Classification and Logistic Regression

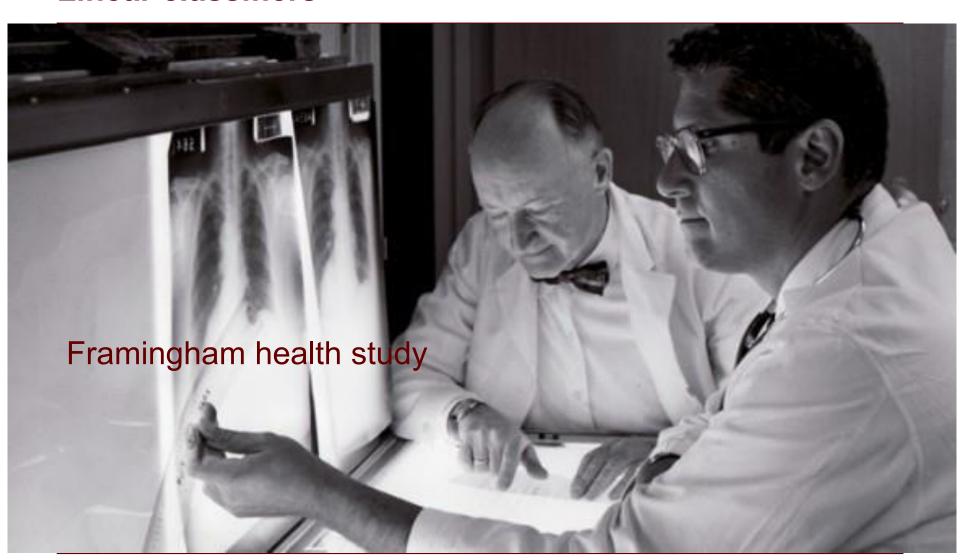
CS556











- Huge impact!
- Used everywhere: marketing, online industry, health care...
- Notable examples: spam filters, all sorts of online services,
 Framingham health study, assessing health care

Linear classifiers in a nutshell



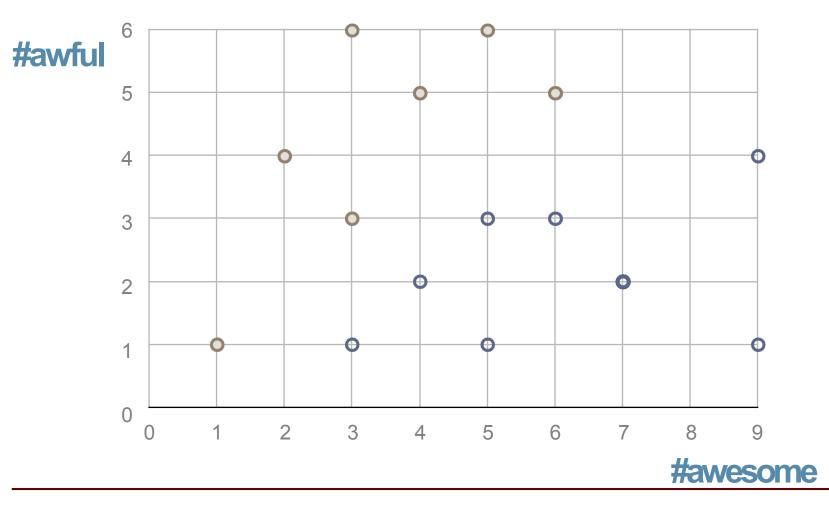
Hamburger was awesome ... service was awful... price was awesome

Pizza was awful ... hot dog was awesome...price was awesome Pizza was awesome ... hot dog was awful...price was awesome

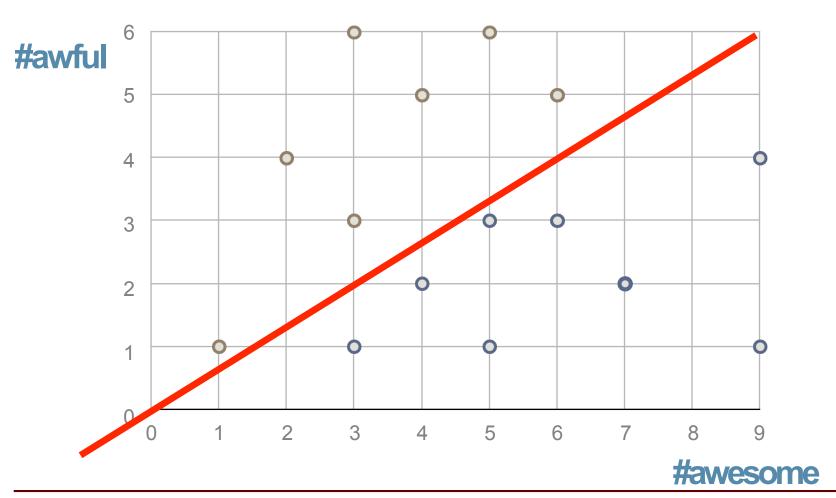
Pizza was awful ... hot dog was awful...price was awful

