

# **BIOSTATISTICS METHODS**

## **FOR TRANSLATIONAL & CLINICAL RESEARCH**



**DAY #4: REGRESSION APPLICATIONS, PART B**  
**COMPARING & COMBINING STATISTICS**

# CORRELATION & REGRESSION

- We have 2 continuous measurements made on each subject, one is the response variable Y, the other predictor X. There are two types of analyses:
- Correlation: is concerned with the association between them, measuring the strength of the relationship; the aim is to determine if they are correlated – the roles are exchangeable.
- Regression: To predict response from predictor.

You normally like to proceed to performing prediction **only if** the association is strong enough. However, in practice, “correlation analysis” only covers association whereas “regression analysis” would cover **both** association and prediction simultaneously.

In practice, we use statistical model and software for Regression, e.g. PROC REG of SAS, but putting more emphasis on “Correlation Results” in efforts called “**Risk Determination**” or “**Risk Assessment**”.

**The “regression component” is based the following model, called the “Normal Error Regression Model”:**

$$Y = \beta_0 + \beta_1 x + \varepsilon$$

$$\varepsilon \in N(0, \sigma^2)$$

The Mean Response is :

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

# PREDICTORS

- Unlike the response variable, we treat values of **Predictor or Covariate** as “fixed” - not random; and no restriction on its measurement scale.
- Examples of binary covariates include **treatment (yes/no), gender, and presence or absence of certain co-morbidity**; **Polytomous or categorical covariates** include **race, and different grades of symptoms**; Continuous covariates include **patient age, blood pressure, etc...**

**The “Correlation component” of Regression & Correlation is based on a joint distribution of X and Y. It is often referred to as a “correlation model”; the most widely used is the “Bivariate Normal Distribution” with density:**

$$f(X, Y) = \frac{1}{2\pi\sigma_x\sigma_y\sqrt{1-\rho^2}} \exp \left\{ -\frac{1}{2(1-\rho^2)} \left[ \left( \frac{X-\mu_x}{\sigma_x} \right)^2 - 2\rho \left( \frac{X-\mu_x}{\sigma_x} \right) \left( \frac{Y-\mu_y}{\sigma_y} \right) + \left( \frac{Y-\mu_y}{\sigma_y} \right)^2 \right] \right\}$$

$$\rho = \frac{\sigma_{xy}}{\sigma_x\sigma_y}$$

$$\sigma_{xy} = \text{Cov}(X, Y)$$

$$= E[(X - \mu_x)(Y - \mu_y)]$$

**“Regression Data” are often “designed”; values of the predictor are fixed in advance. “Correlation Data” are often cross-sectional or observational. For example, if we study the relationship between “height” and “weight” of a person, a sample of  $n$  people are taken but both measurements are random. Rather than a regression model, one should consider a “correlation model”**



$$f(X, Y) = \frac{1}{2\pi\sigma_x\sigma_y\sqrt{1-\rho^2}} \exp\left\{-\frac{1}{2(1-\rho^2)}\left[\left(\frac{X-\mu_x}{\sigma_x}\right)^2 - 2\rho\left(\frac{X-\mu_x}{\sigma_x}\right)\left(\frac{Y-\mu_y}{\sigma_y}\right) + \left(\frac{Y-\mu_y}{\sigma_y}\right)^2\right]\right\}$$

$$\rho = \frac{\sigma_{xy}}{\sigma_x\sigma_y}$$

$$\begin{aligned}\sigma_{xy} &= \text{Cov}(X, Y) \\ &= E[(X - \mu_x)(Y - \mu_y)]\end{aligned}$$

**$\sigma_{xy}$  is the Covariance and  $\rho$  is the Coefficient of Correlation between the two random variables X and Y;  $\rho$  is estimated by the (sample) Coefficient of Correlation r.**

The “regression component” - stipulated by the “Normal Error Regression Model” – and the “correlation component” – stipulated by the “Normal Correlation Model – can be tied together; regression method applied **conditionally** on fixed values of  $X$ .

