FRyerson University CKME 136 Data Analytics Capstone Course

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Coronary Artery Disease Prediction Using Classification Algorithms

Introduction

Using Angiography for the diagnosis of Coronary Artery Disease (CAD) is an accurate, but expensive method with many side effects. A more cost-effective and less invasive method is needed in order to make accurate diagnoses of CAD accessible for doctors and patients. This research paper will assess the predictive capability of three different types of machine learning algorithms. The effectiveness of these three types of algorithms will be compared against known results established using angiography. The three types of classification algorithms that will be evaluated are the Random Forest, the Naïve Bayes, and the Logistic Regression algorithms.

The dataset used in this paper is called Z-Alizadeh Sani ¹ and was collected from 303 random visitors to Tehran's Shaheed Rajaei Cardiovascular, Medical and Research Center who were suspected of having CAD. Angiography was used to find that 87 visitors were healthy and 216 had CAD. These results are used as the basis of comparison to assess the predictive capability of the three classification algorithms.

Literature Review

In 2018, heart disease is the second leading cause of death in Canada ², the United States ³, and in fact it remains the leading cause of death in industrialized nations around the world ⁴. Coronary heart disease is also known as ischaemic heart disease and it refers to the build-up of plaque in the heart's arteries which can cause a heart attack, a stroke or heart failure. Heart disease is more common in men, people who smoke, people who are overweight or obese, people over the age of 55 and for those with a family history of heart disease or heart attack. As such, there are risk factors that can be controlled and others that cannot. Making lifestyle changes can reduce the chance of having heart disease. Some controllable risk factors include smoking, high low-density lipoprotein (LDL, often called 'bad' cholesterol), low high-density lipoprotein (HDL, often called 'good' cholesterol), uncontrolled high blood pressure, physical inactivity, obesity, uncontrolled diabetes, and uncontrolled stress.

Angiography is considered an accurate method used to diagnose the presence of coronary heart disease. An angiogram allows doctors to view blood flow through the heart by injecting a special dye into the coronary arteries. The dye is injected into arteries of the heart through a flexible catheter that is threaded through an artery, usually in the leg. This dye shows narrow spots and blockages on X-ray images ⁵. There are many risks and potential side-effects of angiograms. There is the extremely small chance of developing cancer in the long term due to the exposure to the radiation. There are also potential side-effects with some medications, such as blood thinning and diabetic medications. In addition, there are the risks of allergic reactions, infection, blood clot, and weakness of the blood vessel wall ⁶.

Two other diagnostic tests that are referred to in the dataset used for this paper are the Electrocardiogram (ECG) and the Echocardiogram. An ECG records electrical signals as they travel through the heart and can show evidence of a previous heart attack or one that is in progress and can also show abnormalities indicating inadequate blood flow to the heart. An echocardiogram uses sound waves to produce images of the heart. Such images are used to determine whether all parts of the heart are contributing normally to the heart's pumping activity ⁷.

There have been numerous studies completed using various data mining algorithms for the purpose of heart disease prediction. The use of such algorithms in health care applications has gained in popularity over recent years given their efficiency, cost and non-invasive advantages. Heart disease prediction by using a number of different machine learning algorithms on a few different datasets has been successfully achieved in these studies. Sonam Nikhar and A.M. Karandikar (2016) ⁸ concluded that the Decision Tree classifier provided better results in the diagnosis of heart disease than Neural Network, Support Vector Machine (SVM) or k-Nearest Neighbour (kNN) classifier techniques. Aamanpreet Kaur (2017) ^{9,10} found that the Random Forest algorithm is quite effective at correctly classifying instances of heart disease. N. Bhatla and K. Jyoti (2012) ¹¹ were able to achieve high accuracy with the Naïve Bayes and Decision Tree classifiers while also reducing the number of attributes used from 15 to 4. Mai Shouman et al (2012) ¹² was able to show that applying the k-Nearest Neighbour (kNN) algorithm can achieve higher accuracy than neural network ensemble in the diagnosis of heart disease.

However, while there has been significant success predicting heart disease with various data mining techniques, Mudasir Kirmani (2017) ¹³ found that there is no single classifier which produces the best results for every dataset and no single data mining technique which gives consistent results for all types of healthcare data. As such, further investigation is still needed given the availability of huge amounts of medical data and the subsequent need for data analysis tools to extract useful knowledge.

