R01 - Simple linear regression

STAT 5870 (Engineering) Iowa State University

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Telomere length

http://www.pnas.org/content/101/49/17312

People who are stressed over long periods tend to look haggard, and it is commonly thought that psychological stress leads to premature aging [as measured by decreased telomere length]

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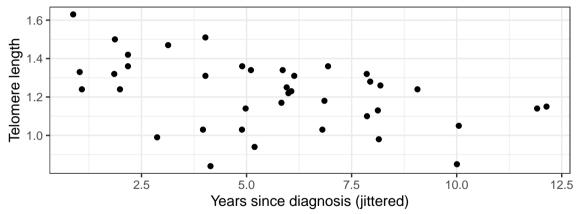
examine the importance of ... caregiving stress (...number of years since a child's diagnosis [of a chronic disease]) [on telomere length]

..

Telomere length values were measured from DNA by a quantitative PCR assay that determines the relative ratio of telomere repeat copy number to single-copy gene copy number (T/S ratio) in experimental samples as compared with a reference DNA sample.

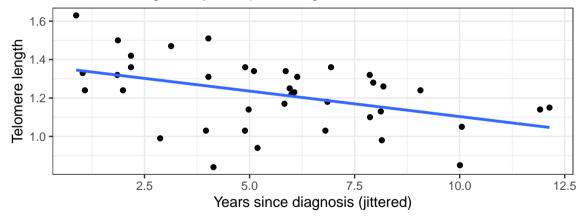
Data

Telomere length vs years post diagnosis



Data with regression line

Telomere length vs years post diagnosis



Simple Linear Regression

The simple linear regression model is

$$Y_i \stackrel{ind}{\sim} N(\beta_0 + \beta_1 X_i, \sigma^2)$$

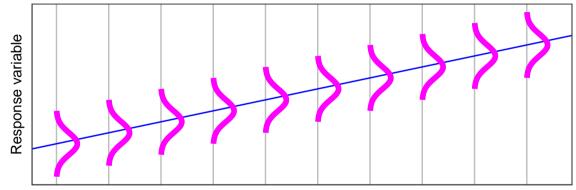
where Y_i and X_i are the response and explanatory variable, respectively, for individual i.

Terminology (all of these are equivalent):

response	explanatory
outcome	covariate
dependent	independent
endogenous	exogenous

Simple linear regression - visualized

Simple linear regression model



Explanatory variable

Parameter interpretation

Recall:

$$E[Y_i|X_i = x] = \beta_0 + \beta_1 x \qquad Var[Y_i|X_i = x] = \sigma^2$$

- If $X_i = 0$, then $E[Y_i | X_i = 0] = \beta_0$. β_0 is the expected response when the explanatory variable is zero.
- If X_i increases from x to x+1, then

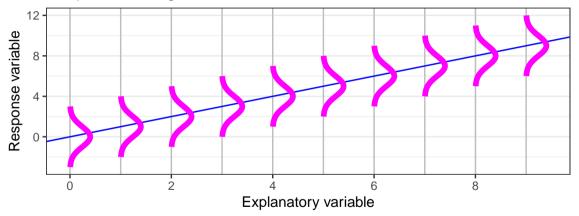
$$E[Y_i|X_i = x + 1] = \beta_0 + \beta_1 x + \beta_1 - E[Y_i|X_i = x] = \beta_0 + \beta_1 x = \beta_1$$

 β_1 is the expected increase in the response for each unit increase in the explanatory variable.

ullet σ is the standard deviation of the response for a fixed value of the explanatory variable.

Simple linear regression - visualized

Simple linear regression model



Errors v residuals

Remove the mean:

$$Y_i = \beta_0 + \beta_1 X_i + e_i$$
 $e_i \stackrel{iid}{\sim} N(0, \sigma^2)$

So the error is

$$e_i = Y_i - (\beta_0 + \beta_1 X_i)$$

which we approximate by the residual

$$r_i = \hat{e}_i = Y_i - (\hat{\beta}_0 + \hat{\beta}_1 X_i)$$

These residuals we will use for a number of purposes including

- assessing model assumptions,
- identifying outliers, and
- estimating error variance.