# CONDITIONAL DISTRIBUTION

$$f(X,Y) = \frac{1}{2\pi\sigma_x\sigma_y\sqrt{1-\rho^2}} \exp\left\{-\frac{1}{2(1-\rho^2)}\left[\left(\frac{X-\mu_x}{\sigma_x}\right)^2 - 2\rho\left(\frac{X-\mu_x}{\sigma_x}\right)\left(\frac{Y-\mu_y}{\sigma_y}\right) + \left(\frac{Y-\mu_y}{\sigma_y}\right)^2\right]\right\}$$

## Theorem :

The conditional distribution of Y for any given  $X=x$  is normal with mean  $\beta_0 + \beta_1 x$  and standard deviation  $\sigma_{y|x}$  :

$$\beta_0 = \mu_y - \mu_x \rho \frac{\sigma_y}{\sigma_x}$$

$$\beta_1 = \rho \frac{\sigma_y}{\sigma_x}$$

$$\sigma_{y|x}^2 = (1-\rho^2)\sigma_y^2$$

$$\beta_1 = \rho \frac{\sigma_y}{\sigma_x}$$

$$\sigma_{y|x}^2 = (1 - \rho^2) \sigma_y^2$$

- (1) The slope and the Coefficient of Correlation measure the same thing but on different “scales”;**
- (2) The information on X helps to “reduce” the variance of Y; the square of the Coefficient of Correlation is the “Coefficient of Determination”.**

**We learned from basic statistics or biostatistics course:**

- (1) We can estimate the Slope; in repeated sampling, this statistic is distributed as normal. We can also estimate its variance (& standard error);**
- (2) We can estimate the Coefficient of Correlation; we do not usually mention the variance of this statistics because its sampling distribution is not normal. However, the Fisher's Transformation can be used to turn the coefficient of correlation into a normally-distributed statistic; and we can estimate variance (& standard error) of this newly transformed statistic.**

# APPLICATIONS

We can use what we learned to tackle these 2 problems:

- (1) We can compare two or several slopes and can combine to form a common value if these two or several statistics are not different.
  - (2) We can compare two or several coefficients of correlation and can combine to form a common value if these two or several statistics are not different.
- (In part C, we would re-visit this issue using Multiple Regression with alternative solution which, in many cases, might be a bit more elegant.)

Example #1:

# OCCUPATIONAL HEALTH

In applications, sometimes data are classified into groups, and within each group a separate regression model of  $Y$  on  $X$  may be postulated. For example, the regression of “forced expiratory volume” ( $Y$ ) on age ( $X$ ) may be considered separately for men in different occupational groups because different occupations may have different effects on the lung health of workers. Differences between the regression lines, especially the slopes, are our primary interest.

Suppose we study “vital capacity” among men working in the cadmium industry; the main purpose of the study was to see whether exposure to fumes was associated with a change in respiratory function. However, we must take into account the effect of “age” because respiratory performance declines with age. The men in the sample were divided into three groups:

- (1) those who were exposed for at least 10 years,
- (2) those who were exposed for less than 10 years,
- (3) Control group consisted of men not exposed to fumes.

We then consider three regression lines:

**Y = Vital Capacities (liters) versus X = Age**



It is well-known that respiratory test performance declines with age. But the question is **whether being exposed to fume in the cadmium industry would accelerate the declining process.** That is to focus on the **difference of slopes.**

We could consider to merge groups 1 and 2 then compare to group 3; however, the result would be **masked** by a phenomenon called “ **healthy worker effects**” (healthier people are more likely to choose more dangerous occupations).

The main focus could be placed on the comparison of group 1 versus group 2 – by showing an **attenuation of health worker effects**: the decline is steeper in group 1 (longer exposure) than in group 2 (shorter exposure).

When possible differences between the regression lines, for example the slopes, are of interest, there are two possibilities: **(1) If the slopes clearly differ**, from one group to another, then we have no choice but to draw separate group-specific inferences.

**(2) If the slopes do not differ**, the lines are parallel with a common slope; that common slope can and should be estimated **using combined data from all groups**.