Sentiment analysis

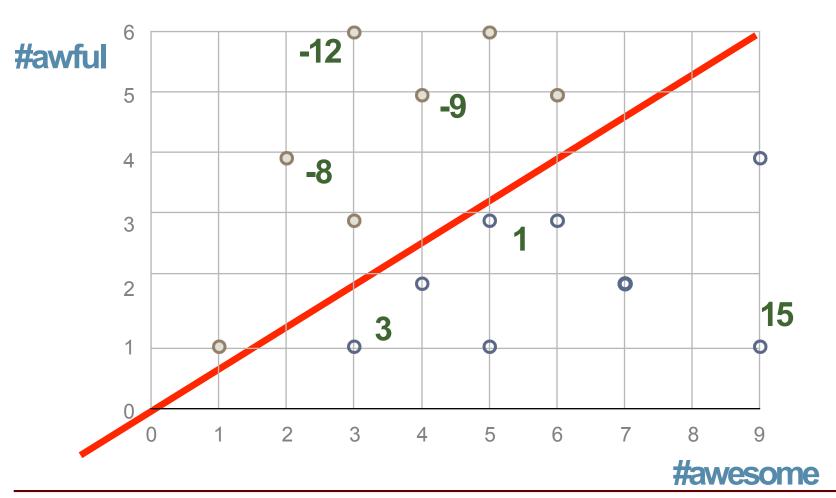
People LOVE US HATE US On Son Son Son Son Son Son Son Son Son So	#awful	#awesome	sentiment
pizza was awful ice cream was awesomeprice was awesome	1	2	+
hot dog was awesomemartin i was awesome price was awful	1	2	+
everything was awful	1	0	_
wine was awfulpasta was awfuldessert was awfulview was	3	1	_