The three classification algorithms that will be used in this paper are the Random Forest, the Naïve Bayes, and the Logistic Regression algorithms. All three of these machine learning algorithms are and/or can be non-parametric methods that can be used for classification and regression ¹⁴. A confusion matrix is an available output for all three types of algorithms which will be used to evaluate their accuracy, sensitivity and specificity. Sensitivity measures the true positive prediction rate, specificity measures the true negative prediction rate, and accuracy is calculated as the number of all correct positive and negative predictions divided by the total number of observations in the dataset ¹⁵.

Dataset

The dataset that is used for this paper is called the Z-Alizadeh Sani dataset. It has been downloaded from the UCI Machine Learning Repository (https://archive.ics.uci.edu/ml/machine-learning-databases/00412/). This dataset contains the records of 303 patients, each of which have 54 attributes. All of the attributes are considered as indicators of Coronary Artery Disease (CAD). These attributes are split into four groups: Demographic, Symptom & Examination, ECG, and Laboratory & Echo Features. The Demographic group includes 16 attributes, the Symptom & Examination group includes 14 attributes, the ECG group includes 7 attributes and the Laboratory & Echo Features group includes 17 attributes. This dataset shows the results of angiography testing in which each patient is diagnosed into one of two possible categories: CAD or Normal. A patient is diagnosed as CAD if their coronary artery diameter narrowing is greater than or equal to 50%, otherwise they are categorized as Normal. The actual diagnoses of CAD or Normal was determined using angiography. The results showed that 87 patients were healthy (that is, 'Normal') and 216 had CAD.

The following four tables present all of the attributes used in the Z-Alizadeh Sani dataset, broken into the four groups, and include their valid ranges. Table 5 defines the discretized features and ranges of some of the attributes in tables 1, 2 and 4. Table 6 provides a summary of the attributes *before* there any changes made to the data types and before any discretization is completed.

DEMOGRAPHIC ATTRIBUTES

Feature Name	Column Header	Range		
Age	Age	30 – 86		
Weight (kgs)	Weight	48 – 120		
Sex	Sex	Male, Female		
Body Mass Index (kg/m²)	BMI	18 – 41		
History of Diabetes Mellitus	DM	Yes, No		
History of Hypertension (High Blood Pressure)	HTN	Yes, No		
Current Smoker	Smoker	Yes, No		
Ex-Smoker	ExSmoker	Yes, No		
FH (Family History of Heart Disease in First-Degree Relatives)	FH	Yes, No		
Obesity	Obesity	Yes, if BMI > 25, No otherwise		
Chronic Renal Failure	CRF	Yes, No		
Cerebrovascular Accident (often referred to as a Stroke)	CVA	Yes, No		
Airway Disease (such as asthma, COPD and bronchiectasis)	AD	Yes, No		
Thyroid Disease	TD	Yes, No		
Congestive Heart Failure	CHF	Yes, No		
Dyslipidemia (elevated cholesterol, triglycerides, or both)	DLP	Yes, No		

Table 1

<u>Note</u>: 'Age' will change to a categorical attribute with 2 levels, as defined on Table 5. 'Weight' will be discretized as 'Average' or 'High' using the below or above mean results using the data from this attribute since it was not discretized in the original dataset.

SYMPTOM & EXAMINATION ATTRIBUTES

Feature Name	Column Header	Range
Blood Pressure (mmHg)	BP	90 - 190
Pulse Rate (pulse per minute)	PR	50 - 110
Edema (excess fluid trapped in the body's tissues)	Edema	Yes, No
Weak Peripheral Pulse (weak pulse in extremities)	WPP	Yes, No
Lung Rales (rattling/bubbling sound in lungs)	LR	Yes, No
Systolic Murmur	SysM	Yes, No
Diastolic Murmur	DiaM	Yes, No
Typical Chest Pain (pressure/squeezing in chest)	TCP	Yes, No
Dyspnea (shortness of breath)	Dyspnea	Yes, No
Heart Failure Functional Class ²⁰ *	Fclass	1, 2, 3, 4
Atypical Chest Pain (not heart-related pain)	ACP	Yes, No
Nonanginal Chest Pain (typically lasting over 30 minutes)	NCP	Yes, No
Exertional Chest Pain (pain from exertion or excitement)	ECP	Yes, No
Low Threshold Angina (low threshold for angina pain)	LTAng	Yes, No

Table 2

<u>Note:</u> Blood Pressure and Pulse Rate will be changed to categorical attributes (with 3 levels). Heart Failure Functional Class will change from a categorical attribute with 4 levels, to having 2 levels. These changes and levels are defined on Table 5.

