P8160 - Less is More: Comparing Logistic and Lasso-Logistic Regression in Breast Cancer Diagnosis

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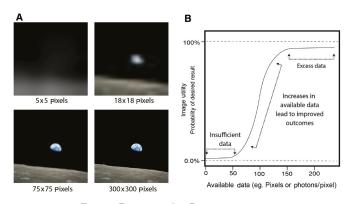
Motivation

As breast cancer is one of the most common kinds of cancer in the United States, great efforts have been made to aid in early and accurate detection.

Improvements in tumor imaging technology used in screening procedures have allow us access to more data than ever before, ideally to construct better ways to evaluate disease severity.

However... data \neq information.

Imaging Data and Information Overload



From: Duncan, J., Diagnosis, 2017

Goal

We want to investigate two questions:

Does having more data always correspond to an advantage in diagnosis prediction?

Can we reduce the amount of information we need to collect while maintaining (or increasing) predictive power?

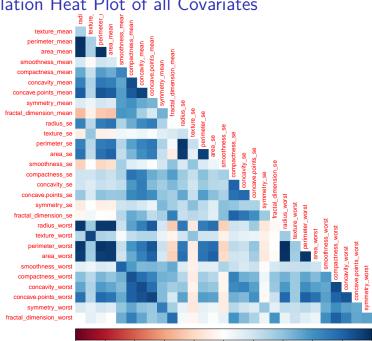
Toward this end, we will develop and evaluate the performance of two predictive models.

- Newton-Raphson Algorithm (Full Model)
- ► Logistic LASSO Algorithm (Optimal Model)

Data

- ▶ 569 rows and 31 columns related to breast tissue images
- Outcome of interest: Diagnosis (B or M)
 - ▶ 357 benign (B) cases and 212 malignant (M) cases
- The available predictors include summary statistics for tumor characteristics such as radius, texture, perimeter, area, smoothness, compactness, concavity, concave points, symmetry, and fractal dimension.

Correlation Heat Plot of all Covariates

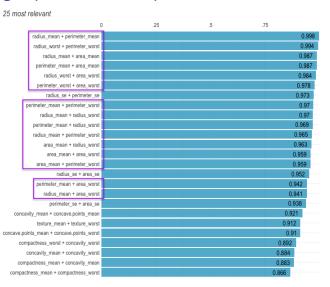


Ranking Cross-Correlations

25 most relevant

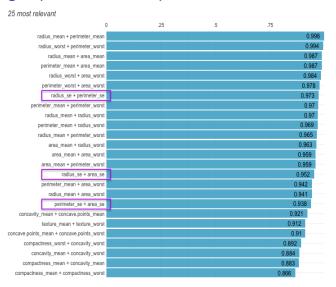
	 25	.5	.75
radius_mean + perimeter_mean			0.998
radius_worst + perimeter_worst			0.994
radius_mean + area_mean			0.987
perimeter_mean + area_mean			0.987
radius_worst + area_worst			0.984
perimeter_worst + area_worst			0.978
radius_se + perimeter_se			0.973
perimeter_mean + perimeter_worst			0.97
radius_mean + radius_worst			0.97
perimeter_mean + radius_worst			0.969
radius_mean + perimeter_worst			0.965
area_mean + radius_worst			0.963
area_mean + area_worst			0.959
area_mean + perimeter_worst			0.959
radius_se + area_se			0.952
perimeter_mean + area_worst			0.942
radius_mean + area_worst			0.941
perimeter_se + area_se			0.938
concavity_mean + concave.points_mean			0.921
texture_mean + texture_worst			0.912
concave.points_mean + concave.points_worst			0.91
compactness_worst + concavity_worst			0.892
concavity_mean + concavity_worst			0.884
compactness_mean + concavity_mean			0.883
compactness_mean + compactness_worst			0.866

