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

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2022-03-28

### Motivation

#### According to NIH, there have been an estimated:

- ▶ 281,550 new cases of breast cancer in women in 2021,
- ▶ 43,600 breast cancer in women related deaths in 2021.

### American Cancer Society Guideline for Breast Cancer Screening:

- ▶ Women between ages 25-40 should have an annual clinical breast examination.
- Women between ages 40-44 should begin annual screening via mammogram
- Women between ages 45-54 should screened annually via mammogram

# Imaging Data and Information Overload

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

### Goal

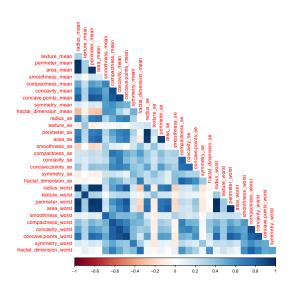
Toward this end, we want to develop and evaluate the performance of two predictive models.

- 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 measurements such as radius, texture, perimeter, area, smoothness, compactness, concavity, concave points, symmetry, and fractal dimension.

### Correlation Heat Plot of all Covariates

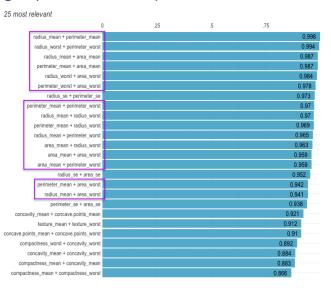


# Ranking Cross-Correlations

#### 25 most relevant

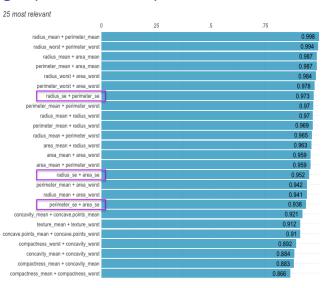
	 25	.5	.75
radius_mean + perimeter_mean			0.998
radius_worst + perimeter_worst			0.994
radius_mean + area_mean			0.987
perimeter_mean + area_mean			0.987
radius_worst + area_worst			0.984
perimeter_worst + area_worst			0.978
radius_se + perimeter_se			0.973
perimeter_mean + perimeter_worst			0.97
radius_mean + 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

# Finally Selected Variables

	Diagnosis			
Variable	<b>B</b> , N = 357 <sup>1</sup>	M, N = 212 <sup>1</sup>	p-value <sup>2</sup>	
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	
<sup>†</sup> Statistics presented: Mean (SD) <sup>2</sup> 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:

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

#### <del>-</del>. . .

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

Lemma 1. Consider the optimization problem

$$\min_{x \in \mathbb{R}} \left\{ \frac{1}{2} (x - b)^2 + c|x| \right\}$$

for  $b \in \mathbb{R}$  and  $c \in \mathbb{R}_{++}$ . It follows that the minimizer is given by

$$\hat{x} = S(b, c),$$

where S is the soft-thresholding operator.

Lemma 2. Consider the optimization problem

$$\min_{\beta_k \in \mathbb{R}} \left\{ \frac{1}{2n} \sum_{i=1}^n w_i \left( z_i - \sum_{j=1}^p \beta_j x_{ij} \right)^2 \right\}$$

for some  $k \in \{1, \dots, p\}$ . It follows that the minimizer is given by

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

# Optimal Model (Logistic LASSO)

**Lemma 3.** With  $\hat{\beta}_k$  defined as above,

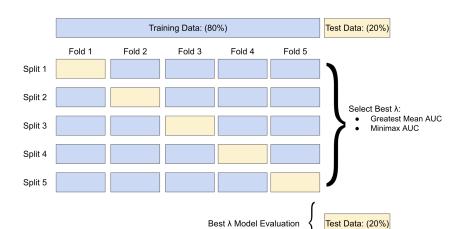
$$\min_{\beta_k \in \mathbb{R}} \left\{ \frac{1}{2n} \sum_{i=1}^n w_i \left( z_i - \sum_{j=1}^p \beta_j x_{ij} \right)^2 + \lambda \sum_{j=1}^p |\beta_j| \right\}$$

$$= \min_{\beta_k \in \mathbb{R}} \left\{ \frac{1}{2} (\beta_k - \hat{\beta}_k)^2 + \left( \frac{1}{n} \sum_{i=1}^n w_i x_{ik}^2 \right)^{-1} \lambda |\beta_k| \right\}.$$

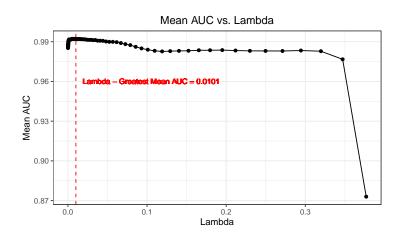
By Lemma 1 and Lemma 3,

$$\begin{aligned} & \underset{\beta_k \in \mathbb{R}}{\arg\min} \left\{ \frac{1}{2n} \sum_{i=1}^n w_i \left( z_i - \sum_{j=1}^p \beta_j x_{ij} \right)^2 + \lambda \sum_{j=1}^p |\beta_j| \right\} \\ &= S\left( \hat{\beta}_k, \left( \frac{1}{n} \sum_{i=1}^n w_i x_{ik}^2 \right)^{-1} \lambda \right) \end{aligned}$$

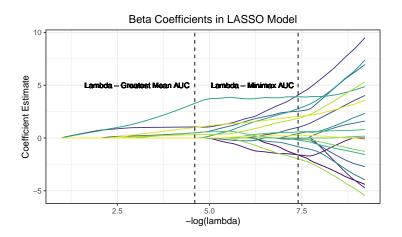
### 5-Fold Cross Validation



### Cross Validation Results: Selecting Best Lambda



### Cross Validation Results: LASSO Coefficients



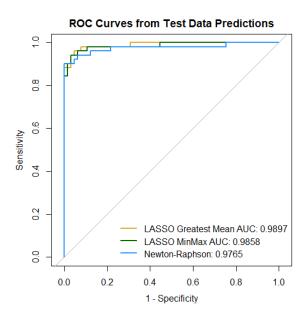
### Coefficients Comparison: All Estimates

	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

- 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. Retrieved March 27, 2022, from

 $https://www.degruyter.com/document/doi/10.1515/dx-2017-0008/html?lang{=}en$ 

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.