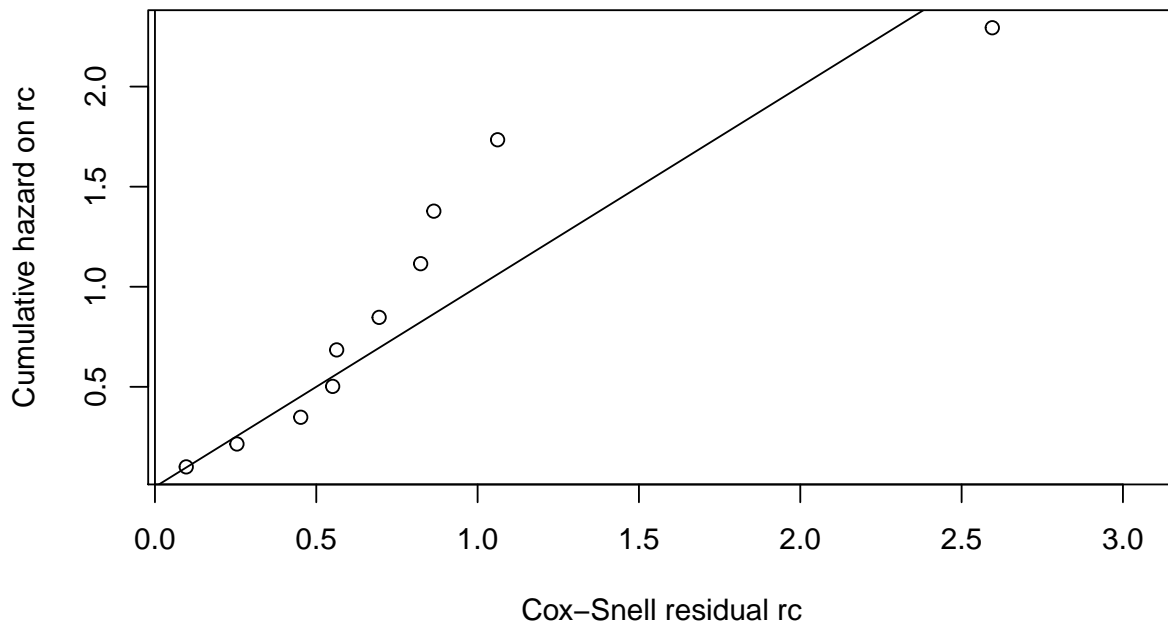


Model 2 – Cox–Snell residual model fit evaluation



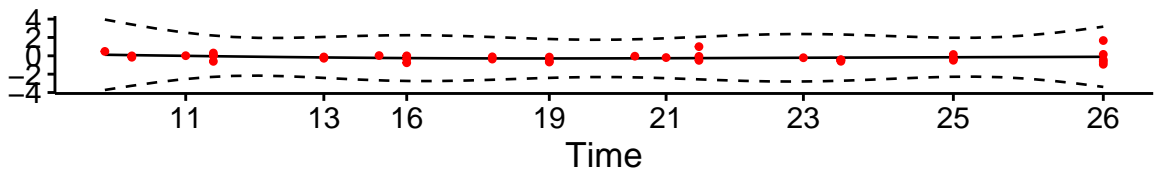
```
## Warning in status - mod3$residuals: longer object length is not a multiple of  
## shorter object length
```

Model 3 – Cox–Snell residual model fit evaluation

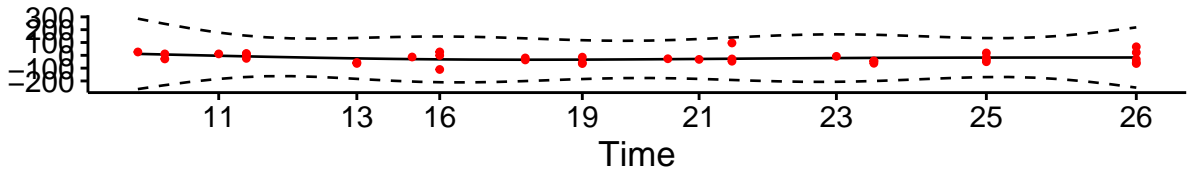


Global Schoenfeld Test p: 0.8463

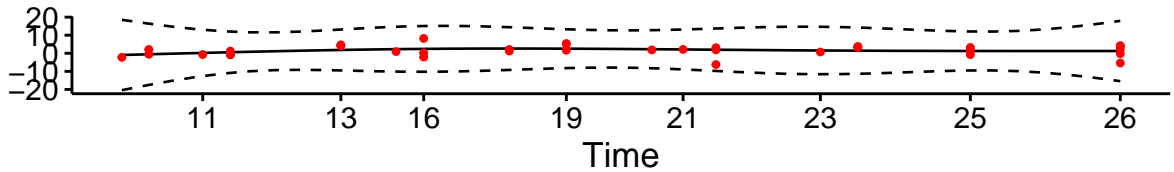
Schoenfeld Residuals – Const Coeff Evaluation for Model 1