ECG ATTRIBUTES

Feature Name	Column Header	Range
Q Wave (Present = Yes, not present = No)	Qwave	Yes, No
ST Elevation (Elevation present = Yes, not present = No)	Stelev	Yes, No
ST Depression (Depression present = Yes, not present = No)	Stdep	Yes, No
T Inversion (Inverted = Yes, not inverted = No)	Tinv	Yes, No
Left Ventricular Hypertrophy	LVH	Yes, No
Poor R Wave Progression	PoorR	Yes, No
Bundle Branch Block	BBB	LBBB, RBBB, No

Table 3

^{*} Heart Failure Functional Class: https://www.chf-solutions.com/heart-failure-classifications/

LABORATORY & ECHOCARDIOGRAPHIC ATTRIBUTES

Feature Name	Column Header	Range			
Fasting Blood Sugar (mg/dl)	FBS	62 - 400			
Creatine (mg/dl)	Cr	0.5 - 2.2			
Triglyceride (mg/dl)	TG	37 - 1050			
Low Density Lipoprotein (mg/dl)	LDL	18 - 232			
High Density Lipoprotein (mg/dl)	HDL	15 - 111			
Blood Urea Nitrogen (mg/dl)	BUN	6 to 52			
Erythrocyte Sedimentation Rate (mm/h)	ESR	1 to 90			
Hemoglobin (g/dl)	Hb	8.9 - 17.6 3.0 - 6.6			
Potassium (mEq/lit)	K				
Sodium (mEq/lit)	, , ,				
White Blood Cell (cells/ml)	WBC	3700 - 18000			
Lymphocyte (%) ²¹	Lymph	7 to 60			
Neutrophil (%) ²²	Neut	32 - 89			
Platelet (1000/ml)	PLT	25 - 742			
Ejection Fraction (%)		15 - 60			
Region With Regional Wall Motion Abnormality					
Valvular Heart Disease	VHD	Normal, Mild, Moderate, Severe			

Table 4

<u>Note:</u> For those attributes in Table 4 that are not already categorical, all but two of them will be changed to categorical attributes, as defined in Table 5. Lymphocyte and Neutrophil will be discretized using accepted ranges provided by the medical community per the references provided.

Discretized Features and Their Range of Values

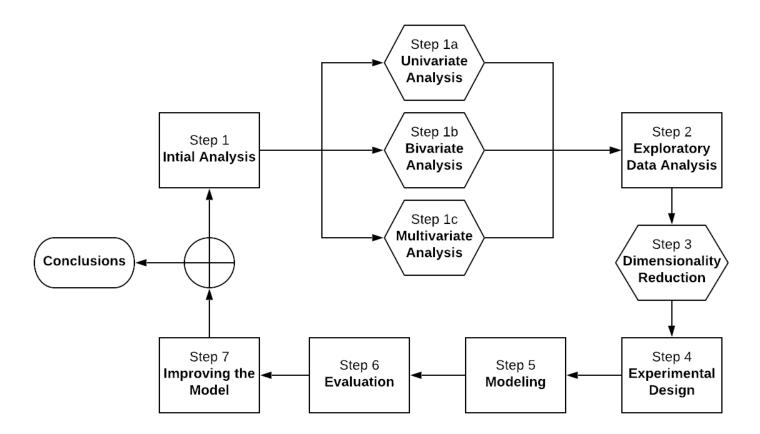
Feature Name	Low	Medium	High		
Fasting Blood Sugar	FBS < 70	70 ≤ FBS ≤ 105	FBS > 105		
Creatine	Cr < 0.7	0.7 ≤ Cr ≤ 1.5	Cr > 1.5		
Triglyceride		TG ≤ 200	TG > 200		
Low Density Lipoprotein		LDL ≤ 130	LDL > 130		
High Density Lipoprotein	HDL < 35	HDL≥35			
Blood Urea Nitrogen	BUN < 7	7 ≤ BUN ≤ 20	BUN > 20		
Erythrocyte Sedimentation Rate		If male & ESR ≤ age/2 or if female & ESR ≤ (age/2) + 5	If male & ESR > age/2 or if female & ESR > (age/2) +5		
Hemoglobin	If male & Hb < 14 or if female & Hb < 12.5	If male & 14 ≤ Hb ≤ 17 or if female & 12.5 ≤ Hb ≤15	If male & Hb > 17 or if female & Hb > 15		
Potassium	K < 3.8	3.8 ≤ K ≤ 5.6	K > 5.6		
Sodium	Na < 136	136 ≤ Na ≤ 146	Na > 146		
White Blood Cell	WBC < 4000	4000 ≤ WBC ≤ 11000	WBC > 11000		
Lymphocyte (%) ²¹	Lymph < 18	18 ≤ Lymph ≤ 45	Lymph > 45		
Neutrophil (%) ²²	Neut < 44	45 ≤ Neut ≤ 75	Neut > 75		
Platelet	PLT < 150	150 ≤ PLT ≤ 450	PLT > 450		
Ejection Fraction (%)	EF ≤ 50	EF > 50			
Regional Wall Motion Abnormality		RWMA = 0	RWMA ≠ 0		
Blood Pressure	BP < 90	90 ≤ BP ≤ 140	BP > 140		
Pulse Rate	PR < 60	60 ≤ BP ≤ 100	PR > 100		
Heart Failure Functional Class		1	2, 3, 4		
Weight		\leq 70.52 kg if female, > 70.52 kg if f or \leq 76.21 kg if male or > 76.21 kg			
* Age		If male & age \leq 45 or If male & age \leq if female & age \leq 55 if female & age			

