

Algorithms and Inference

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Statistical Learning



Algorithms and inference

- Suppose we have observed y_1, \dots, y_n , realizations of Y_1, \dots, Y_n i.i.d. Y , and our interest is on $\mathbb{E}(Y) = \mu$
- Averaging is the *algorithm*

$$\bar{y} = \frac{1}{n} \sum_{i=1}^n y_i$$

- The standard error provides an *inference* on the algorithm's accuracy

$$\widehat{\text{se}} = \sqrt{\frac{1}{n} \frac{\sum_{i=1}^n (y_i - \bar{y})^2}{n-1}}$$

- “It is a surprising, and crucial, aspect of statistical theory that the same data that supplies an estimate can also assess its accuracy” (Efron and Hastie, 2016)



Gaussian model

Model

Y_1, \dots, Y_n i.i.d. $Y \sim N(\mu, \sigma^2)$

μ is the *parameter of interest*

σ^2 is the *nuisance parameter*

Estimator and its standard error

$\bar{Y} \sim N(\mu, \sigma^2/n)$

$\text{se}(\bar{Y}) = \sqrt{\text{Var}(\bar{Y})} = \sigma\sqrt{1/n}$

Estimator of the standard error

$\hat{\text{se}}(\bar{Y}) = \hat{\sigma}\sqrt{1/n}$

$$\hat{\sigma}^2 = \frac{\sum_{i=1}^n (Y_i - \bar{Y})^2}{n-1} \sim \sigma^2 \chi_{n-1}^2 / (n-1)$$


Confidence interval

Pivotal statistic

$$T = \frac{\bar{Y} - \mu}{\sigma \sqrt{1/n}} \cdot \frac{\sigma}{\hat{\sigma}} \sim \frac{N(0, 1)}{\sqrt{\chi_{n-1}^2/(n-1)}} \sim t_{n-1}$$

with $\Pr(-t_{n-1}^{1-\alpha/2} \leq T \leq t_{n-1}^{1-\alpha/2}) = 1 - \alpha$

where $t_{n-1}^{1-\alpha/2}$ is the $1 - \alpha/2$ quantile of the Student t distribution with $n - 1$ degrees of freedom

$1 - \alpha$ **confidence interval for μ**

$$[\underline{\mu}, \bar{\mu}] = \bar{Y} \pm t_{n-1}^{1-\alpha/2} \cdot \hat{\text{se}}(\bar{Y})$$

Coverage

$$\Pr([\underline{\mu}, \bar{\mu}] \ni \mu) = 1 - \alpha$$



Simulation

```
sim = function(n=25, mu=0, sigma=1, alpha=0.05){  
  ys = rnorm(n, mean=mu, sd=sigma)  
  bary = mean(ys)  
  hatse = sqrt( var(ys) / n )  
  k = qt(alpha/2, df = n-1, lower.tail = F)  
  ci = bary + c(-1,1) * k * hatse  
  cover = (mu >= ci[1] & mu <= ci[2])  
  return(cover)  
}
```

```
set.seed(123)  
B = 1000  
mean( replicate(B, sim(n=25) ) )
```



Outline

① Leukemia data

② Kidney data



Leukemia data

- $n = 72$ leukemia patients: 45 with ALL (acute lymphoblastic leukemia) and 27 with AML (acute myeloid leukemia, a worse prognosis)
- Each patient has genetic activity measured for $p = 7128$ genes
- The histograms in the next slide compare the genetic activities in the two groups for gene 136
- Is the perceived difference genuine, or perhaps, as people like to say, “a statistical fluke”?



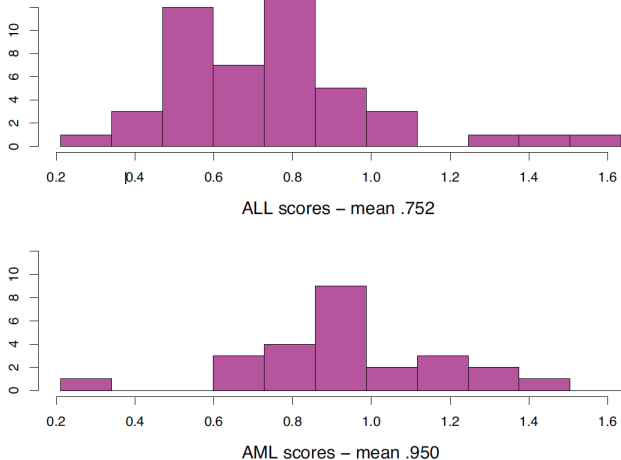


Figure 1.4 Scores for gene 136, leukemia data. Top **ALL** ($n = 47$), bottom **AML** ($n = 25$). A two-sample t -statistic = 3.01 with p -value = .0036.