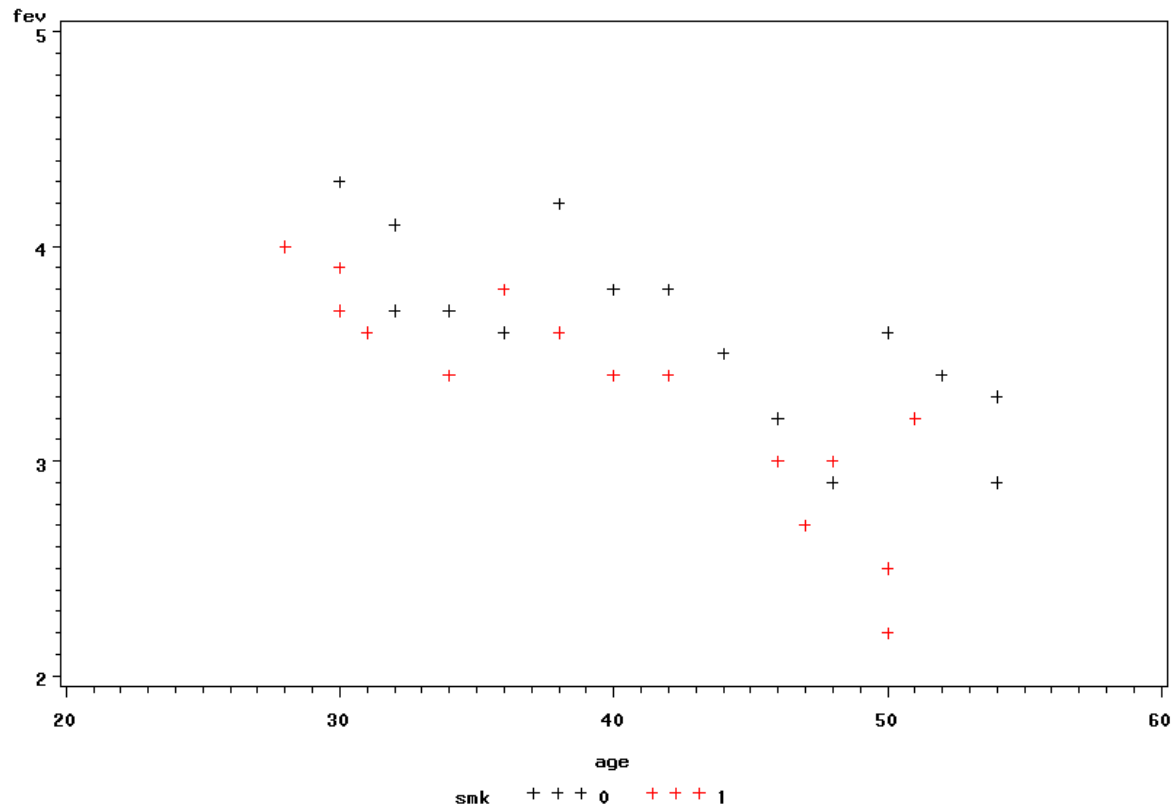
In practice, the fitted regression lines would rarely have precisely the same slope or position – as seen from the scatter diagram. The question is to what extent the differences can be attributed to random variation. There are simple ways to “compare” and, if applicable, to “combine” data forming the common slope.

## Example #2:

# SMOKING & LUNG HEALTH

Response Variable: Forced Expired Volume (FEV),  
a measure of lung health.

Main Predictor: Age



- (1) Generally, FEV is linearly related to Age,**
- (2) Non-smokers have better lung health as compared to smokers; the Regression Line for smokers looks “steeper”. To see if smoking could modify the natural effect of Age on Lung Health; we might want to compare the slopes. If they are not different, the common parameter would be more precisely estimated using all data.**

## Example #3:

# WHITE CELLS & LEUKEMIA

Leukemia is a cancer characterized by an over-proliferation of white blood cells; the higher the white blood count (WBC), the more severe the disease; **WBC is an important predictor of Survival Time (the Response)**. Another important factor is the presence (“AG positive”) or absence (“AG negative”) of Auer rods and/or significant granulation of the leukemic cells in the bone marrow at diagnosis.

AG-Positive, n = 17		AG-Negative, n = 16	
White Blood count (WBC)	Survival Time (weeks)	White Blood Count (WBC)	Survival Time (weeks)
2,300	65	4,400	56
750	156	3,000	65
4,300	100	4,000	17
2,600	134	1,500	7
6,000	16	9,000	16
10,500	108	5,300	22
10,000	121	10,000	3
17,000	4	19,000	4
5,400	39	27,000	2
7,000	143	28,000	3
9,400	56	31,000	8
32,000	26	26,000	4
35,000	22	21,000	3
100,000	1	79,000	30
100,000	1	100,000	4
52,000	5	100,000	43
100,000	65		