Schoenfeld Residuals – Const Coeff Evaluation for Model 1

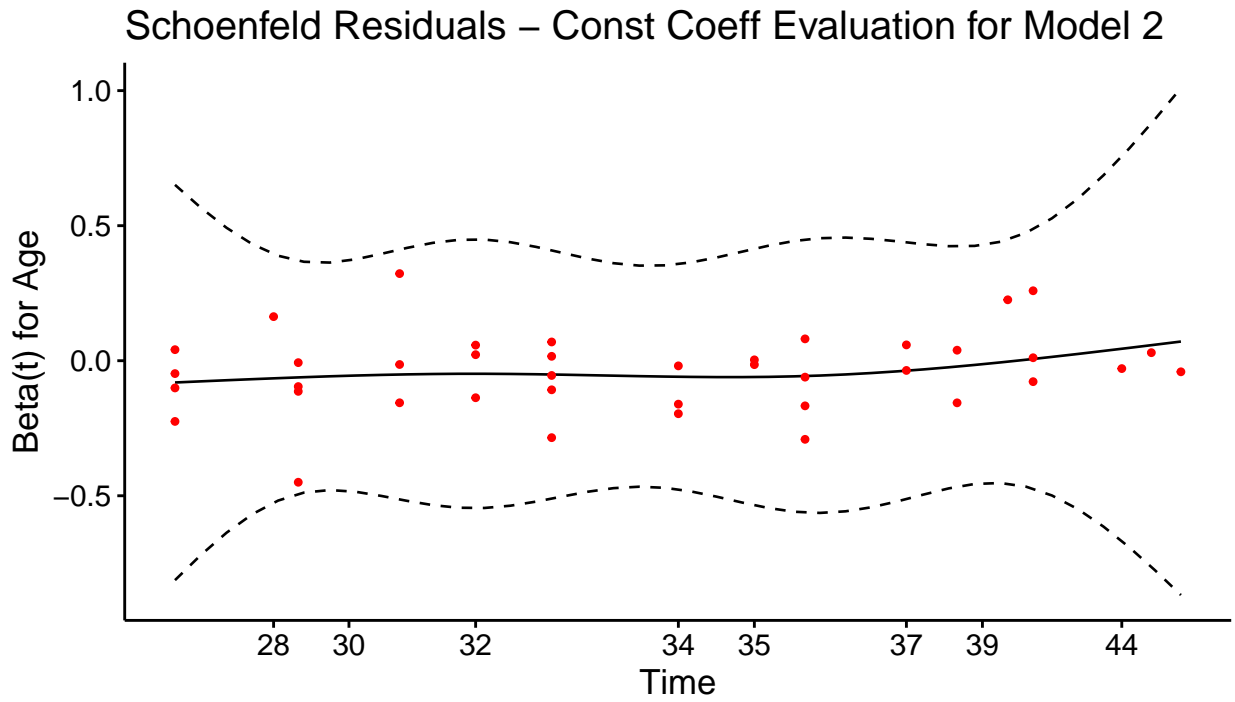


Schoenfeld Residuals – Const Coeff Evaluation for Model 1

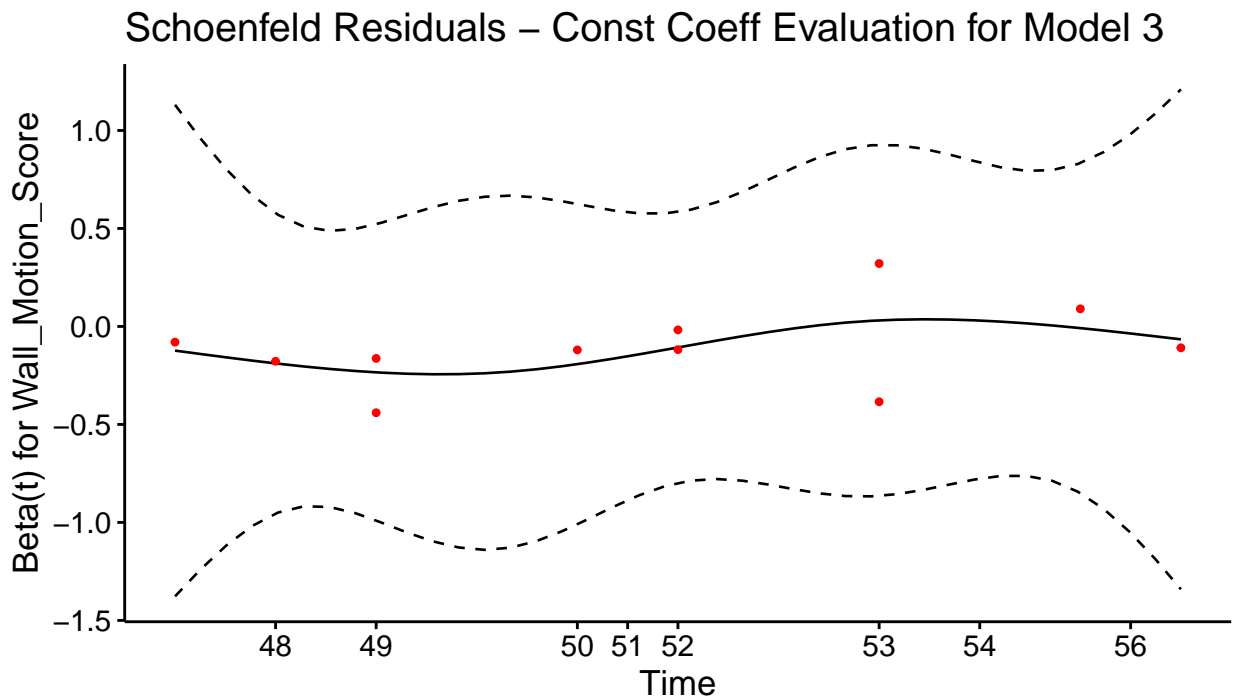


lotion_Score:Fraction_Bracket_Short_Score_Motion_Score

Global Schoenfeld Test p: 0.1363



Global Schoenfeld Test p: 0.2132



R Code

```
knitr::opts_chunk$set(echo = TRUE)
knitr::opts_chunk$set(fig.height=4.5, fig.width=7)

library(readxl)
library(knitr)
library(tidyverse)
library(dplyr)
library(kableExtra)
library(survival)
library(survminer)
library(ggplot2)
library(VIM)
library(missForest)
library(ggplot2)
library(ggpubr)
library(MASS)
library(SurvCorr)

df = data.frame(read_excel("df.xlsx"))
df[df=="?"] = " "

df.new = data.frame(read_excel("df.new.xlsx"))

s.df = Surv(df.new$Survival, df.new$Status)

Indicator = c("0", "1")
Age = c("< 63 Years", "\u2265 63 Years")
Effusion = c("Fluid is absent", "Fluid is present")
WMS = c("< 11", "\u2265 11")
FS = c("< 0.2", "\u2265 0.2")

groupings = data.frame(Indicator, Age, Effusion, WMS, FS)

kable(groupings, caption="Stratification Groupings", align="c") %>%
  kable_styling(position = "center", latex_options="hold_position")

missing.data = aggr(df) #visualize the missing information

set.seed(7522)
df.i = missForest(df, maxiter = 30, ntree = 1000)

round_df <- function(x, digits) {
  # round all numeric variables
  # x: data frame
  # digits: number of digits to round
  numeric_columns <- sapply(x, mode) == 'numeric'
  x[numeric_columns] <- round(x[numeric_columns], digits)
  x}

df.impute = round_df(df.i$ximp, 2) #imputed values table
```

```

df.new = df.impute[,c(-5,-12)] #remove incomplete strata from original data
Age.s = ifelse(df.impute$Age < 63,0,1) #new age strata based on imputed data #new age strata based on i
WMS.s = ifelse(df.impute$WMS < 14,0,1) #new WMS strata based on imputed data
Fshort.s = ifelse(df.impute$F.Shortening < 0.2,0,1) #new fshort strata based on imputed data
LVDD.s = ifelse(df.impute$LVDD < 4.75,0,1) #new lvdd strata based on imputed data
EPSS.s = ifelse(df.impute$EPSS < 11.1,0,1) #new epss strata based on imputed data

df.new$Age.s = Age.s
df.new$WMS.s = WMS.s
df.new$F.Short.s = Fshort.s
df.new$LVDD.s = LVDD.s
df.new$EPSS.s = EPSS.s

km.plots = list()

km.all = survfit(s.df~1,type="kaplan-meier", data=df.new)
km.plots[[1]] = ggsurvplot(km.all,
                           palette = "#2E9FDF",
                           conf.int = TRUE,
                           title="Post-Myocardial Infarction Survival",
                           font.title=c(14,"bold.italic"),
                           font.subtitle = c(10,"italic"),
                           font.x = c(9, "bold.italic"),
                           font.y = c(9, "bold.italic"),
                           ylab="Survival Proportion",
                           xlab="Time to Death (Months)",
                           surv.median.line = "hv",
                           legend.title = "Groups",
                           legend.labs = "All")

km.age = survfit(s.df~Age.s, type="kaplan-meier", data = df.new)
km.plots[[2]] = ggsurvplot(km.age,