score(x,y)=2x-3y

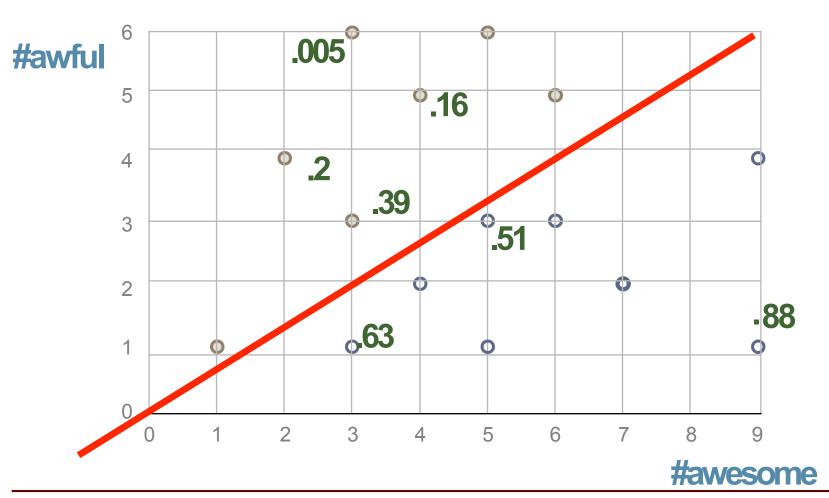


score(x,y)=2x-3y

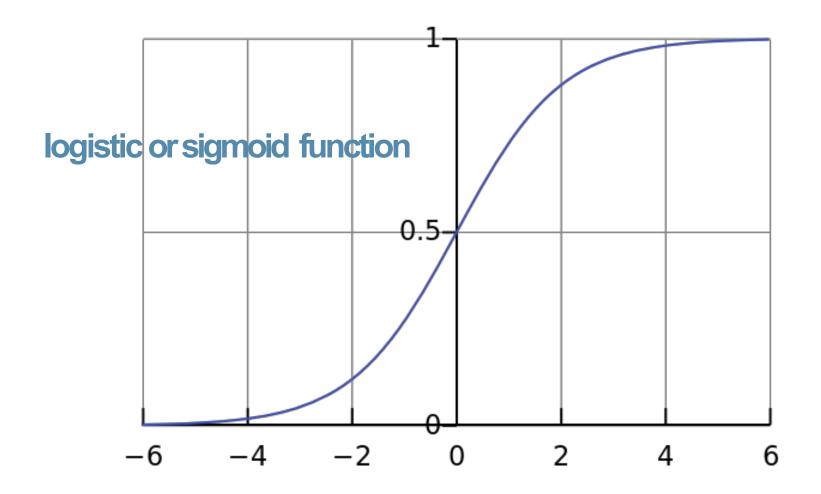


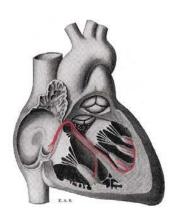


score(x,y)=2x-3y



Squeezing scores into probabilities



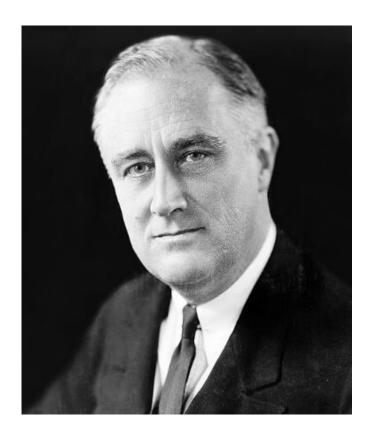


The Framingham Heart Study

Evaluating Risk Factors to Save Lives

Franklin Delano Roosevelt (FDR)

- President of the United States, 1933-1945
 - Longest-serving president
 - Led country through Great Depression
 - Commander in Chief of U.S. military in World War II
- Died while president, April 12, 1945



FDR's Blood Pressure

- Before presidency, blood pressure of 140/100
 - Healthy blood pressure is less than 120/80
 - Today, this is already considered high blood pressure
- One year before death, 210/120
 - Today, this is called Hypertensive Crisis, and emergency care is needed
 - FDR's personal physician:
 - "A moderate degree of arteriosclerosis, although no more than normal for a man of his age"
- Two months before death: 260/150
- Day of death: 300/190

Early Misconceptions

- High blood pressure dubbed essential hypertension
 - Considered important to force blood through arteries
 - Considered harmful to lower blood pressure
- Today, we know better
- "Today, presidential blood pressure numbers like FDR's would send the country's leading doctors racing down hallways ... whisking the nation's leader into the cardiac care unit of Bethesda Naval Hospital."
- -- Daniel Levy, Framingham Heart Study Director

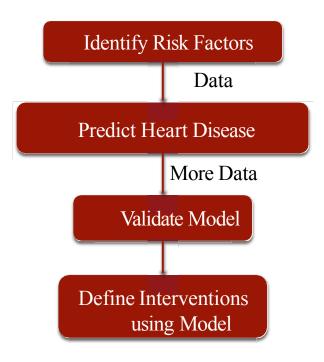
How Did we Learn?

- In late 1940s, U.S. Government set out to better understand cardiovascular disease (CVD)
- Plan: track large cohort of initially healthy patients over time
- City of Framingham, MA selected as site for study
 - Appropriate size
 - Stable population
 - Cooperative doctors and residents
- 1948: beginning of Framingham Heart Study

The Framingham Heart Study

- 5,209 patients aged 30-59 enrolled
- Patients given questionnaire and exam every 2 years
 - Physical characteristics
 - Behavioral characteristics
 - Test results
- Exams and questions expanded over time
- We will build models using the Framingham data to predict and prevent heart disease

Analytics to Prevent Heart Disease



Coronary Heart Disease (CHD)

- We will predict 10-year risk of CHD
 - Subject of important 1998 paper, introducing the Framingham Risk Score
- CHD is a disease of the blood vessels supplying the heart
- Heart disease has been the leading cause of death worldwide since 1921
 - 7.3 million people died from CHD in 2008
 - Since 1950, age-adjusted death rates have declined 60%

Risk Factors

- Risk factors are variables that increase the chances of a disease
- Term coined by William Kannell and Roy Dawber from the Framingham Heart Study
- Key to successful prediction of CHD: identifying important risk factors