Hypothesis testing

- The classic answer to this question is via a two-sample t -statistic

$$t = \frac{\overline{\text{AML}} - \overline{\text{ALL}}}{\hat{\text{sd}}}$$

- Compare the observed value $t = 3.01$ with the null distribution, i.e. Student's t distribution with 70 degrees of freedom
- The p -value is 0.0036. A small p -value is a statement of statistical surprise: something very unusual has happened if in fact there is no difference in gene 136 expression between ALL and AML patients
- We are less surprised by $t = 3.01$ or $p = 0.0036$ if gene 136 is just one out of thousands candidates that might have produced “interesting” results
- The next slide shows the histogram of the two-sample t -statistics for the 7178 genes. Now $t = 3.01$ looks less unusual; 400 other genes have t exceeding 3.01, about 5.6% of them



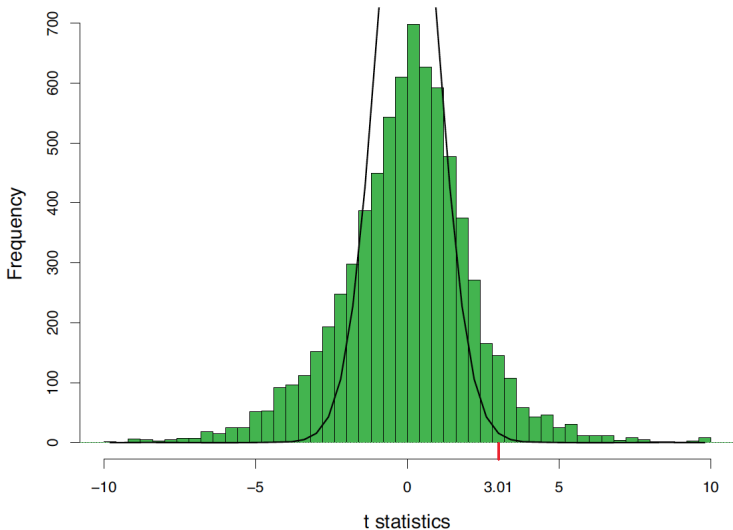


Figure 1.5 Two-sample t -statistics for 7128 genes, leukemia data. The smooth curve is the theoretical null density for the t -statistic.



Outline

① Leukemia data

② Kidney data



Kidney data

- Kidney function generally declines with age. The rate of decline is an important question in kidney transplantation: in the past, potential donors past age 60 were prohibited
- $Y = \text{tot}$ (kidney function overall score)
- $X = \text{age}$ (age in years)
- $(x_1, y_1), \dots, (x_n, y_n)$ for $n = 157$ healthy volunteers



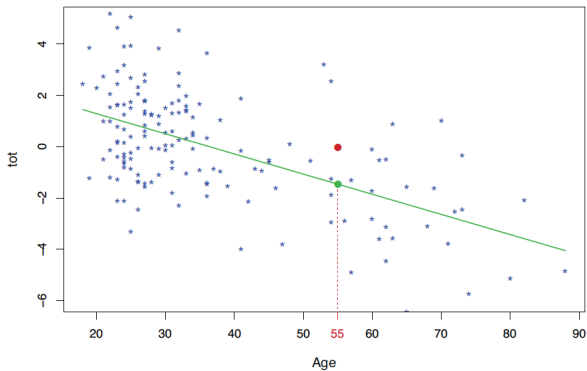


Figure 8.1 Kidney data; a new volunteer donor is aged 55.
Which prediction is preferred for his kidney function?

Source: Efron and Hastie (2016)



Linear model

Model

$$\mathbf{y} = \mathbf{X}\boldsymbol{\beta} + \boldsymbol{\epsilon}, \quad \boldsymbol{\epsilon} \sim \mathcal{N}(\mathbf{0}, \sigma^2 \mathbf{I}_n)$$

$\mathbf{x}^\top = (x_1, \dots, x_p)$: values of interest

$\mu_x = \mathbf{x}^\top \boldsymbol{\beta}$: parameter of interest

Estimator and its standard error

$$\hat{\mu}_x = \mathbf{x}^\top \hat{\boldsymbol{\beta}} = \mathbf{x}^\top (\mathbf{X}^\top \mathbf{X})^{-1} \mathbf{X}^\top \mathbf{y} \sim N(\mathbf{x}^\top \boldsymbol{\beta}, \sigma^2 \mathbf{x}^\top (\mathbf{X}^\top \mathbf{X})^{-1} \mathbf{x})$$

$$\text{se}(\hat{\mu}_x) = \sigma \sqrt{\mathbf{x}^\top (\mathbf{X}^\top \mathbf{X})^{-1} \mathbf{x}}$$