                           palette = c("darkcyan","darkgoldenrod3","darkorange3"),
                           subtitle="Survival, Stratified by Age Group",
                           font.subtitle = c(10,"italic"),
                           font.x = c(9, "bold.italic"),
                           font.y = c(9, "bold.italic"),
                           ylab="Survival Proportion",
                           xlab="Time to Death (Months)",
                           surv.median.line = "hv",
                           legend.title = "Groups",
                           legend.labs = c("< 63 Years","\u2265 63 Years"))

km.effusion = survfit(s.df~P.Effusion, type="kaplan-meier", data = df.new)
km.plots[[3]] = ggsurvplot(km.effusion,
                           palette = c("darkcyan","darkgoldenrod3"),
                           subtitle="Survival, Stratified by Presence of Pericardial Effusion",
                           font.subtitle = c(10,"italic"),
                           font.x = c(9, "bold.italic"),
                           font.y = c(9, "bold.italic"),
                           ylab="Survival Proportion",

```

```

        xlab="Time to Death (Months)",
        surv.median.line = "hv",
        legend.title = "Groups",
        legend.labs = c("Present","Absent"))

km.wms = survfit(s.df~WMS.s, type="kaplan-meier", data = df.new)
km.plots[[4]] = ggsurvplot(km.wms,
        palette = c("darkcyan","darkgoldenrod3","darkorange3"),
        subtitle="Survival, Stratified by Wall Motion Score",
        font.subtitle = c(10,"italic"),
        font.x = c(9, "bold.italic"),
        font.y = c(9, "bold.italic"),
        ylab="Survival Proportion",
        xlab="Time to Death (Months)",
        surv.median.line = "hv",
        legend.title = "Groups",
        legend.labs = c("< 14","\u2265 14"))

km.fshort = survfit(s.df~Fshort.s, type="kaplan-meier", data = df.new)
km.plots[[5]] = ggsurvplot(km.fshort,
        palette = c("darkcyan","darkgoldenrod3"),
        subtitle="Survival, Stratified by Fractal Shortening",
        font.subtitle = c(10,"italic"),
        font.x = c(9, "bold.italic"),
        font.y = c(9, "bold.italic"),
        ylab="Survival Proportion",
        xlab="Time to Death (Months)",
        surv.median.line = "hv",
        legend.title = "Groups",
        legend.labs = c("< 0.2","\u2265 0.2"))

arrange_ggsurvplots(km.plots, print=TRUE, ncol=2, nrow=3)

ks1 = data.frame(t(summary(km.all)$table))
ks2 = data.frame(summary(km.age)$table)
ks3 = data.frame(summary(km.effusion)$table)
ks4 = data.frame(summary(km.wms)$table)
ks5 = data.frame(summary(km.fshort)$table)

ksall = rbind(ks1, ks2, ks3, ks4,ks5)
km.as = ksall[,c(1,4,5,7,8,9)]
colnames(km.as) = c("Records","Events","Mean","Median","Median 0.95 LCL","Median 0.95 UCL")
rownames(km.as) = c("All","Age:<55", "Age: 55-65", "P.Eff: Absent","P.Eff: Present", "WMS: < 14", "WMS:
>= 14")

kable(km.as, caption="Kaplan-Meier Results",align="c", digits=2) %>%
  kable_styling(position = "center", latex_options="hold_position")
haz.plots = list()

haz.plots[[1]] = ggsurvplot(km.all,
        fun = "cumhaz",
        palette = "#2E9FDF",