Hypothesized CHD Risk Factors

- We will investigate risk factors collected in the first data collection for the study
 - Anonymized version of original data
- Demographic risk factors
 - *male*: sex of patient
 - age age in years at first examination
 - *education*: Some high school (1), high school/GED (2), some college/vocational school (3), college (4)

Hypothesized CHD Risk Factors

Behavioral risk factors

• currentSmoker, cigsPerDay: Smoking behavior

Medical history risk factors

- *BPmeds*: On blood pressure medication at time of first examination
- prevalentStroke: Previously had a stroke
- prevalentHyp: Currently hypertensive
- diabetes: Currently has diabetes

Hypothesized CHD Risk Factors

Risk factors from first examination

- totChol: Total cholesterol (mg/dL)
- *sysBP*: Systolic blood pressure
- diaBP: Diastolic blood pressure
- BMI: Body Mass Index, weight (kg)/height (m)²
- *heartRate*: Heart rate (beats/minute)
- glucose: Blood glucose level (mg/dL)

An Analytical Approach

Randomly split patients into training and testing sets

• Use logistic regression on training set to predict whether or not a patient experienced CHD within 10 years of first examination

• Evaluate predictive power on test set

Logistic Regression



Predicting 10-year CHD

- The independent variables are the risk factors
- The dependent variable is modeled as a binary variable
 - 1 if CHD in 10 years, 0 if not
- This is a categorical variable
 - Two (or few) possible outcomes

Logistic Regression

- Predicts the probability of 10-year CHD
 - Denote dependent variable "10-year CHD" by y
 - P(y = 1)
- Then P(y = 0) = 1 P(y = 1)
- Independent variables (risk factors): $x_1, x_2, ..., x_k$
- Uses the Logistic Response Function

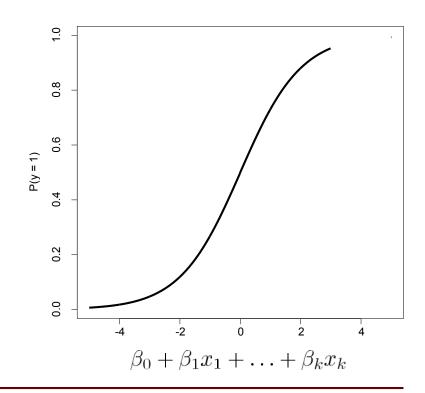
$$P(y=1) = \frac{1}{1 + e^{-(\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_k x_k)}}$$

• Nonlinear transformation of linear regression equation to produce number between 0 and 1

Understanding the Logistic Function

$$P(y=1) = \frac{1}{1 + e^{-(\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_k x_k)}}$$

- Positive values are predictive of class 1
- Negative values are predictive of class 0



Understanding the Logistic Function

$$P(y=1) = \frac{1}{1 + e^{-(\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_k x_k)}}$$

- The coefficients are selected to
 - Predict a high probability for the cases of 10-year CHD
 - Predict a low probability for the cases of no 10-year CHD

Understanding the Logistic Function

$$P(y=1) = \frac{1}{1 + e^{-(\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_k x_k)}}$$

We can instead talk about Odds (like in gambling)

Odds =
$$\frac{P(y=1)}{P(y=0)}$$

- Odds > 1 if y = 1 is more likely
- Odds < 1 if y = 0 is more likely

The Logit

It turns out that

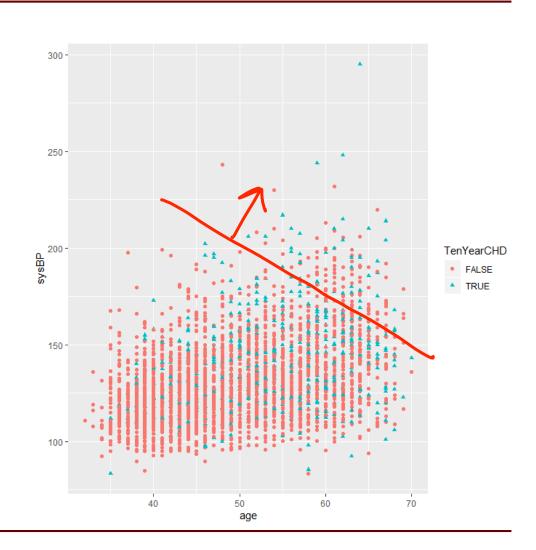
Odds =
$$e^{\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_k x_k}$$

$$log(Odds) = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \ldots + \beta_k x_k$$