**Look at the effects of “AG factor”, that morphologic characteristic of white cells! We might want to compare the two coefficients of correlation (between WBC and survival time); there is a strong effect modification here.**



AG-Positive, n = 17		AG-Negative, n = 16	
White Blood count (WBC)	Survival Time (weeks)	White Blood Count (WBC)	Survival Time (weeks)
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10,000	121	10,000	3
17,000	4	19,000	4
5,400	39	27,000	2
7,000	143	28,000	3
9,400	56	31,000	8
32,000	26	26,000	4
35,000	22	21,000	3
100,000	1	79,000	30
100,000	1	100,000	4
52,000	5	100,000	43
100,000	65		

It can be easily seen that, among AG-positive patients, WBC and Survival Time are negatively correlated – as noted that “the higher the white blood count (WBC), the more severe the disease”. But that is not necessarily true for AG-negative patients: **AG modifies the effect of WBC.**

## Example #4:

# KIDNEY ACTIVITIES

One way to learn about the function of the kidney is to study the rates at which it produces and consumes different substances. One important aspect is the rate at which oxygen is consumed since this is considered to be a measure of how hard the kidney is working. Another aspect of interest is the rate at which kidney reabsorbs ionic sodium from the urine. This later activity, known as sodium pumping, requires energy. Thus there should be a direct relationship between the intensity of sodium pumping and the rate of oxygen consumption.

## **An interesting and important Regression Model:**

**In the regression of Oxygen Consumption Y (Response) against Sodium Re-absorption X (Predictor Explanatory Variable), the Intercept was interpreted as the base rate of oxygen consumption (which is less important) and the reciprocal of the Slope was used as the estimate of the “pumping efficiency”, an important measure of renal health. Investigators could use pumping efficiency as an outcome to compare different experiment conditions or to evaluate (new) medication/supplement.**

## Tasks:

Studies could be carried in human or animals (dogs or rats); in animals, it could be live or kidneys could be removed from animals and maintained with fluids. Experiments could be done or observations could be made to obtain several data points from each kidney. Then we could compare the slopes across subjects in the same experiment condition and combined to form a common pumping efficiency. How do we do that? How do we compare several slopes?

# THE DEMAND CURVE

A fundamental concept of consumer demand, in Behavioral Economics, is the **Demand Curve** relating the consumption of a commodity ( $Q$ , the dependent variable) to its (unit) price ( $P$ , the independent variable). According to the theory, the consumption of most goods will decrease with increases in price (Watson and Holman, 1977). At the backbone of the Demand Curve is the concept of **Elasticity**.

# ELASTICITY

At the discrete level, a section of the demand curve is characterized by a parameter called **Elasticity (E)** which is defined as the ratio of two rates or proportions:

$$E = \frac{\frac{(Q_2 - Q_1)}{(1/2)(Q_1 + Q_2)}}{\frac{(P_2 - P_1)}{(1/2)(P_1 + P_2)}}$$

**“Elasticity”** could be used to compare **liability** between products; that with the same price increase, consumption of one product would reduce faster than that of the other. For example, the difference could represent different levels of dependency or **addiction**.

# ELASTICITY on Continuous Scale

For a point on the demand curve, i.e. continuous scale, the elasticity  $E$  becomes:

$$E = \frac{(Q_2 - Q_1)}{\frac{(1/2)(Q_1 + Q_2)}{(P_2 - P_1)}} \longrightarrow E = \left[ \frac{P}{Q} \right] \left[ \frac{dQ}{dP} \right]$$
$$= \frac{d[\ln Q]}{d[\ln P]}$$

which represents the **slope** on the demand curve when both price ( $P$ ) & consumption ( $Q$ ) are expressed on the **log scale** (we do not have to graph with both on log scale).



# **DEMAND CURVE FOR TOBACCO RESEARCH**

**The demand curve established for food consumption has been adopted for use in tobacco research in areas of product liability and relative reinforcing efficacy (RRE), a concept in psychopharmacological research (Bickel and Madden 1999).**

**There are studies both in humans (surveys of smokers) & animals (experiments with rats)**

# ANIMAL EXPERIMENTS

- Human research suggests that there are sex differences in the addiction-related behavioral effects of nicotine; a study was conducted to examine this issue in rats.
- Male and female rats were trained to self-administer nicotine (0.06 mg/kg) under a FR 3 schedule during daily 23-hour sessions.
- Rats were then exposed to saline extinction and reacquisition of NSA, followed by weekly reductions in the unit dose (0.03 to 0.00025 mg/kg) until extinction levels of responding were achieved.
- Fifteen rats (8 males, 7 females) were tested at 8 doses: 0.03, 0.02, 0.01, 0.007, 0.004, 0.002, 0.001, 0.0005, mg/kg/infusion.