```

```

conf.int = TRUE,
title="Post-Myocardial Infarction Hazard",
font.title=c(14,"bold.italic"),
font.subtitle = c(10,"italic"),
font.x = c(9, "bold.italic"),
font.y = c(9, "bold.italic"),
ylab="Survival Proportion",
xlab="Time to Death (Months)",
legend.title = "Groups",
legend.labs = "All")

haz.plots[[2]] = ggsurvplot(km.age,
  fun = "cumhaz",
  palette = c("darkcyan","darkgoldenrod3"),
  subtitle="Hazard, Stratified by Age Group",
  font.subtitle = c(10,"italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)",
  legend.title = "Groups",
  legend.labs = c("< 55 Year","55 - 65 Years"))

haz.plots[[3]] = ggsurvplot(km.effusion,
  fun = "cumhaz",
  palette = c("darkcyan","darkgoldenrod3"),
  subtitle="Hazard, Stratified by Pericardial Effusion Presence",
  font.subtitle = c(10,"italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)",
  legend.title = "Groups",
  legend.labs = c("Present","Absent"))

haz.plots[[4]] = ggsurvplot(km.wms,
  fun = "cumhaz",
  palette = c("darkcyan","darkgoldenrod3"),
  subtitle="Hazard, Stratified by Wall Motion Index",
  font.subtitle = c(10,"italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)",
  legend.title = "Groups",
  legend.labs = c("< 14","\u2265 14"))

haz.plots[[5]] = ggsurvplot(km.fshort,
  fun = "cumhaz",
  palette = c("darkcyan","darkgoldenrod3"),
  subtitle="Survival, Stratified by Fractal Shortening",
  font.subtitle = c(10,"italic"),
  font.x = c(9, "bold.italic"),

```

```

        font.y = c(9, "bold.italic"),
        ylab="Survival Proportion",
        xlab="Time to Death (Months)",
        legend.title = "Groups",
        legend.labs = c("< 0.2", "\u2265 0.2"))

arrange_ggsurvplots(haz.plots, print=TRUE, ncol=2, nrow=3)

months=df.new$Survival
status=df.new$Status
months.u=months[status == 1]
months.u = sort(months.u)
nu = length(months.u)

#Weibull model plot

weib.fit=survreg(Surv(months,status)~1,dist="weib")
alphahat=1/weib.fit$scale
scalehat=exp(weib.fit$coefficients)
Shat.w = 1- pweibull(months.u,alphahat,scalehat)
plot(km.all,conf.int=F,xlab="time until death (in months)",
     ylab="proportion survived",
     main= "Survival Curves - Weibull and Kaplan-Meier",
     lwd=2,
     col = "darkcyan")
lines(months.u, Shat.w, col="darkgoldenrod3",lwd=2)
legend(40, 0.8, legend=c("Kaplan-Meier", "Weibull"),
      col=c("darkcyan","darkgoldenrod3"), lty=1:1, cex=0.8,lwd=2)
abline(h=0)

#log-normal model plot

lognorm.fit=survreg(Surv(months,status)~1,dist="lognormal")
muhat=lognorm.fit$coefficients
sigmahat=lognorm.fit$scale
Shat.l = 1- pnorm(log(months.u),muhat,sigmahat)
plot(km.all,conf.int=F,xlab="time until death (in months)",
     ylab="proportion survived",
     main="Survival Curves - Log-normal and Kaplan-Meier",
     lwd=2,
     col="darkcyan")
lines(months.u, Shat.l, col="darkgoldenrod3",lwd=2)
legend(40, 0.8, legend=c("Kaplain", "Weibull"), lwd=2,
      col=c("darkcyan","darkgoldenrod3"), lty=1:1, cex=0.8)
abline(h=0)

#log-logistic model plot

loglog.fit=survreg(Surv(months,status)~1,dist="loglogistic")
muhat=loglog.fit$coefficients
sigmahat=loglog.fit$scale
Shat.ll = 1- plogis(log(months.u),muhat,sigmahat)