- This is called the "Logit" and looks like linear regression
- The bigger the Logit is, the bigger P(y = 1)

Model for Healthcare Quality

- Plot of the independent variables
 - age
 - sysBP
- Red are no 10-year CHD
- Cyan are cases of 10year CHD



Threshold Value

- The outcome of a logistic regression model is a probability
- Often, we want to make a binary prediction
 - Will the patient incur in 10-year CHD?
- We can do this using a threshold value t
- If $P(10\text{-year CHD} = 1) \ge t$, predict CHD
- If P(10-year CHD = 1) < t, predict No CHD
- What value should we pick for t?

Threshold Value

- Often selected based on which errors are "better"
- If t is large, predict 10-year CHD rarely (when P(y=1) is large)
 - More errors where we say no risk, but there may be risk of 10-year CHD
 - Detects "accurately" patients who would most likely incur into 10-year CHD
- If t is small, predict 10-year CHD commonly (also for P(y=1) small)
 - More errors where we say 10-year CHD, but there may be no risk
 - Detects all patients who might be incurring in 10-year CHD (high recall)
- With no preference between the errors, select t = 0.5
 - Predicts the more likely outcome

Evaluation of classifiers

Selecting a Threshold Value

 Compare actual outcomes to predicted outcomes using a confusion matrix (classification matrix)

	Predicted = 0	Predicted = 1
Actual = 0	True Negatives (TN)	False Positives (FP)
Actual = 1	False Negatives (FN)	True Positives (TP)

N = number of observations

Overall accuracy =
$$(TN + TP)/N$$
 Overall error rate = $(FP + FN)/N$

Sensitivity =
$$TP/(TP + FN)$$
 = True Positive Rate

Specificity =
$$TN/(TN + FP) = 1 - False Positive Rate$$

Model Strength

- Model rarely predicts 10-year CHD risk above 50%
 - Accuracy very near a baseline of always predicting no CHD
- Model can differentiate low-risk from high-risk patients (AUC = 0.74)
- Some significant variables suggest interventions
 - Smoking
 - Cholesterol
 - Systolic blood pressure
 - Glucose

Risk Model Validation

- So far, we have used *internal validation*
 - Train with some patients, test with others
- · Weakness: unclear if model generalizes to other populations

Framingham cohort white, middle class

• Important to test on other populations

Framingham Risk Model Validation

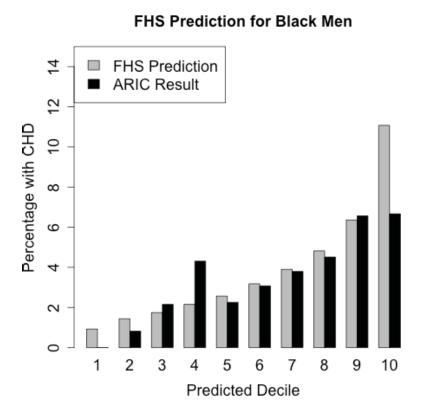
Framingham Risk Model tested on diverse cohorts

Study	Population
Atherosclerosis Risk in Communities (ARIC) Study	White and Black
Honolulu Heart Program (HHP)	Japanese American
Puerto Rico Heart Health Program (PR)	Hispanic
Strong Heart Study (SHS)	Native American

- Cohort studies collecting same risk factors
- Validation Plan
 - Predict CHD risk for each patient using FHS model
 - Compare to actual outcomes for each risk decile

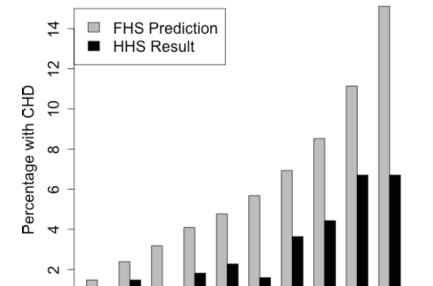
Validation for Black Men

- 1,428 black men in ARIC study
- Similar clinical characteristics, except higher diabetes rate
- Similar CHD rate
- Framingham risk model predictions accurate



Validation for Japanese American Men

- 2,755 Japanese American men in HHS
- Lower CHD rate
- Framingham risk model systematically overpredicts CHD risk