# SURVEYS OF SMOKERS

- **Data are collected by the cigarette purchase task (CPT) survey, also called TPT, in which participants were asked to respond to the following set of questions**
- **How many cigarettes would you smoke if they were \_\_\_\_\_ each?: 0¢ (free), 1¢, 5¢, 13¢, 25¢, 50¢, \$1, \$2, \$3, \$4, \$5, \$6, \$11, \$35, \$70, \$140, \$280, \$560, \$1,120.**
- **This set of questions are asked during an online survey in the preceding order until the respondents gives “0” as an answer, then no more further questions will be asked.**

# STANDARDIZATION

A Standardized Demand Curve could be formed as follows; let:

$$t = \ln(P/P_B)$$

$$S(t) = \frac{Q}{Q_B}$$

“Survival Fraction”  $S(t) = Q/Q_B$  going down from 1.0 as “time”  $t$  increases. In this setup, individual curves have the same shape as the “global curve”.

# HOW TO EXPRESS ELASTICITY?

$$S(t) = \frac{Q}{Q_0} \text{ \& } t = \ln(Q/Q_0) = \ln Q - \ln Q_0$$

$$\ln[S(t)] = \ln(Q) - \ln(Q_0)$$

$$\begin{aligned} h(t) &= -\frac{d}{dt} \ln[S(t)] = -\frac{d(\ln Q)}{d(\ln P)} \\ &= -\text{Elasticity} \end{aligned}$$

If we view the Standardized Demand Curve as a survival curve, **Elasticity Function** is simply the negative of the Hazard Function.

# A Possibility: WEIBULL MODEL

St Demand Curve  $S(t) = \exp[-(\alpha t)^\beta]$

Elasticity  $E(t) = -\alpha\beta(\alpha t)^{\beta-1}$

**Could it be more simple? Yes, if  $\beta=1$ , the standardized demand curve has linear elasticity.**

# DATA ANALYSIS STRATEGIES

- There could be two choices:
  - (1) Starting with individual curves, then combining results to form the population or global curve.
  - (2) Going right to population curve, and treat individual data as repeated observations.
- The first strategy is more simple – a straight application of Simple Linear Regression - and it would show individual differences.

# DATA TRANSFORMATION

$$t = \ln(P/P_0)$$

$$S(t) = \frac{Q}{Q_B}$$

$$S(t) = \exp[-(\alpha t)^\beta]$$

$$E(t) = -\alpha\beta(\alpha t)^{\beta-1}$$



$$\ln[-\ln S(t)] = \beta \ln \alpha + \beta \ln t$$

$$\ln\left[-\ln \frac{Q}{Q_B}\right] = \beta \ln \alpha + \beta \ln[\ln(P/P_B)]$$

**We have a simple linear regression after two double log transformations; goodness-of-fit could be judged visually ( $R^2$  is a good measure). We combine individual results by calculating weighted averages of slopes and intercepts using inverse of variance as the weight; and use these weighted averages to form Standardized Demand & Elasticity functions.**



# **METHOD**

## **FOR COMPARING & COMBINING STATISTICS**

# Review A: COMPARISON OF TWO POPULATION MEANS

- In this type of problems, we have two independent samples  $(n_1, \bar{y}_1, s_1^2)$  and  $(n_2, \bar{y}_2, s_2^2)$ ; the  $n$ 's being the sample sizes,  $\bar{y}$  the sample means, and  $s^2$  the sample variances (the  $s$  are standard deviations).
- Often called the “two-sample problem”
- Considered as samples with population means  $\mu_1$  and  $\mu_2$
- The aim is to compare the two population means.
- (“Y” is the “response”, a measure of interest)

$$t = \frac{\bar{y}_2 - \bar{y}_1}{\sqrt{\left(\frac{(n_1 - 1)s_1^2 + (n_2 - 1)s_2^2}{n_1 + n_2 - 2}\right)\left(\frac{1}{n_1} + \frac{1}{n_2}\right)}}$$