Table 5

^{*} Since women under age 55, and men under age 45 are less affected by CAD, the range of age is partitioned at these values

	Data Set Attribute Data Types Before Any Changes								
	Column #	Heading	Int, Factor or Numeric?	Values on CSV File	Values in R	Levels			
int	1	Age	int	2 digit numbers	2 digit numbers				
int	2	Weight	int	2 digit numbers	2 digit numbers				
Factor	3	Sex	Factor	Fmale, Male	2 and 1	2	Male = 2		
num	In	BMI	num	floating Pt #'s	floating Pt #'s				
	5	DM	int	0 and 1	0 and 1		1 = Y		
int	6	HTN	int	0 and 1	0 and 1				
	7	Smoker	int	0 and 1	0 and 1				
	8	ExSmoker	int	0 and 1	0 and 1				
	9	FH	int	0 and 1	0 and 1				
	10	Obesity	Factor	Y and N	1 and 2	2	Y = 2		
	11	CRF	Factor	Y and N	1 and 2	2			
Factor	12	CVA	Factor	Y and N	1 and 2	2			
	13	AD	Factor	Y and N	1 and 2	2			
	14	TD	Factor	Y and N	1 and 2	2			
	15	CHF	Factor	Y and N	1 and 2	2			
	16	DLP	Factor	Y and N	1 and 2	2			
	17	ВР	int	3 digit numbers	3 digit numbers				
int	18	PR	int	2 digit numbers	2 digit numbers				
	19	Edema	int	0 and 1	0 and 1				
	20	WPP	Factor	Y and N	1 and 2	2			
	21	LR	Factor	Y and N	1 and 2	2			
Factor	22	SysM	Factor	Y and N	1 and 2	2			
	23	DiaM	Factor	Y and N	1 and 2	2			
int	24	TCP	int	0 and 1	0 and 1	_			
Factor	25	Dyspnea	Factor	Y and N	1 and 2	2			
int	26	Fclass	int	0, 1, 2, 3	0, 1, 2, 3	2			
IIIC	27	ACP	Factor	Y and N	1 and 2				
	28	NCP	Factor	Y and N	1 and 2				
Factor		ECP		N N	1	1	NI - 1		
	30		Factor	Y and N	1 and 2	1	N = 1		
		LTAng	Factor						
	31	Qwave	int	0 and 1	0 and 1				
int	32	Stelev	int	0 and 1	0 and 1				
		Stdep	int	0 and 1	0 and 1				
	34	Tinv	int	0 and 1	0 and 1	2			
	35	LVH	Factor	Y and N	1 and 2	2			
Factor	36	PoorR	Factor	Y and N	1 and 2	2			
	37	BBB	Factor	N, LBBB, RBBB	1, 2 (0 or 3?)	3	LBBB = 1	N = 2	RBBB :
int	38	FBS	int	Reg numbers	Reg numbers				
num	39	Cr	num	Decimals	Decimals				
int	40	TG	int	Reg numbers	Reg numbers				
	41	LDL	int	Reg numbers	Reg numbers				
num	42	HDL	num	Reg numbers	Reg numbers				
int	43	BUN	int	Reg numbers	Reg numbers				
	44	ESR	int	Reg numbers	Reg numbers				
num	45	Hb							
	46	K	num	Decimals	Decimals				
	47	Na	int	Reg numbers	Reg numbers				
	48	WBC	int	Reg numbers	Reg numbers				
	49	Lymph	int	Reg numbers	Reg numbers				
int	50	Neut	int	Reg numbers	Reg numbers				
	51	PLT	int	Reg numbers	Reg numbers				
	52	EF	int	Reg numbers	Reg numbers				
	53	RWMA	int	Reg numbers	Reg numbers				
Factor	54	VHD	Factor	mild, Moderate, Severe, N	1, 2, 3, 4	4	mild = 1	Moderate = 2	
							N = 3	4 = Severe	
Factor	55	CAD	Factor	Cad, Normal	1 and 2	2	CAD = 1	Normal = 2	