```

```

plot(km.all,conf.int=F,xlab="time until death (in months)",
     ylab="proportion survived",
     main="Survival Curves - Log-logistic and Kaplan-Meier",
     lwd=2,
     col="darkcyan")
lines(months.u, Shat.ll, col="darkgoldenrod3",lwd=2,)
legend(40, 0.8, legend=c("Kaplan", "Weibull"), lwd=2,
      col=c("darkcyan","darkgoldenrod3"), lty=1:1, cex=0.8)
abline(h=0)

param.est = data.frame(read_excel("param.est.xlsx"))
kable(param.est,
      col.names = c("Model","Quantile", "Point Estimate", "95% LCL", "95% UCL", "Interval Length")
      )

library("gtsummary")
LL=0.0
UL=26.0

#Subset the data based on the time region

months=df.new$Survival[df.new$Survival>=LL & df.new$Survival<=UL]
status=df.new$Status[df.new$Survival>=LL & df.new$Survival<=UL]
Age=df.new$Age[df.new$Survival>=LL & df.new$Survival<=UL]
Pericardial_Effusion=df.new$P.Effusion[df.new$Survival>=LL & df.new$Survival<=UL]
Wall_Motion_Score=df.new$WMS[df.new$Survival>=LL & df.new$Survival<=UL]
Fractional_Shortening=df.new$F.Shortening[df.new$Survival>=LL & df.new$Survival<=UL]

#Create initial model fit

cph.fit1=coxph(Surv(months,status)~Age+Pericardial_Effusion+Wall_Motion_Score+Fractional_Shortening,x=T)

#Reduce with StepAIC procedure

cph.fit2=stepAIC(cph.fit1,~.^2,direction="both",trace=FALSE)
mod1=cph.fit2

t1 <-
  mod1 %>%
  tbl_regression(exponentiate = TRUE)%>%
  add_nevent()

LL=26
UL=46

#Subset the data based on the time region

months=df.new$Survival[df.new$Survival>LL & df.new$Survival<=UL]
status=df.new$Status[df.new$Survival>LL & df.new$Survival<=UL]
Age=df.new$Age[df.new$Survival>LL & df.new$Survival<=UL]
Pericardial_Effusion=df.new$P.Effusion[df.new$Survival>LL & df.new$Survival<=UL]

```

```

Wall_Motion_Score=df.new$WMS[df.new$Survival>LL & df.new$Survival<=UL]
Fractional_Shortening=df.new$F.Shortening[df.new$Survival>LL & df.new$Survival<=UL]

#Create initial model fit

cph.fit1=coxph(Surv(months,status)~Age+Pericardial_Effusion+Wall_Motion_Score+Fractional_Shortening,x=T)

#Reduce with StepAIC procedure

cph.fit2=stepAIC(cph.fit1,~.^2,direction="both",trace=FALSE)
mod2=cph.fit2

t2 <-
  mod2 %>%
  tbl_regression(exponentiate = TRUE) %>%
  add_nevent()

#Subset the data based on the time region

months=df.new$Survival[df.new$Survival>UL]
status=df.new$Status[df.new$Survival>UL]
Age=df.new$Age[df.new$Survival>UL]
Pericardial_Effusion=df.new$P.Effusion[df.new$Survival>UL]
Wall_Motion_Score=df.new$WMS[df.new$Survival>UL]
Fractional_Shortening=df.new$F.Shortening[df.new$Survival>UL]

#Create initial model fit

cph.fit1=coxph(Surv(months,status)~Age+Pericardial_Effusion+Wall_Motion_Score+Fractional_Shortening,x=T)

#Reduce with StepAIC procedure

cph.fit2=stepAIC(cph.fit1,~.^2,direction="backward",trace=FALSE)
mod3=cph.fit2

t3 <-
  mod3 %>%
  tbl_regression(exponentiate = TRUE)%>%
  add_nevent()

# merge tables

tbl_merge(
  tbls = list(t1, t2, t3),
  tab_spanner = c("Model 1", "Model 2", "Model 3")
)

df.sum = data.frame(read_excel("df.sum.xlsx"))

kable(df.sum, "latex",
      booktabs = TRUE,
      longtable = TRUE,
      linesep = "\\addlinespace",

```



```

        caption = "Summary of Dataset Covariates") %>%
kable_styling(latex_options = c("hold_position", "repeat_header"),
              full_width = TRUE)

kable(df, "latex",
      booktabs = TRUE,
      longtable = TRUE,
      caption = "Original Dataset") %>%
kable_styling(latex_options = c("hold_position", "repeat_header"),
              full_width = TRUE)
kable(df.new, "latex",
      booktabs = TRUE,
      longtable = TRUE,
      caption = "Imputed Dataset") %>%
kable_styling(latex_options = c("hold_position", "repeat_header"),
              full_width = TRUE)

km.sum = data.frame(km.all$n.risk, km.all$n.event, km.all$n.censor, km.all$surv, km.all$std.err, km.all$

kable(km.sum,
      caption="Kaplan-Meier Estimate Summary",
      col.names = c("Ni", "Di", "Ci", "Survival", "Std. Err", "95% LCL", "95% UCL")) %>%
kable_styling(latex_options = "hold_position")

#Q-Q Plots - Weibull, Log-lognormal, Log-logistic
#qq.surv function:
#Author: Jong Sung Kim, Date: 8/10/2004
# Edited by D. Leif Rustvold, Date: 6/7/2006

qq.surv <- function(time, status, pdgy = 0, distribution = "weibull", scale = 0, adjpb =
                    0.025, ...)
{
  ## Purpose: qqplot for distributions that satisfy a log-linear form
  ## for one sample. It fits each sample with own intercept and slope
  ## (location and scale).
  ##-----
  ## Arguments
  ## =====
  ## time:   observed time
  ## status: censoring indicator
  ##
  ## Options
  ## =====
  ## pdgy:   Flag to generate for pedagogical purposes additional lines
  ##          incorporating the effect of how we treat censored
  ##          observations on the MLE's (equivalently estimated line).
  ##          pdgy=0 is the default, for no additional lines.
  ##          pdgy=1 generates additional lines.
  ## distribution: Distribution for fit.
  ##          May take values "weibull", "loglogistic", or "lognormal".
  ##          The default is "weibull" distribution (exponential model with