$$s_i^2 = \frac{\sum_{j=1}^{n_i} (y_{ij} - \bar{y}_i)^2}{n_i - 1}; \quad s_p^2 = \frac{(n_1 - 1)s_1^2 + (n_2 - 1)s_2^2}{n_1 + n_2 - 2}$$

$$t = \frac{\bar{y}_2 - \bar{y}_1}{s_p \sqrt{\frac{1}{n_1} + \frac{1}{n_2}}}$$

**The “key” term is the numerator, the difference between sample means; we find the difference by subtracting one mean from the other mean.**

# Review B: COMPARISON OF POPULATION SEVERAL MEANS

- **Suppose we want to know whether there are differences in the means of more than two independent groups. For example, do families of different ethnic groups have different income levels?**
- **Important Question: How to measure the “difference” (between sample means)?**

# COMPONENTS OF TOTAL VARIATION

- The total variation in the combined sample can be decomposed into two components as follows:

$$(x_{ij} - \bar{x}) = (x_{ij} - \bar{x}_i) + (\bar{x}_i - \bar{x}):$$

- (1) The first term reflects the **variation within the samples**; the following sum is called the “**within sum of squares**”:
- $$SSW = \sum (x_{ij} - \bar{x}_i)^2 = \sum (n_i - 1)s_i^2$$

- (2) The difference, **SSB = SST - SSW**, is called the “**between sum of squares**” which measures the differences between samples:

$$SST = \sum_{i,j} (x_{ij} - \bar{x})^2$$

$$SSB = \sum_{i,j} (\bar{x}_i - \bar{x})^2 = \sum_i n_i (\bar{x}_i - \bar{x})^2$$

# ANOVA: ANALYSIS OF VARIANCE

- **SST** measures the “total variation” in the combined sample with  $(n-1)$  degrees of freedom,  $n=\sum n_i$  is the total size. It is decomposed into:  
 **$SST=SSW+SSB$**
- **SSW** measures the variation within samples with  $\sum(n_i-1)=(n-k)$  degrees of freedom, and
- **SSB** measures the variation between sample means with  $(k-1)$  degrees of freedom;  $k=\#$  of groups

**SSB** measures the **variation, or difference,**  
**between sample means:**

$$SSB = \sum_{i,j} (\bar{x}_i - \bar{x})^2 = \sum_i n_i (\bar{x}_i - \bar{x})^2$$

**(which is a concept similar to the “variance”:  
variation among sample means); decision is  
based on the F-statistic:  $F = MSB/MSW$ .**

**Grand mean is a “weighted Average”; the weight is the inverse of the variance:**

$$\begin{aligned}\bar{\mathbf{x}} &= \frac{\sum_{i,j} \mathbf{x}_{ij}}{N} \\ &= \frac{\sum_i \mathbf{n}_i \bar{\mathbf{x}}_i}{\sum_i \mathbf{n}_i} \\ &= \frac{\sum_i \frac{\mathbf{n}_i}{\sigma^2} \bar{\mathbf{x}}_i}{\sum_i \frac{\mathbf{n}_i}{\sigma^2}}\end{aligned}$$



**Between Sum of Squares is a weighted sum of deviations (from weighted mean); the weight is the inverse of the variance: :**

$$\begin{aligned}SSB &= \sum n_i (\bar{x}_i - \bar{x})^2 \\ &= \sum \frac{\sigma^2}{\text{Var}(\bar{x}_i)} (\bar{x}_i - \bar{x})^2 \\ &= \text{Weighted Sum of Deviations}\end{aligned}$$

# GROUP-SPECIFIC RESULTS

Suppose there are  $k$  groups with  $n_i$  pairs of observations (i.e. “correlation data”) in **the  $i$ th group; a regression line is fitted with the following results for the slope:**