Approach



Approach Details

Step 1: Initial Analysis

- Download the Z-Alizadeh Sani dataset
- Review attributes to understand, create and update the data dictionary
- Define the four groups of attributes and how each set of attributes are related
- Clarify the dependent variable that is going to be predicted
- Import dataset as a .csv file into R Studio
- Update column headers, as needed, for clarity and/or correspondence with data dictionary
- Remove 'Length' attribute from data set. This attribute was ultimately not included in the original study, presumably due to the fact that they have the Body Mass Index values, and as such, 'length' (that is, height) is redundant.
- Remove dependent attribute from data set in R not used for algorithms
- Determine whether there are any missing values
- Determine whether there are any errors, duplicates and/or incomplete or inappropriate entries
- Assign the correct data types (numeric, categorical, etc)
- Create four attribute groups as four data frames to be used throughout the project
- ** Need to change the attribute scale from numeric to factor, and assign them as 'low', 'medium' and 'high' as defined in the 'Discretized Features & Their Range of Values' table shown in the Dataset section (Table 5). Will discretize lymphocyte and neutrophil percentages into 'low', 'medium' and 'high' limits using accepted medically defined cut-offs. **
- ** Change the data types to 'ordered factor' for those that are factor data types, but retain some ordering (such as 'low, 'medium', 'high')

• Will need to further discretize categorical attributes with greater than 2 levels (refer to chart made up of all attributes and their initial data types, levels, understandings).

Step 1a: Univariate Analysis

- For numeric attributes, check the min, mean, max, first and third quartiles.
- Create box plots to visualize these numeric summary values and see outliers (if any). Determine appropriate actions for any outliers (some attributes have been discretized which addresses those outliers)
- Check the number of levels and their meanings for categorical attributes and make sure they are correct at this stage.
- Check on the distributions of the attributes. Given the high number of attributes (at this stage), the Shapiro-Wilk test of normality will be more useful than visualizing all of the individual distributions. Although the algorithms used later are appropriate for non-parametric data, it is still useful to have an understanding of the distributions for interpretation of results later.
- Check for any 'very low' variance for individual attributes. This will help determine which attributes can be removed later (if any).
- Check whether there is an imbalance in the dependent variable. Since the number of 'Normal' results to 'Coronary Artery Disease (CAD)' results are 87 to 216, respectively, any imbalance will need to be evaluated.
- Create visualizations of the attributes within each of the four attribute groups. Keep visualizations that appear to display something 'interesting' about the dataset. This will be updated/improved as I learn more about the relationships of the attributes.

Step 1b: Bivariate Analysis

- Investigate the pairwise relations between the input variables as well as between the input variables and output variable. Create scatter plots to visualize these relations.
- Complete correlation analysis and check the significance of relationships between attributes. As appropriate, use Pearson Correlation Coefficient for two quantitative variables; Chi-square for two categorical variables using Cramer's V test to show the strength of any association between two factor variables; and ANOVA for one categorical and one quantitative variable.