Predicted Decile

10

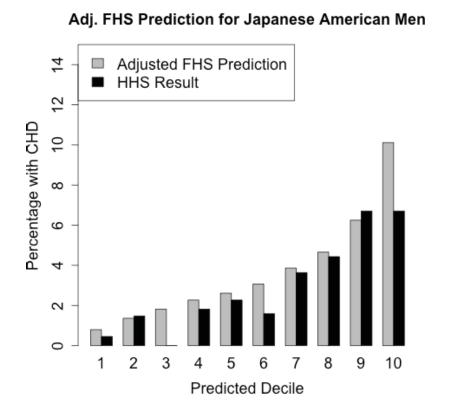
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3

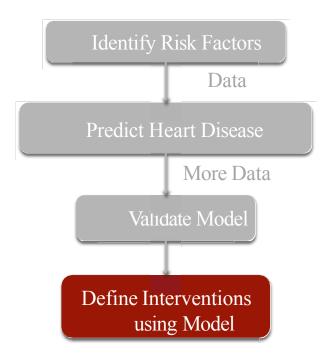
FHS Prediction for Japanese American Men

Recalibrated Model

- Recalibration adjusts model to new population
- Changes predicted risk, but does not reorder predictions
- More accurate risk estimates



Interventions



Drugs to Lower Blood Pressure

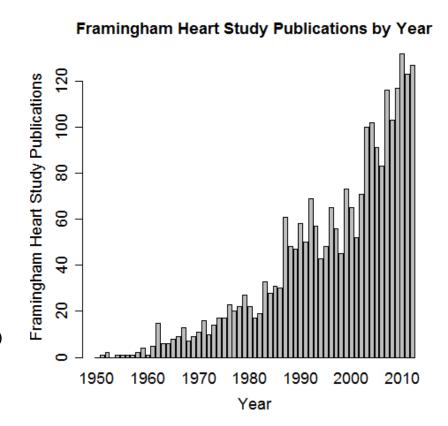
- In FDR's time, hypertension drugs too toxic for practical use
- In 1950s, the diuretic chlorothiazide was developed
- Framingham Heart Study gave Ed Freis the evidence needed to argue for testing effects of BP drugs
- Veterans Administration (VA) Trial: randomized, double blind clinical trial
- Found decreased risk of CHD
- Now,>\$1B market for diuretics worldwide

Drugs to Lower Cholesterol

- Despite Framingham results, early cholesterol drugs too toxic for practical use
- In 1970s, first statins were developed
- Study of 4,444 patients with CHD: statins cause 37% risk reduction of second heart attack
- Study of 6,595 men with high cholesterol: statins cause 32% risk reduction of CVD deaths
- Now, >\$20B market for statins worldwide

The Heart Study Through the Years

- More than 2,400 studies use Framingham data
- Many other risk factors evaluated
 - Obesity
 - Exercise
 - Psychosocial issues
 - •
- Texas Heart Institute Journal: top 10 cardiology advances of 1900s



Available Online

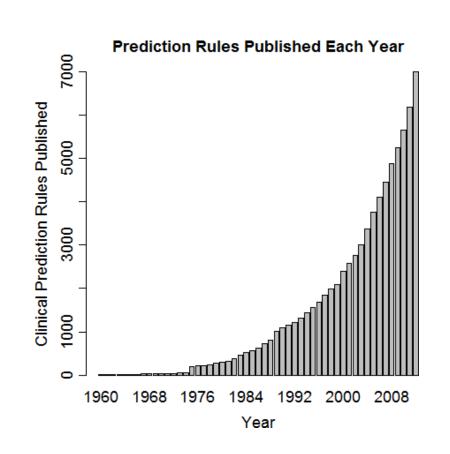
200 to 239 mg/dL 'Borderline-high.'