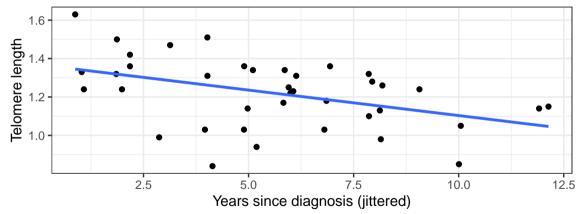
Estimators

The least squares (minimize $\sum_{i=1}^{n} r_i^2$), maximum likelihood, and Bayesian estimators (prior $1/\sigma^2$) are

$$\hat{\beta}_{1} = SXY/SXX
\hat{\beta}_{0} = \overline{Y} - \hat{\beta}_{1}\overline{X}
\hat{\sigma}^{2} = SSE/(n-2) df = n-2
\overline{X} = \frac{1}{n} \sum_{i=1}^{n} X_{i}
\overline{Y} = \frac{1}{n} \sum_{i=1}^{n} Y_{i}
SXY = \sum_{i=1}^{n} (X_{i} - \overline{X})(Y_{i} - \overline{Y})
SXX = \sum_{i=1}^{n} (X_{i} - \overline{X})(X_{i} - \overline{X}) = \sum_{i=1}^{n} (X_{i} - \overline{X})^{2}
SSE = \sum_{i=1}^{n} r_{i}^{2}$$

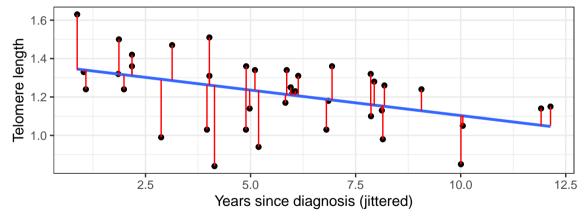
Residuals

Telomere length vs years post diagnosis



Residuals

Telomere length vs years post diagnosis



How certain are we about $\hat{\beta}_0$ and $\hat{\beta}_1$?

We quantify this uncertainty using their standard errors (or posterior scale parameters):

$$\begin{array}{ll} SE(\hat{\beta}_0) &= \hat{\sigma}\sqrt{\frac{1}{n} + \frac{\overline{X}^2}{(n-1)s_X^2}} & df = n-2 \\ SE(\hat{\beta}_1) &= \hat{\sigma}\sqrt{\frac{1}{(n-1)s_X^2}} & df = n-2 \\ \\ s_X^2 &= SXX/(n-1) \\ s_Y^2 &= SYY/(n-1) \\ SYY &= \sum_{i=1}^n (Y_i - \overline{Y})^2 \\ \\ r_{XY} &= \frac{SXY/(n-1)}{s_X s_Y} & \text{correlation coefficient} \\ R^2 &= r_{XY}^2 = \frac{SST - SSE}{SST} & \text{coefficient of determination} \\ SST &= SYY = \sum_{i=1}^n (Y_i - \overline{Y})^2 \end{array}$$

The coefficient of determination (R^2) is the proportion of the total response variation explained by the model.

Default Bayesian analysis of the simple linear regression model

If we assume the default prior $p(\beta_0, \beta_1, \sigma^2) \propto 1/\sigma^2$, then the marginal posteriors for the mean parameters are

$$\beta_j | y \sim t_{n-2}(\hat{\beta}_j, SE(\hat{\beta}_j)^2).$$

We can construct a 100(1-a)% two-sided credible interval for β_j via

$$\hat{\beta}_j \pm t_{n-2,1-a/2} SE(\hat{\beta}_j)$$

where $P(T_{n-2} < t_{n-2,1-a/2}) = 1 - a/2$ for $T_{n-2} \sim t_{n-2}$.