Step 1c: Multivariate Analysis

• Will consider the One-Way-ANOVA for the parametric attributes and the Kruskal-Wallis test for the non-parametric attributes.

Step 2: Exploratory Data Analysis

- Determine which attributes should be normalized, if any. After discretizing the attributes, as needed, it does not appear that there will be a need to normalize any other attributes (such as age, weight, neutrophil and lymphocyte since these will ultimately be discretized during the initial analysis).
- Review whether additional sub-setting can clarify attribute relationships further. For example, subset those patients who have been diagnosed with CAD versus those that do not. Subset further between men and women. Save these dataframes for later investigation and/or predictions.

Step 3: Dimensionality Reduction

- This will be an important step in this project. There are a lot of attributes to consider, even within the four attribute groups. I anticipate that the correlation and variation analyses will enable to removal of some attributes.
- Feature Selection will be used to aid in the dimensionality reduction. In particular, Forward Selection and/or Backward Elimination will be used.

- FSelector and FSelectorRcpp packages will be used for feature selection. I will use these packages on the discretized data set, as well as the data set before discretization to compare the results. I will also compare those results with the feature selection done with the caret package in R (using Recursive Feature Elimination).
- Save final dataframes for each of the four attribute groups. Also save a separate dataframe with all four attribute groups combined, both discretized and before discretization.

Step 4: Experimental Design

- In order to ensure that there are balanced splits between the training and testing splits of the data set, I am using the createDataPartition function in R to create the initial split for 75% testing and 25% testing.
- Given that this data set is not very large, the splitting to be used for the training of the algorithms will be the done by using the 'k-fold cross-validation' method. Specifically, I will use a 10-fold cross validation, repeated 10 times.
- The 10-fold cross validation should help with addressing any 'overfitting' concerns since there are a lot of attributes (especially given that the random forest algorithm will be evaluated)
- Address the class imbalance for the output variable (the dependent variable). That is, given that there was a total of 87 patients (28.7%) that were found to be 'Normal' and 216 (71.3%) were diagnosed with 'CAD', that means that there is some class imbalance. Planning to use the 'SMOTE' function in R to assist with this problem. SMOTE stands for "Synthetic Minority Over-Sampling Technique" and this function will synthesize new minority instances between the existing (actual) minority instances.

Step 5: Modeling

- The research question in this paper that is, the prediction of the presence of coronary artery disease using a number of independent variables requires that we use a classification algorithm for the answer. In addition, given that the data set is non-linear, it is appropriate to use the Random Forest, the Naïve Bayes, and Logistic Regression algorithms. The effectiveness, efficiency and stability of these three algorithms will be compared.
- Run the three algorithms on the 'combined' dataframe (that is, with the four final attribute group dataframes together).

Step 6: Evaluation

- The accuracy, sensitivity (true positive prediction rate) and specificity (true negative prediction rate) are very relevant metrics to use for evaluating algorithms used for predicting medical diagnoses. The Confusion Matrix provides these evaluation results and all three algorithms output a confusion matrix. This is another reason to run these three machine learning algorithms for this research question.
- ** However, when it comes to the diagnosis of coronary heart disease, being falsely diagnosed with CAD is more
 'tolerable' than being falsely diagnosed as 'Normal'. As such, emphasis should be placed on the <u>Sensitivity</u>
 results for this data set.
- I will also generate the F-score (F1) to help evaluate the results. That is, the F1-Score considers <u>both</u> 'Precision' and Sensitivity. It is the 'harmonic mean' of the precision and sensitivity. In addition, F1 is a good evaluation measure to use when there is an uneven class distribution (as is the case with this data set).

Precision provides a measure of how many of those patients who were predicted to have CAD, actually have CAD. Precision = True Positive CAD/False Positive CAD + True Negative CAD.

The F1 Score = 2*(Precision*Sensitivity)/Precision + Sensitivity.

Precision, Sensitivity and the F1-Score can be generated with the function "confusionMatrix()" from the 'Caret' package in R.

- For additional evaluation and understanding of the interpretation of the impacts of the final variables selected
 for use on these three algorithms. I will use the logistic regression algorithm in the base R package to see
 whether the log odds of being diagnosed with Coronary Artery Disease increases or decreases with a one unit
 increase of each of the final attributes.
- I will measure the performance efficiency of the machine language algorithms using the 'microbenchmark' package in R.
- Lastly, I will evaluate the stability of the algorithms and confirm whether they generate consistent results. I will graph the results to help determine stability and any evidence of overfitting.