Estimator of the standard error

$$\hat{\text{se}}(\hat{\mu}_x) = \sigma \sqrt{\mathbf{x}^\top (\mathbf{X}^\top \mathbf{X})^{-1} \mathbf{x}}$$

$$\hat{\sigma}^2 = (\mathbf{y} - \mathbf{X}\hat{\boldsymbol{\beta}})^\top (\mathbf{y} - \mathbf{X}\hat{\boldsymbol{\beta}}) / (n - p) \sim \sigma^2 \chi_{n-p}^2 / (n - p)$$



Confidence interval

Pivotal statistic

$$T = \frac{\hat{\mu}_x - \mu_x}{\hat{\text{se}}(\hat{\mu}_x)} \cdot \frac{\sigma}{\hat{\sigma}} \sim \frac{\mathcal{N}(0, 1)}{\sqrt{\chi_{n-p}^2/(n-p)}} \sim t_{n-p}$$

$$\text{with } \Pr(-t_{n-p}^{1-\alpha/2} \leq T \leq t_{n-p}^{1-\alpha/2}) = 1 - \alpha$$

$1 - \alpha$ **confidence interval for μ_x**

$$[\underline{\mu}_x, \bar{\mu}_x] = \hat{\mu}_x \pm t_{n-p}^{1-\alpha/2} \cdot \hat{\text{se}}(\hat{\mu}_x)$$

Coverage

$$\Pr([\underline{\mu}_x, \bar{\mu}_x] \ni \mu_x) = 1 - \alpha$$



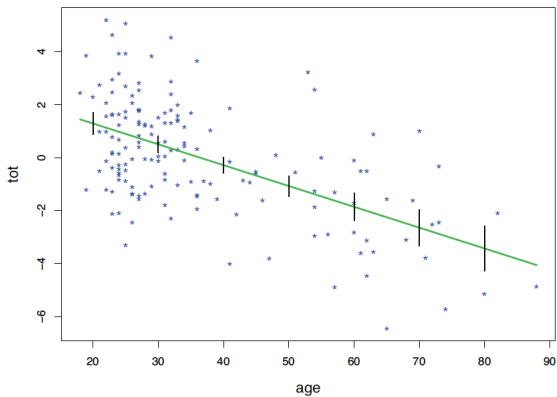


Figure 1.1 Kidney fitness `tot` vs `age` for 157 volunteers. The line is a linear regression fit, showing ± 2 standard errors at selected values of `age`.

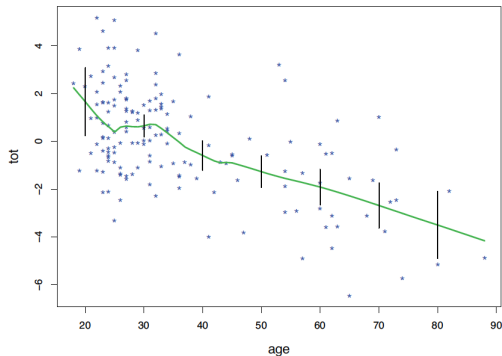
Source: Efron and Hastie (2016)



Bootstrap standard errors

Table 1.1 Regression analysis of the kidney data; (1) linear regression estimates; (2) their standard errors; (3) **lowess** estimates; (4) their bootstrap standard errors.

| age | 20 | 30 | 40 | 50 | 60 | 70 | 80 |
|------------------------|------|-----|------|-------|-------|-------|-------|
| 1. linear regression | 1.29 | .50 | -.28 | -1.07 | -1.86 | -2.64 | -3.43 |
| 2. std error | .21 | .15 | .15 | .19 | .26 | .34 | .42 |
| 3. lowess | 1.66 | .65 | -.59 | -1.27 | -1.91 | -2.68 | -3.50 |
| 4. bootstrap std error | .71 | .23 | .31 | .32 | .37 | .47 | .70 |



Bootstrap replications

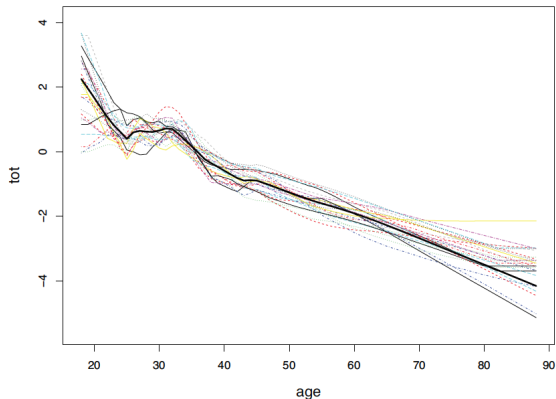


Figure 1.3 25 bootstrap replications of `lowess(x, y, 1/3)`.

Source: Efron and Hastie (2016)