Identifying Equivalence Groups



Best Proxy: radius_worst

Identifying Equivalence Groups



Best Proxy: radius_se

Selected Variables

	Diagnosis	Received		
Variable	B , N = 357 ¹	M , N = 212 ⁷	p-value ²	
texture_mean	17.91 (4.00)	21.60 (3.78)	<0.001	
smoothness_mean	0.09 (0.01)	0.10 (0.01)	<0.001	
compactness_mean	0.08 (0.03)	0.15 (0.05)	<0.001	
concave points_mean	0.03 (0.02)	0.09 (0.03)	<0.001	
symmetry_mean	0.17 (0.02)	0.19 (0.03)	<0.001	
fractal_dimension_mean	0.06 (0.01)	0.06 (0.01)	0.5	
radius_se	0.28 (0.11)	0.61 (0.35)	<0.001	
texture_se	1.22 (0.59)	1.21 (0.48)	0.6	
smoothness_se	0.01 (0.00)	0.01 (0.00)	0.2	
compactness_se	0.02 (0.02)	0.03 (0.02)	<0.001	
concavity_se	0.03 (0.03)	0.04 (0.02)	<0.001	
concave points_se	0.01 (0.01)	0.02 (0.01)	<0.001	
symmetry_se	0.02 (0.01)	0.02 (0.01)	0.028	
fractal_dimension_se	0.00 (0.00)	0.00 (0.00)	<0.001	
radius_worst	13.38 (1.98)	21.13 (4.28)	<0.001	
smoothness_worst	0.12 (0.02)	0.14 (0.02)	<0.001	
compactness_worst	0.18 (0.09)	0.37 (0.17)	<0.001	
concavity_worst	0.17 (0.14)	0.45 (0.18)	<0.001	
symmetry_worst	0.27 (0.04)	0.32 (0.07)	<0.001	
fractal_dimension_worst	0.08 (0.01)	0.09 (0.02)	<0.001	
⁷ Statistics presented: Mean (SD)				

² Statistical tests performed: Wilcoxon rank-sum test

Full Model: Newton-Raphson

Consider the following log-likelihood, gradient, and hessian matrix. First, let

$$\pi_i = P(Y_i = 1 | x_{i,1}, \dots x_{i,p}) = \frac{e^{\beta_0 + \sum_{j=1}^p \beta_j x_{i,j}}}{1 + e^{\beta_0 + \sum_{j=1}^p \beta_j x_{i,j}}}.$$

The log-likelihood:

$$I(\mathbf{X}|\vec{\beta}) = \sum_{i=1}^{n} \left[y_i \left(\beta_0 + \sum_{i=1}^{p} \beta_j x_{i,j} \right) - \log \left(1 + \exp \left(\beta_0 + \sum_{i=1}^{p} \beta_j x_{i,j} \right) \right) \right]$$

The gradient

$$\nabla I(\mathbf{X}|\vec{\beta}) = \begin{bmatrix} \sum^{n} y_{i} - \pi_{i} & \sum^{n} x_{i,1}(y_{i} - \pi_{i}) & \dots & \sum^{n} x_{i,p}(y_{i} - \pi_{i}) \end{bmatrix}_{1 \times (p+1)}^{T}$$
The hessian: produces a matrix $(p+1 \times p+1)$

$$abla^2 I(\mathbf{X}|ec{eta}) = -\sum_{i=1}^n egin{pmatrix} 1 \ X \end{pmatrix} ig(1 \quad Xig) \, \pi_i (1-\pi_i)$$

Optimal Model: Logistic LASSO

For vector $\alpha \in \mathbb{R}^{p+1}$, define $g: \mathbb{R}^{p+1} \to \mathbb{R}$ to be

$$g(\beta) \equiv -\frac{1}{2n}\sum_{i=1}^n w_i(z_i - \mathbf{X}_i^t\beta)^2 + O(\alpha),$$

the Taylor expansion of our log-likelihood centered around lpha, where

$$z_i \equiv \mathbf{X}_i^t \alpha + \frac{y_i - \pi_i}{w_i},$$
 (effective response) $w_i \equiv \pi_i (1 - \pi_i),$ and (effective weights) $\pi_i \equiv \frac{e^{\mathbf{X}_i^t \alpha}}{1 + e^{\mathbf{X}_i^t \alpha}}$

for $i \in \{1, ..., n\}$.

Optimal Model: Logistic LASSO

It follows that for any $\lambda \in \mathbb{R}_+$,

$$\operatorname*{arg\,min}_{\beta_k \in \mathbb{R}} \left\{ g(\boldsymbol{\beta}) + \lambda \sum_{j=1}^p |\beta_j| \right\} = S\left(\hat{\beta}_k, \lambda_k\right), \text{ where }$$

$$\hat{\beta}_k \equiv \left(\sum_{i=1}^n w_i x_{ik}^2\right)^{-1} \sum_{i=1}^n w_i x_{ik} \left(z_i - \sum_{j \neq k} \beta_j x_{ij}\right),$$

$$\lambda_k \equiv \left(\frac{1}{n} \sum_{i=1}^n w_i x_{ik}^2\right)^{-1} \lambda,$$

and S is the soft-thresholding (or *shrinkage*) function. This is analogous to a penalized, weighted Gaussian regression.

