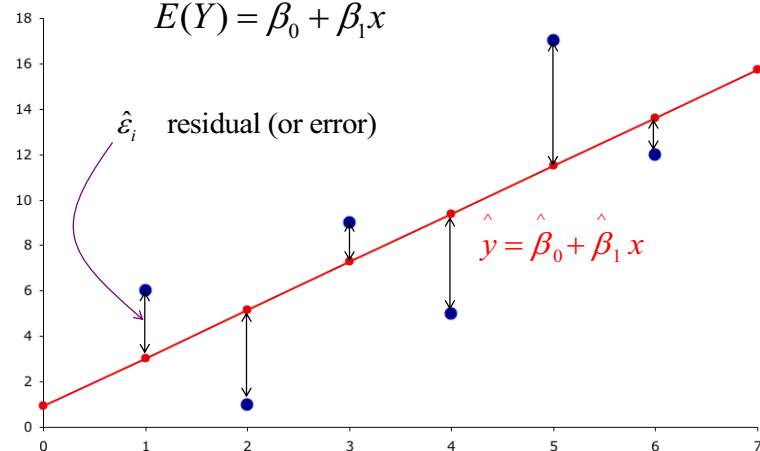


Linear Regression

Model: $Y = \beta_0 + \beta_1 x + \varepsilon \quad \varepsilon \sim N(0, \sigma)$

$$E(Y) = \beta_0 + \beta_1 x$$



Least Squares Estimation of β_0, β_1

- Least Squares Estimates for β_0 & β_1 minimize the sum of squared errors (SSE)

$$\hat{y} = \hat{\beta}_0 + \hat{\beta}_1 x$$

$$SSE = \sum_{i=1}^n \left(y_i - \hat{y}_i \right)^2 = \sum_{i=1}^n \left(y_i - \left(\hat{\beta}_0 + \hat{\beta}_1 x_i \right) \right)^2$$

Least Squares Estimates (LSEs)

$$\hat{\beta}_1 = \frac{\sum (x_i - \bar{x})(y_i - \bar{y})}{\sum (x_i - \bar{x})^2} = \frac{S_{xy}}{S_{xx}}$$

$$\hat{\beta}_0 = \bar{y} - \hat{\beta}_1 \bar{x}$$

$$\hat{\sigma}^2 = s^2 = \frac{\sum (y_i - \hat{y}_i)^2}{n-2} = \frac{SSE}{n-2} = MSE$$

Inference for the Slope (β_1)

- Standard Error of $\hat{\beta}_1$: $SE_{\hat{\beta}_1} = s / \sqrt{S_{xx}}$
- 2-Sided Test
 - $H_0: \beta_1 = 0$
 - $H_A: \beta_1 \neq 0$
- Test Statistic: $t_s = \frac{\hat{\beta}_1}{SE_{\hat{\beta}_1}}$
- Rejection Rule (RR): $|t_s| \geq t_{\alpha/2, n-2}$
- P-value: $2P(t \geq |t_{obs}|)$
- (1- α)100% Confidence Interval for β_1

$$\hat{\beta}_1 \pm t_{\alpha/2} SE_{\hat{\beta}_1} \equiv \hat{\beta}_1 \pm t_{\alpha/2} \frac{s}{\sqrt{S_{xx}}}$$

- Narrower CIs can be achieved through study design

Analysis of Variance in Regression

$$(y_i - \bar{y}) = (y_i - \hat{y}_i) + (\hat{y}_i - \bar{y})$$

$$\sum (y_i - \bar{y})^2 = \sum (y_i - \hat{y}_i)^2 + \sum (\hat{y}_i - \bar{y})^2$$

Total (SST)

$$\bullet df_{\text{Total}} = n-1$$

Error (SSE)

$$df_{\text{Error}} = n-2$$

Regression (SSR)

$$df_{\text{Regression}} = 1$$

Analysis of Variance in Regression

Source of Variation	Sum of Squares	Degrees of Freedom	Mean Square	F
Model	SSR	1	MSR = SSR/1	F = MSR/MSE
Error	SSE	n-2	MSE = SSE/(n-2)	
Total	SST = S _{yy}	n-1		

- Analysis of Variance - F-test

$$\bullet H_0: \beta_1 = 0 \quad H_A: \beta_1 \neq 0$$

$$F_s = \frac{MSR}{MSE}$$

$$RR: F_s \geq F_{\alpha, 1, n-2}$$

$$Pvalue: P(F \geq F_s)$$

Multiple Regression

- Numeric Response variable (Y)
- p Numeric predictor variables
- Model:

$$Y = \beta_0 + \beta_1 x_1 + \dots + \beta_p x_p + \varepsilon$$

- Partial Regression Coefficients: $\beta_i \equiv$ effect on the mean response of increasing the i^{th} predictor variable by 1 unit, **holding all other predictors constant**

Analysis of Variance

- Only changes from SLR are the d.f.

$$- df_{\text{Model}} = p \quad df_{\text{Error}} = n-p-1$$

Source of Variation	Sum of Squares	Degrees of Freedom	Mean Square	F
Model	SSR	p	MSR = SSR/p	F = MSR/MSE
Error	SSE	n-p-1	MSE = SSE/(n-p-1)	
Total	SST = S _{yy}	n-1		

$$R^2 = \frac{SST - SSE}{SST} = \frac{SSR}{SST}$$

Testing the Overall Model

- Tests whether **any** of the explanatory variables are associated with the response
- $H_0: \beta_1 = \dots = \beta_p = 0$
- H_A : Not all $\beta_i = 0$

$$F_s = \frac{MSR}{MSE}$$

$$RR: F_s \geq F_{\alpha, p, n-p-1}$$

$$Pvalue: P(F \geq F_s)$$

Models with Dummy Variables

- Since genotype is a categorical variable, we need to recode its values in order to include it in the regression model.
- If a categorical variable has k levels, need to create $k-1$ dummy variables that take on the values 1 if the level of interest is present, 0 otherwise.
- The baseline level of the categorical variable for which all $k-1$ dummy variables are set to 0
- The regression coefficient corresponding to a dummy variable is the difference between the mean for that level and the mean for baseline group, controlling for all numeric predictors

Analysis of Covariance

- Combination of 1-Way ANOVA and Linear Regression
- Goal: Comparing numeric responses among k groups, adjusting for numeric concomitant variable(s), referred to as **Covariate(s)**
- Clinical trial applications: Response is Post-Trt score, covariate is Pre-Trt score
- Epidemiological applications: Outcomes compared across exposure conditions (i.e., genotype), adjusted for other risk factors (age, BMI, sex,...)