```

```

##          scale=1). Enter "loglogistic" to fit loglogistic distribution;
##          Enter "lognormal" to fit lognormal distribution.
## scale: Scale parameter. scale=0 is the default. This estimates
##          the scale. With distribution "weibull", scale=1 fits the
##          exponential model.
## adjpb: Replaces the zero survival probability when the max is exact.
##          Or when the min is censored, it replaces the survival
##          probability by 1 - adjpb. Default is 0.025.
##          This has nothing to do with the MLE line, but is solely for
##          plotting the point on the graph.
##-----
## Author: Jong Sung Kim, Date: 8/10/2004
## Edited by D. Leif Rustvold, Date: 6/7/2006
d <- data.frame(time, status)
# data frame
d <- na.exclude(d)
# Missing observations excluded
d <- d[order(d$time), ]
# Rearranging the observed times into a nondecreasing order
# Unordered times sometimes mess up QQ-plots.
time <- d$time
# sorted time
status <- d$status
# status corresponding to sorted time
data <- Surv(time, status)
# Surv object
t.c <- class(data)
if(!is.null(t.c) && t.c == "Surv")
  data <- list(data)
t.s <- summary(survfit(Surv(time, status)~1, type = "kaplan-meier",
                      na.action = na.exclude))

survp <- t.s$surv
survtime <- t.s$time
rare <- F
# rare = T indicates that the smallest observation is censored
if(time[1] < survtime[1]) {
  print("Smallest observation is censored!")
  survp <- c(1 - adjpb, survp)
  survtime <- c(time[1], survtime)
  rare <- T
}
#####
#####
xlabs <- ifelse(distribution == "weibull",
               "Standard Extreme Value Quantiles", ifelse(distribution ==
                                                         "loglogistic", "Standard Log-logistic Qu
                                                         distribution == "lognormal", "Standard
                                                         "")))

if(pdgy == 1) {
  #####
  t.s.exactall <- summary(survfit(Surv(time, status >= 0)~1, type
                                = "kaplan-meier", na.action = na.exclude))
  exactall.surv <- t.s.exactall$surv

```

```

exactall.survtime <- t.s.exactall$time
exactall.length <- length(exactall.survtime)
exactall.survp[exactall.length] <- adjpb
t.ss.exactall <- exactall.survp
#quant.exactall <- qweibull(1 - t.ss.exactall, 1)
quant.exactall <- switch(distribution,
                        weibull = qweibull(1 - t.ss.exactall, 1),
                        lognormal = qlnorm(1 - t.ss.exactall),
                        loglogistic = exp(logis((1 - t.ss.exactall))))
exactall.sevq <- log(quant.exactall)
# standard extreme value quantile
exactall.logtime <- log(exactall.survtime)
print(data.frame(exactall.logtime, exactall.sevq))
#####
ok <- status == 1
t.s.exact <- summary(survfit(Surv(time[ok], status[ok])~1, type
                           = "kaplan-meier", na.action = na.exclude))
exact.survp <- t.s.exact$surv
exact.survtime <- t.s.exact$time
exact.length <- length(exact.survtime)
exact.survp[exact.length] <- adjpb
t.ss.exact <- exact.survp
#quant.exact <- qweibull(1 - t.ss.exact, 1)
quant.exact <- switch(distribution,
                     weibull = qweibull(1 - t.ss.exact, 1),
                     lognormal = qlnorm(1 - t.ss.exact),
                     loglogistic = exp(qlogis(1 - t.ss.exact)))
exact.sevq <- log(quant.exact)
# standard extreme value quantile
exact.logtime <- log(exact.survtime)
print(data.frame(exact.logtime, exact.sevq))
#####
n <- length(time)
t.ss <- rep(0, n)
for(i in 1:n) {
  # This loop assigns probabilities to censored time points,
  # and takes care of tied observations as well
  idx <- time[i] >= survtime
  t.ss[i] <- min(survp[idx], na.rm = T)
}
#sevq <- log(qweibull(1 - t.ss, 1))
sevq <- log(switch(distribution,
                  weibull = qweibull(1 - t.ss, 1),
                  lognormal = qlnorm(1 - t.ss),
                  loglogistic = exp(qlogis(1 - t.ss))))
# standard extreme value quantile
logtime <- log(time)
print(data.frame(logtime, sevq))
##### Multiple Plot starts #####
xrange <- range(c(exactall.sevq, exact.sevq, sevq))
yrange <- range(c(exactall.logtime, exact.logtime, logtime))
par(mar = c(5, 5, 2, 2))
plot(sevq, logtime, type = "n", lty = 1, xlim = xrange, ylim

```

```

    = yrange, xlab = xlabs, ylab = "Ordered Log Time",
    ...)
points(sevq[ok], logtime[ok], pch = 1)
# exact points portion
points(sevq[!ok], logtime[!ok], pch = "\255", font = 8)
# censored points portion
points(exactall.sevq, exactall.logtime, pch = 3, col = 6)
# exactall
exactallfit <- survreg(Surv(time, status >= 0) ~ 1, dist =
    distribution, scale = scale)
# treating censored as exac
t
abline(exactallfit$coef, exactallfit$scale, lty = 3, col = 6)
points(exact.sevq, exact.logtime, pch = 5, col = 5)
# exact points only
exactlyfit <- survreg(Surv(time[ok], status[ok]) ~ 1, dist
    = distribution, scale = scale)
# deleting censored
abline(exactlyfit$coef, exactlyfit$scale, lty = 2, col = 5
)
fit <- survreg(Surv(time, status) ~ 1, dist = "weibull", scale
    = scale)
# censoring taken into account
abline(fit$coef, fit$scale, lty = 1, col = 1)
}
else {
  n <- length(time)
  t.ss <- rep(0, n)
  for(i in 1:n) {
    # This loop assigns probabilities to censored time points,
    # and takes care of tied observations as well
    idx <- time[i] >= survtime
    t.ss[i] <- min(survp[idx], na.rm = T)
  }
  #sevq <- log(qweibull(1 - t.ss, 1))
  sevq <- log(switch(distribution,
    weibull = qweibull(1 - t.ss, 1),
    lognormal = qlnorm(1 - t.ss),
    loglogistic = exp(qlogis(1 - t.ss))))
  # standard extreme value quantile
  logtime <- log(time)
  print(data.frame(logtime, sevq))
  par(mar = c(5, 5, 2, 2))
  plot(sevq, logtime, type = "n", xlab = xlabs, ylab =
    "Ordered Log Time", ...)
  ok <- status == 1
  # exact status only
  points(sevq[ok], logtime[ok], pch = 1)
  # exact points only
  points(sevq[!ok], logtime[!ok], pch = "\255", font = 8)
  # censored points only
  fit <- survreg(Surv(time, status) ~ 1, dist = distribution,
    scale = scale)