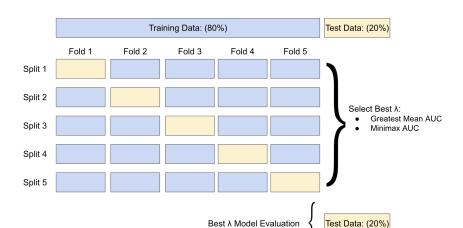
Optimal Model: Logistic LASSO

Our coordinate descent algorithm proceeds as follows.

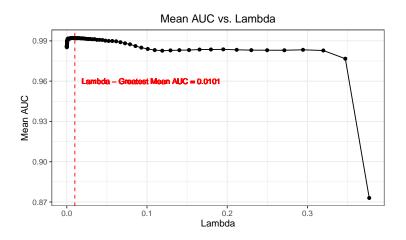
- ▶ Outer Loop: Decrement over $\lambda \in (\lambda_{max}, \dots, \lambda_{min})$
- ▶ Middle Loop: Update $\alpha = \beta$ and Taylor expand g around α .
- Inner Loop: Update $\beta_k = S\left(\hat{\beta}_k, \lambda_k\right)$ sequentially for $k \in \{0, 1, \dots, p, 0, 1, \dots, p, 0, 1, \dots\}$ until convergence.

Note: the middle loop terminates when a given Taylor expansion no longer yields updates (within the specified tolerance) to β in the inner loop.

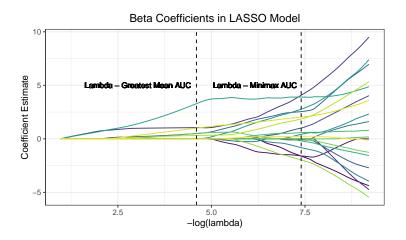
5-Fold Cross Validation



Cross Validation Results: Selecting Best Lambda



Cross Validation Results: LASSO Coefficients



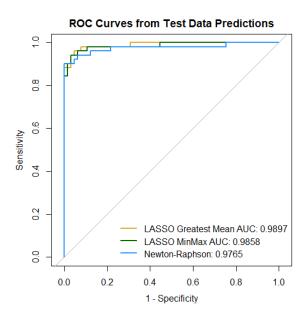
Coefficients Comparison: All Estimates

	NewtonRaphson	LASSO_GreatestMeanAUC	LASSO_MinimaxAUC
intercept	-0.0881	-0.9302	-1.2291
texture_mean	8.2691	0.9994	2.0355
smoothness_mean	-2.5447	0.0000	-0.1919
compactness_mean	-10.9297	0.0000	-1.6093
concave points_mean	20.9342	1.0512	4.0737
symmetry_mean	-1.9716	0.0000	0.0000
fractal_dimension_mean	3.9221	0.0000	0.0000
radius_se	18.5308	0.0000	2.7696
texture_se	-2.0504	0.0000	0.2269
smoothness_se	1.2753	0.0000	0.5802
compactness_se	4.7358	0.0000	-1.5925
concavity_se	-6.4914	0.0000	-0.0893
concave points_se	8.9092	0.0000	1.0126
symmetry_se	-13.8273	0.0000	-1.9717
fractal_dimension_se	-12.2152	0.0000	-0.8241
radius_worst	9.2654	3.2848	3.9047
smoothness_worst	0.6705	0.4955	0.0000
compactness_worst	-14.3912	0.0000	0.0000
concavity_worst	14.9611	0.4943	2.5436
symmetry_worst	12.3102	0.3112	1.8506
fractal_dimension_worst	7.7655	0.0000	0.4242

Coefficients Comparison: Model Performance

Measures	NewtonRaphson	LASSO_GreatestMeanAUC	LASSO_MinimaxAUC
Specificities	0.2462	0.6923	0.5538
AUC	0.9765	0.9897	0.9858
Selected Lambda	0	0.0101	0.0006
Number of Variables (w/o Intercept)	20	6	16

ROC Plot



Discussion

- Lasso-logistic model, with fewer predictors, out-performed the logistic model with all selected predictors.
 - however, full model was not bad (AUC > 0.96)
- ▶ Ideal performance is to accurately classify every patient
- Balancing sensitivity and specificity
 - ► False positives vs false negatives
 - Decision boundaries
- ► Future work

Resources

Duncan, J. R. (2017, September 1). Information overload: When less is more in medical imaging. De Gruyter. https://www.degruyter.com/document/doi/10.1515/dx-2017-0008/html?lang=en

Cancer Stat Facts: Female Breast Cancer. *National Cancer Institute - NIH* https://seer.cancer.gov/statfacts/html/breast.html

American Cancer Society. (2019). Breast cancer facts & figures 2019–2020. Am Cancer Soc, 1-44.