Step 7: Improving the Model

- I will check whether building a machine learning ensemble improves my results. I will do this using the 'SuperLearner' R package.
- In the event of very high accuracy results, I will need to determine whether overfitting is a problem with the Random Forest algorithm, even though there have been preventative measures taken (such as 10-fold cross validation and dimensionality reduction).
- Will need to pay attention to the number of 'events per variable' (EPV) on the predictive accuracy performance for the logistic regression algorithm. The EPV is equal to the number of events (87, being the lesser of 216 and 87, from the 303) divided by the number of predictor variables considered in developing the prediction model. ¹⁶. It is the number of events divided by the number of degrees of freedom required to represent all of the variables in the model. For example, a three-level categorical variable would require two degrees of freedom.
- Will want to continue to determine whether additional feature selection is possible (Forward Selection and/or Backward Elimination). Will need to continually evaluate whether more attributes can be eliminated without adversely affecting predictive accuracy.

Final Step: Conclusions

- Describe the inferences that are possible using these three machine learning algorithms.
- Summarize which algorithm(s) provide the best accuracy, sensitivity and specificity results.
- Provide overview of threats to validity of results (such as sample size and number of attributes used)
 and what solutions are available to reduce the effects of such threats.
- Provide conclusions regarding whether the accuracy, sensitivity and specificity results enable a
 conclusion that the inferences made are appropriate (that is, that the algorithms measure what they
 claim to that they provide 'construct validity').
- Summarize the conclusions in terms that are relevant to the users of such data. That is, to provide concrete, specific recommendations of which attributes should be used, and why.

References

- [1] UC Irvine Machine Learning Repository. "Heart Disease Data Set", retrieved 05/06/2019 from https://archive.ics.uci.edu/ml/datasets/Heart+Disease.
- [2] Canada.ca. "Report from the Canadian Chronic Disease Surveillance System: Heart Disease in Canada, 2018", retrieved on 06/01/2019 from https://www.canada.ca/content/dam/phac-aspc/documents/services/publications/diseases-conditions/report-heart-disease-canada-2018/pub1-eng.pdf
- [3] Healthline.com. "What Are The 12 Leading Causes of Death in the United States?" retrieved on 06/01/2019 from https://www.healthline.com/health/leading-causes-of-death.

- cdc.gov. "Prevalence of Uncontrolled Risk Factors for Cardiovascular Disease: Unites States, 1999 2010", retrieved on 06/01/2019 from https://www.cdc.gov/nchs/data/databriefs/db103.pdf.
- [4] Academicoup.com. British Medical Bulletin. "Global and Regional Causes of Death", retrieved on 06/01/2019 from https://academic.oup.com/bmb/article/92/1/7/332071.
- [5] Mayoclinic.org. "Coronary Artery Disease", retrieved on 06/01/2019 from https://www.mayoclinic.org/diseases-conditions/coronary-artery-disease/diagnosis-treatment/drc-20350619.
- [6] Imagingpathways.health.wa.gov.au. "Information for Consumers Angiography (Angiogram)", retrieved on 06/01/2019 from http://www.imagingpathways.health.wa.gov.au/index.php/consumer-info/imaging-procedures/angiography.
- [7] Mayoclinic.org. "Coronary Artery Disease", retrieved on 06/01/2019 from https://www.mayoclinic.org/diseases-conditions/coronary-artery-disease/diagnosis-treatment/drc-20350619.
- [8] Media.neliti.com. "Prediction of Heart Disease Using Machine Learning Algorithms", retrieved on 06/13/2019 from https://media.neliti.com/media/publications/239484-prediction-of-heart-disease-using-machin-4b2e96d4.pdf.
- [9] Ijiet.com. "A Comprehensive Approach to Predict Heart Disease Using Data Mining", retrieved on 06/03/2019 from http://ijiet.com/wp-content/uploads/2017/05/3.pdf.
- [10] UC Irvine Machine Learning Repository. "Heart Disease Data Set Cleveland Database", retrieved 06/03/2019 from https://archive.ics.uci.edu/ml/datasets/Heart+Disease.
- [11] Pdfs.semanticscholar.org. "A Novel Approach for Heart Disease Diagnosis Using Data Mining and Fuzzy Logic", retrieved on 06/03/2019 from https://pdfs.semanticscholar.org/cd47/47f0ebabb741c3b30b83b28b71b3aeaf7b04.pdf.
- [12] Ijiet.com. "Applying k-Nearest Neighbour in Diagnosing Heart Disease Patients", retrieved on 06/13/2019 from http://www.ijiet.org/papers/114-K0009.pdf
- [13] Computersciencejournal.org. "Cardiovascular Disease Prediction Using Data Mining Techniques: A Review", retrieved on 06/10/2019 from http://www.computerscijournal.org/vol10no2/cardiovascular-disease-prediction-using-data-mining-techniques-a-review/.
- [14] Towardsdatascience.com. "Comparative Study on Classic Machine Learning Algorithms", retrieved on 06/03/2019 from https://towardsdatascience.com/comparative-study-on-classic-machine-learning-algorithms-24f9ff6ab222.
- [15] Classeval.wordpress.com. "Basic Evaluation Measures From The Confusion Matrix", retrieved on 06/03/2019 from https://classeval.wordpress.com/introduction/basic-evaluation-measures/.
- [16] Ncbi.nlm.nih.gov. "Events per variable (EPV) and the relative performance of different strategies for estimating the out-of-sample validity of logistic regression models", retrieved on 06/10/2019 from https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5394463/