We can compute posterior probabilities via

$$P(\beta_j > b_j | y) = P\left(T_{n-2} > \frac{b_j - \hat{\beta}_j}{SE(\hat{\beta}_j)}\right) \quad \text{or} \quad P(\beta_j < b_j | y) = P\left(T_{n-2} < \frac{b_j - \hat{\beta}_j}{SE(\hat{\beta}_j)}\right)$$

often $b_i = 0$.

p-values and confidence interval

We can construct a 100(1-a)% two-sided confidence interval for β_i via

$$\hat{\beta}_j \pm t_{n-2,1-a/2} SE(\hat{\beta}_j).$$

We can compute one-sided p-values,

$$H_0: eta_j \geq b_j$$
 vs $H_A: eta_j < b_j$ has

and $H_0: \beta_j \leq b_j$ vs $H_A: \beta_j > b_j$ has

$$p$$
-value $=P\left(T_{n-2}<rac{\hat{eta}_j-b_j}{SE(\hat{eta}_j)}
ight)$

$$p ext{-value} = P\left(T_{n-2} > rac{\hat{eta}_j - b_j}{SE(\hat{eta}_j)}
ight)$$

or two-sided p-values for $H_0: \beta_j = b_j$ vs $H_A: \beta_j \neq b_j$:

$$=2\times\min\left\{P\left(T_{n-2}>\frac{\hat{\beta}_j-b_j}{SE(\hat{\beta}_j)}\right),P\left(T_{n-2}<\frac{\hat{\beta}_j-b_j}{SE(\hat{\beta}_j)}\right)\right\}\\ =2\times P\left(T_{n-2}<-\left|\frac{\hat{\beta}_j-b_j}{SE(\hat{\beta}_j)}\right|\right)$$

software default is usually $b_i = 0$.

Calculations "by hand" in R

= nrow(Telomeres)

```
Xbar = mean(Telomeres$years)
Ybar = mean(Telomeres$telomere.length)
s X = sd(Telomeres$vears)
s_Y = sd(Telomeres$telomere.length)
r_XY = cor(Telomeres$telomere.length, Telomeres$years)
SXX = (n-1)*s X^2
SYY = (n-1)*s Y^2
SXY = (n-1)*s X*s Y*r XY
beta1 = SXY/SXX
beta0 = Ybar - beta1 * Xbar
R2 = r XY^2
SSE = SYY*(1-R2)
sigma2 = SSE/(n-2)
sigma = sqrt(sigma2)
SE_beta0 = sigma*sqrt(1/n + Xbar^2/((n-1)*s_X^2))
SE beta1 = sigma*sqrt(
                                1/((n-1)*s_X^2))
```

Calculations "by hand" in R (continued)

```
# 95% CI for beta0
beta0 + c(-1, 1) * qt(.975, df = n-2) * SE_beta0
[1] 1.251761 1.483603
# 95% CI for beta1
beta1 + c(-1, 1) * qt(.975, df = n-2) * SE_beta1
[1] -0.044785794 -0.007962836
# pvalue for HO: beta0 <= 0 and P(beta0 > 0 | y)
pt(beta0 / SE_beta0, df = n - 2)
[1] 1
# pvalue for HO: beta1 <= 0 and P(beta1 > 0 | y)
pt(beta1 / SE beta1, df = n - 2)
[1] 0.003102353
# pvalue for H1: beta0 = 0
2 * pt(-abs(beta1 / SE_beta1), df = n - 2)
[1] 0.006204706
```