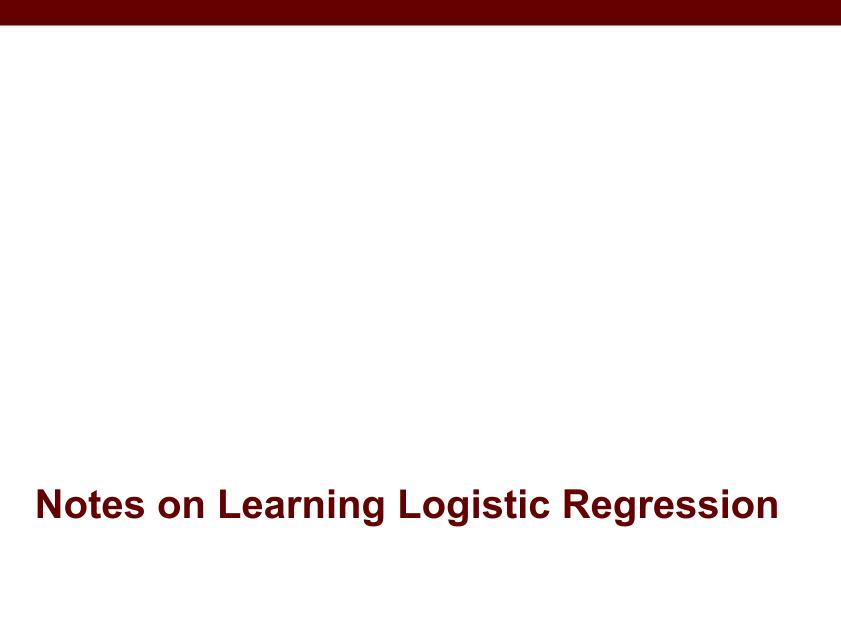
Risk Assessment Tool for Estimating Your 10-year Risk of Having a Heart Attack

The risk assessment tool below uses information from the Framingham Heart Study to predict a person's chance of having a heart attack in the next 10 years. This tool is designed for adults aged 20 and older who do not have heart disease or diabetes. To find your risk score, enter your information in the calculator below. Age: vears Gender: Female
Male mg/dL Total Cholesterol: mg/dL HDL Cholesterol: Smoker: No No Yes mm/Ha Systolic Blood Pressure: Are you currently on any medication to treat high blood pressure. No No Yes Calculate Your 10-Year Risk **€**TOP Total cholesterol - Total cholesterol is the sum of all the cholesterol in your blood. The higher your total cholesterol, the greater your risk for heart disease. Here are the total values that matter to you: Less than 200 mg/dL 'Desirable' level that puts you at lower risk for heart disease. A cholesterol level of 200 mg/dL or greater increases your risk.

Clinical Decision Rules

- Paved the way for clinical decision rules
- Predict clinical outcomes with data
 - Patient and disease characteristics
 - Test results
- More than 75,000 published across medicine
- Rate increasing





Maximum Likelihood Estimation

- We use maximum likelihood estimation (MLE) to estimate the parameters of the logistic regression
- The labels that we are predicting are binary, and the output of our logistic regression function is supposed to be the probability that the label is one
- This means that we can (and should) interpret each label as a Bernoulli random variable: Y ~ Ber(p) where $p = \sigma(\theta^T x)$
- We write then the probability of one data point as:

$$P(Y = y | X = \mathbf{x}) = \sigma(\theta^T \mathbf{x})^y \cdot \left[1 - \sigma(\theta^T \mathbf{x})\right]^{(1-y)}$$

Maximum Likelihood Estimation

So the likelihood of all data is:

$$L(\theta) = \prod_{i=1}^{n} P(Y = y^{(i)} \mid X = \mathbf{x}^{(i)})$$
$$= \prod_{i=1}^{n} \sigma(\theta^{T} \mathbf{x}^{(i)})^{y^{(i)}} \cdot \left[1 - \sigma(\theta^{T} \mathbf{x}^{(i)})\right]^{(1-y^{(i)})}$$

Taking the log, we get the log likelihood for logistic regression

$$LL(\theta) = \sum_{i=1}^{n} y^{(i)} \log \sigma(\theta^T \mathbf{x}^{(i)}) + (1 - y^{(i)}) \log[1 - \sigma(\theta^T \mathbf{x}^{(i)})]$$

- to maximize with numerical techniques such as
 - Line Search
 - Simulated Annealing
 - Gradient Descent
 - Newton's Method

Regularization

 It would still be appropriate to regularize the optimization with an L2 (or L1) loss on the parameter values, to penalize very large coefficients and avoid overfitting:

$$cost = -\sum_{i=1}^{n} \{y^{(i)} \log \sigma(\theta^{T} \mathbf{x}^{(i)}) + (1 - y^{(i)}) \log[1 - \sigma(\theta^{T} \mathbf{x}^{(i)})]\} + \lambda \sum_{i=1}^{n} \theta^{2}$$

Thank you

Slide credits:

- MIT open course ware
- Prof. Alessandro Panconesi
- Will Monroe's lecture notes at Stanford
- Prof. Fabio Galasso