- [17] R. Alizadehsani et al., "A data mining approach for diagnosis of coronary artery disease," Computer Methods and Programs in Biomedicine, vol.111, no.1, pp.52-61, Jul. 2013.
- [18] R. Alizadehsani, M.H. Zangooei, M.J. Hosseini, J. Habibi, A. Khosravi, M. Roshanzamir, F. Khozeimeh, N. Sarrafzadegan, S. Nahavandi, Coronary artery disease detection using computational intelligence methods, Knowledge-Based Systems, 109 (2016) 187-197.
- [19] Z. Arabasadi, R. Alizadehsani, M. Roshanzamir, H. Moosaei, and A. A. Yarifard, "Computer aided decision making for heart disease detection using hybrid neural network-Genetic algorithm," Computer Methods and Programs in Biomedicine, vol. 141, pp. 19-26, 2017/04/01/ 2017.
- [20] Chf-solutions.com. "Heart Failure Classifications", retrieved on 06/02/2019 from https://www.chf-solutions.com/heart-failure-classifications/.
- [21] Healthline.com. "Everything you should know about Lymphocytes", retrieved on 06/21/2019 from https://www.healthline.com/health/lymphocytes#results.
- [22] Healthline.com. "Understanding Neutrophils: Function, Counts and More", retrieved on 06/21/2019 from https://www.healthline.com/health/neutrophils.
- [23] ncbi.nlm.nih.gov. "Neutrophil Lymphocyte Ratio and Cardiovascular Disease Risk: A Systematic Review and Meta-Analysis", retrieved on 06/26/2019 from https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6252240/.
- [24] Healthline.com. "Blood Differential Test", retrieved on 06/26/2019 from https://www.healthline.com/health/blood-differential#test-results .
- [25] ncbi.nlm.nih.gov. "What is the normal value of the neutrophil-to-lymphocyte ratio?", retrieved on 06/26/2019 from https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5217256/.
- [26] ncbi.nlm.nih.gov. "The value of echocardiographic regional wall motion abnormalities in detecting coronary artery disease in patients with or without a dilated left ventricle", retrieved on 06/29/2019 from https://www.ncbi.nlm.nih.gov/pubmed/3157304.
- [27] Mayoclinic.org. "Ejection fraction: What does it measure?", retrieved on 06/29/2019 from https://www.mayoclinic.org/ejection-fraction/expert-answers/faq-20058286.
- [28] Researchgate.net. "Survey of Machine Learning Algorithms for Disease Diagnostic", retrieved on 07/01/2019 from https://www.researchgate.net/publication/312629315 Survey of Machine Learning Algorithms for Disease Diagnos tic.
- [29] Stefan Lessmann et al., "Benchmarking state-of-the-art classification algorithms for credit scoring: A ten-year update", European Journal of Operational Research (doi:10.1016/j.ejor.2015.05.030)