```

```

    # censoring taken into account
    abline(fit$coef, fit$scale, lty = 1, col = 1)
  }
  ymax <- max(logtime)
  yrange <- diff(range(logtime))
  yn <- ymax - yrange * seq(0, by = 0.05, length = 5)
  if(pdgy == 1) {
    xmin <- min(c(sevq, exact.sevq, exactall.sevq))
    xrange <- diff(range(c(sevq, exact.sevq, exactall.sevq)))
  }
  else {
    xmin <- min(sevq)
    xrange <- diff(range(sevq))
  }
  x1 <- xmin + 0.05 * xrange
  x2 <- xmin + 0.1 * xrange
  x3 <- xmin + 0.15 * xrange
  points(x1, yn[1], pch = "\255", font = 8)
  text(x3, yn[1], "censored", adj = 0)
  points(x1, yn[2], pch = 1)
  text(x3, yn[2], "exact", adj = 0)
  if(pdgy == 1) {
    lines(c(x1, x2), rep(yn[3], 2), lty = 1, col = 1, lwd = 3)
    text(x3, yn[3], "censoring taken into account", adj = 0)
    lines(c(x1, x2), rep(yn[4], 2), lty = 3, col = 6, lwd = 3)
    text(x3, yn[4], "treating censored as exact", adj = 0)
    lines(c(x1, x2), rep(yn[5], 2), lty = 2, col = 5, lwd = 3)
    text(x3, yn[5], "deleting censored", adj = 0)
  }
  on.exit()
  paste("Q-Q plot for", distribution, "done")
}

months=df.new$Survival
status=df.new$Status
qq.surv(months, status, distribution = "weibull",
  adjpb=0,
  main="Q-Q plot - Weibull fit")
qq.surv(months, status, distribution = "lognormal",
  adjpb=0,
  main="Q-Q plot - Log-normal fit")
qq.surv(months, status, distribution = "loglogistic",
  adjpb=0,
  main="Q-Q plot - Log-logistic fit")

p= ggsurvplot(km.age,
  fun = "cumhaz",
  palette = c("darkcyan","darkgoldenrod3","darkorange3"),
  subtitle="Hazard, Stratified by Age Group",
  font.subtitle = c(10,"italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),

```

```

        ylab="Survival Proportion",
        xlab="Time to Death (Months)",
        legend.title = "Groups",
        legend.labs = c("Age < 63", "Age => 63")
    )
p$plot + geom_vline(xintercept=26)+
  geom_vline(xintercept=46)

p = ggsurvplot(km.effusion,
  fun = "cumhaz",
  palette = c("darkcyan", "darkgoldenrod3"),
  subtitle="Hazard, Stratified by Pericardial Effusion Presence",
  font.subtitle = c(10, "italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)",
  legend.title = "Groups",
  legend.labs = c("Present", "Absent"))
p$plot + geom_vline(xintercept=26)+
  geom_vline(xintercept=46)

p = ggsurvplot(km.wms,
  fun = "cumhaz",
  palette = c("darkcyan", "darkgoldenrod3", "darkorange3"),
  subtitle="Hazard, Stratified by Wall Motion Index",
  font.subtitle = c(10, "italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)")
p$plot + geom_vline(xintercept=26)+
  geom_vline(xintercept=46)

p = ggsurvplot(km.fshort,
  fun = "cumhaz",
  palette = c("darkcyan", "darkgoldenrod3", "darkorange3"),
  subtitle="Hazard, Stratified by Fractional Shortening",
  font.subtitle = c(10, "italic"),
  font.x = c(9, "bold.italic"),
  font.y = c(9, "bold.italic"),
  ylab="Survival Proportion",
  xlab="Time to Death (Months)"
)
p$plot + geom_vline(xintercept=26)+
  geom_vline(xintercept=46)

#Cox-Snell residual analysis for overall model fit

status=df.new$Status[df.new$Survival>=0 & df.new$Survival<=26]
rc=abs(status - mod1$residuals)
km.rc = survfit(Surv(rc,status)~1)