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Calculations by hand

```
\begin{array}{ll} SXX & = (n-1)s_{\frac{\pi}{2}}^2 = (39-1) \times 2.9354274^2 = 327.4358974 \\ SYY & = (n-1)s_{V}^2 = (39-1) \times 0.1797731^2 = 1.2280974 \end{array}
         SXY = (n-1)s_X s_Y r_{XY} = (39-1) \times 2.9354274 \times 0.1797731 \times -0.4306534 = -8.6358974
                     = SXY/SXX = -8.6358974/327.4358974 = -0.0263743
                     =\overline{Y}-\hat{\beta}_1\overline{X}=1.2202564-(-0.0263743)\times5.5897436=1.3676821
                      =r_{YY}^2 = (-0.4306534)^2 = 0.1854624
                      = SYY(1 - R^2) = 1.2280974(1 - 0.1854624) = 1.0003316
                       = SSE/(n-2) = 1.0003316/(39-2) = 0.027036
                      =\sqrt{\hat{\sigma}^2}=\sqrt{0.027036}=0.1644262
     SE(\hat{\beta}_0) = \hat{\sigma}\sqrt{\frac{1}{n} + \frac{\overline{X}^2}{(n-1)s_+^2}} = 0.1644262\sqrt{\frac{1}{39} + \frac{5.5897436^2}{(39-1)*2.9354274^2}} = 0.0572111
     SE(\hat{\beta}_1) = \hat{\sigma} \sqrt{\frac{1}{(n-1)s^2}} = 0.1644262 \sqrt{\frac{1}{(39-1)*2.9354274^2}} = 0.0090867
 \begin{array}{ll} p_{H_A:\beta_0\neq 0} & = 2P \left(T_{n-2} < - \left| \frac{\hat{\beta}_0}{SE(\hat{\beta}_0)} \right| \right) = 2P(t_{37} < -23.9058799) = 4.2740348 \times 10^{-24} \\ p_{H_A:\beta_1\neq 0} & = 2P \left(T_{n-2} < - \left| \frac{\hat{\beta}_1}{SE(\hat{\beta}_1)} \right| \right) = 2P(t_{37} < -2.9025065) = 0.0062047 \\ \end{array} 
  CI_{95\%\beta_0} = \hat{\beta}_0 \pm t_{n-2,1-a/2} SE(\hat{\beta}_0)
                      = 1.3676821 \pm 2.0261925 \times 0.0572111 = (1.2517613, 1.4836028)
                     = \hat{\beta}_1 \pm t_{n-2,1-a/2} SE(\hat{\beta}_1)
                       = -0.0263743 \pm 2.0261925 \times 0.0090867 = (-0.0447858, -0.0079628)
```

Regression in R

```
m = lm(telomere.length ~ years, Telomeres)
summary(m)
Call:
lm(formula = telomere.length ~ years, data = Telomeres)
Residuals:
    Min
              1Q Median
                                30
                                        Max
-0.42218 -0.08537 0.02056 0.10738 0.28869
Coefficients:
            Estimate Std. Error t value Pr(>|t|)
(Intercept) 1.367682 0.057211 23.906 <2e-16 ***
vears
           -0.026374 0.009087 -2.903 0.0062 **
---
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Residual standard error: 0.1644 on 37 degrees of freedom
Multiple R-squared: 0.1855.Adjusted R-squared: 0.1634
F-statistic: 8.425 on 1 and 37 DF. p-value: 0.006205
confint(m)
                 2.5 %
                             97.5 %
(Intercept) 1.25176134 1.483602799
           -0.04478579 -0.007962836
vears
```

Conclusion

Telomere ratio at the time of diagnosis of a child's chronic illness is estimated to be 1.37 with a 95% credible interval of (1.25, 1.48). For each year since diagnosis, the telomere ratio decreases on average by 0.026 with a 95% credible interval of (0.008, 0.045). The proportion of variability in telomere length described by a linear regression on years since diagnosis is 18.5%.

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The correlation between chronicity of caregiving and mean telomere length is -0.445 (P < 0.01). [$R^2 = 0.198$ was shown in the plot.]

Remark I'm guessing our analysis and that reported in the paper don't match exactly due to a discrepancy in the data.

Summary

• The simple linear regression model is

$$Y_i \stackrel{ind}{\sim} N(\beta_0 + \beta_1 X_i, \sigma^2)$$

where Y_i and X_i are the response and explanatory variable, respectively, for individual i.

- Know how to use R to obtain $\hat{\beta}_0$, $\hat{\beta}_1$, $\hat{\sigma}^2$, R^2 , p-values, Cls, etc.
- Interpret regression output:
 - β_0 is the expected value for the response when the explanatory variable is 0.
 - β_1 is the expected increase in the response for each unit increase in the explanatory variable.
 - ullet σ is the standard deviation of responses around their mean.
 - ullet R² is the proportion of the total variation of the response variable explained by the model.