```

```

summary.km.rc=summary(km.rc)
rcu=summary.km.rc$time
surv.rc = summary.km.rc$urv
plot(rcu,-log(surv.rc),type="p",
      xlab="Cox-Snell residual rc",ylab="Cumulative hazard on rc",
      main="Model 1 - Cox-Snell residual model fit evaluation")
abline(a=0,b=1); abline(v=0); abline(h=0)

status=df.new$Status[df.new$Survival>26 & df.new$Survival<=46]
rc=abs(status - mod2$residuals)
km.rc = survfit(Surv(rc,status)~1)
summary.km.rc=summary(km.rc)
rcu=summary.km.rc$time
surv.rc = summary.km.rc$urv
plot(rcu,-log(surv.rc),type="p",
      xlab="Cox-Snell residual rc",ylab="Cumulative hazard on rc",
      main="Model 2 - Cox-Snell residual model fit evaluation")
abline(a=0,b=1); abline(v=0); abline(h=0)

months=df.new$Survival[df.new$Survival>46]
rc=abs(status - mod3$residuals)
km.rc = survfit(Surv(rc,status)~1)
summary.km.rc=summary(km.rc)
rcu=summary.km.rc$time
surv.rc = summary.km.rc$urv
plot(rcu,-log(surv.rc),type="p",
      xlab="Cox-Snell residual rc",ylab="Cumulative hazard on rc",
      main="Model 3 - Cox-Snell residual model fit evaluation")
abline(a=0,b=1); abline(v=0); abline(h=0)

#Schoenfeld residuals; test for constant coefficients

test.ph <- cox.zph(mod1)
ggcoxzph(test.ph, main="Schoenfeld Residuals - Const Coeff Evaluation for Model 1")

test.ph <- cox.zph(mod2)
ggcoxzph(test.ph, main="Schoenfeld Residuals - Const Coeff Evaluation for Model 2")

test.ph <- cox.zph(mod3)
ggcoxzph(test.ph, main="Schoenfeld Residuals - Const Coeff Evaluation for Model 3")

```

Table 6: Kaplan-Meier Estimate Summary

Ni	Di	Ci	Survival	Std. Err	95% LCL	95% UCL
130	0	1	1.0000000	0.0000000	1.0000000	1.0000000
129	0	4	1.0000000	0.0000000	1.0000000	1.0000000
125	0	6	1.0000000	0.0000000	1.0000000	1.0000000
119	0	6	1.0000000	0.0000000	1.0000000	1.0000000
113	0	6	1.0000000	0.0000000	1.0000000	1.0000000
107	0	1	1.0000000	0.0000000	1.0000000	1.0000000
106	0	2	1.0000000	0.0000000	1.0000000	1.0000000
104	0	2	1.0000000	0.0000000	1.0000000	1.0000000
102	0	1	1.0000000	0.0000000	1.0000000	1.0000000
101	0	2	1.0000000	0.0000000	1.0000000	1.0000000
99	0	1	1.0000000	0.0000000	1.0000000	1.0000000
98	0	1	1.0000000	0.0000000	1.0000000	1.0000000
97	1	0	0.9896907	0.0103628	0.9697921	1.0000000
96	2	1	0.9690722	0.0181389	0.9352253	1.0000000
93	1	0	0.9586520	0.0211163	0.9197860	0.9991604
92	4	0	0.9169715	0.0306589	0.8634932	0.9737618
88	2	0	0.8961312	0.0347021	0.8372075	0.9592021
86	1	1	0.8857111	0.0366202	0.8243677	0.9516193
84	3	0	0.8540786	0.0422132	0.7862595	0.9277475
81	2	0	0.8329902	0.0457657	0.7615248	0.9111624
79	3	1	0.8013577	0.0509330	0.7252240	0.8854839
75	0	1	0.8013577	0.0509330	0.7252240	0.8854839
74	1	1	0.7905285	0.0527189	0.7129238	0.8765809
72	1	1	0.7795490	0.0545427	0.7005136	0.8675015
70	3	1	0.7461397	0.0601212	0.6632005	0.8394512
66	1	0	0.7348346	0.0620295	0.6507137	0.8298302
65	3	0	0.7009191	0.0677649	0.6137427	0.8004782
62	4	0	0.6556985	0.0755277	0.5654770	0.7603149
58	5	0	0.5991728	0.0856211	0.5066071	0.7086518
53	4	0	0.5539522	0.0941871	0.4605747	0.6662612
49	1	1	0.5426471	0.0964177	0.4492070	0.6555237
47	4	0	0.4964643	0.1061866	0.4031827	0.6113280
43	3	0	0.4618273	0.1141043	0.3692784	0.5775709
40	3	0	0.4271902	0.1226655	0.3358987	0.5432933
37	5	0	0.3694618	0.1388157	0.2814553	0.4849865
32	3	0	0.3348248	0.1500085	0.2495342	0.4492676
29	2	0	0.3117334	0.1582935	0.2285829	0.4251312
27	4	0	0.2655507	0.1774769	0.1875335	0.3760244
23	2	0	0.2424593	0.1887825	0.1674738	0.3510192
21	2	0	0.2193680	0.2016218	0.1477585	0.3256822
19	1	1	0.2078223	0.2087471	0.1380404	0.3128801
17	3	0	0.1711478	0.2370240	0.1075514	0.2723494
14	1	0	0.1589229	0.2483443	0.0976777	0.2585697
13	1	0	0.1466981	0.2609313	0.0879669	0.2446412
12	1	0	0.1344732	0.2750653	0.0784332	0.2305536
11	1	0	0.1222484	0.2911216	0.0690939	0.2162949
10	1	0	0.1100236	0.3096174	0.0599707	0.2018517
9	2	0	0.0855739	0.3572240	0.0424885	0.1723498
7	1	0	0.0733490	0.3891253	0.0342114	0.1572601
6	2	0	0.0488994	0.4845119	0.0189185	0.1263923
4	2	0	0.0244497	0.6962412	0.0062464	0.0957006
2	1	0	0.0122248	0.9823466	0.0017481	0.0854929
1	1	0	0.0000000	Inf	NA	NA