Estimated slope =  $b_{1i}$

$$s^2(b_{1i}) = \frac{MSE_i}{\sum_i (x - \bar{x})^2}$$

# GENERAL METHODOLOGY

Let  $w_i$  be the inverse of the variance of the  $i$ th slope; under the null hypothesis that the true slopes of the  $k$  groups are all equal, the following statistic  $G$  follows approximately a **Chi-square distribution with  $(k-1)$  degrees of freedom**:

$$w_i = \frac{1}{s^2(b_{1i})}$$

$$\tilde{b}_1 = \frac{\sum w_i b_{1i}}{\sum w_i}$$

$$G = \sum w_i (b_{1i} - \tilde{b}_1)^2$$

**Weighted average** →

**Similar to ANOVA** →

If the statistic  $G$  is not statistically significant, the null hypothesis that the true slopes of the  $k$  groups are all equal is “tentatively accepted”, the common slope is best estimated by the weighted average (of the  $k$  individual slopes). **The sampling distribution of this weighted average is approximately normal.**

$$w_i = \frac{1}{s^2(b_{1i})}$$

$$\tilde{b}_1 = \frac{\sum w_i b_{1i}}{\sum w_i}$$

$$\sigma^2(\tilde{b}_1) = \frac{1}{\sum w_i}$$

## Recall:

$$b_1 = \frac{\sum (x - \bar{x})y}{\sum (x - \bar{x})^2}$$

$$\begin{aligned} \text{Var}(b_1) &= \frac{\sum (x - \bar{x})^2 \text{Var}(y)}{\{\sum (x - \bar{x})^2\}^2} \\ &= \frac{\sigma^2}{\sum (x - \bar{x})^2} \end{aligned}$$

Therefore, we can use as the weight, the  
Sum of Squares of X:

$$SSX = \sum (x_i - \bar{x})^2$$

We **could** use the same method to compare and combine **intercepts** and **coefficients of correlation**; for the later one we take advantage of the Fisher's transformation:

$$z = \frac{1}{2} \ln \left\{ \frac{1+r}{1-r} \right\}$$

$$\text{Var}(z) = \frac{1}{n-3}$$

**The result becomes much more simple when we only need to compare two slopes (or any two statistics).**

$$\tilde{b} = \frac{w_1 b_1 + w_2 b_2}{w_1 + w_2}$$

$$\begin{aligned} b_1 - \tilde{b} &= b_1 - \frac{w_1 b_1 + w_2 b_2}{w_1 + w_2} \\ &= \frac{w_2 (b_1 - b_2)}{w_1 + w_2} \end{aligned}$$

$$b_2 - \tilde{b} = \frac{w_1 (b_1 - b_2)}{w_1 + w_2}$$



$$\mathbf{b}_1 - \tilde{\mathbf{b}} = \frac{\mathbf{w}_2(\mathbf{b}_1 - \mathbf{b}_2)}{\mathbf{w}_1 + \mathbf{w}_2}$$

$$\mathbf{b}_2 - \tilde{\mathbf{b}} = \frac{\mathbf{w}_1(\mathbf{b}_1 - \mathbf{b}_2)}{\mathbf{w}_1 + \mathbf{w}_2}$$

$$\begin{aligned} \mathbf{G} &= \mathbf{w}_1(\mathbf{b}_1 - \tilde{\mathbf{b}})^2 + \mathbf{w}_2(\mathbf{b}_2 - \tilde{\mathbf{b}})^2 \\ &= \left[ \frac{\mathbf{w}_1\mathbf{w}_2^2 + \mathbf{w}_2\mathbf{w}_1^2}{(\mathbf{w}_1 + \mathbf{w}_2)^2} \right]^2 (\mathbf{b}_1 - \mathbf{b}_2)^2 \\ &= \frac{\mathbf{w}_1\mathbf{w}_2}{(\mathbf{w}_1 + \mathbf{w}_2)} (\mathbf{b}_1 - \mathbf{b}_2)^2 \end{aligned}$$

$$\begin{aligned} \mathbf{G} &= \frac{\mathbf{w}_1 \mathbf{w}_2}{(\mathbf{w}_1 + \mathbf{w}_2)} (\mathbf{b}_1 - \mathbf{b}_2)^2 \\ &= \frac{(\mathbf{b}_1 - \mathbf{b}_2)^2}{\frac{1}{\mathbf{w}_1} + \frac{1}{\mathbf{w}_2}} \\ &= \frac{(\mathbf{b}_1 - \mathbf{b}_2)^2}{\text{Var}(\mathbf{b}_1) + \text{Var}(\mathbf{b}_2)} \end{aligned}$$

**This is equivalent to referring the following statistic to percentiles of the Standard Normal distribution – a very common practice:**

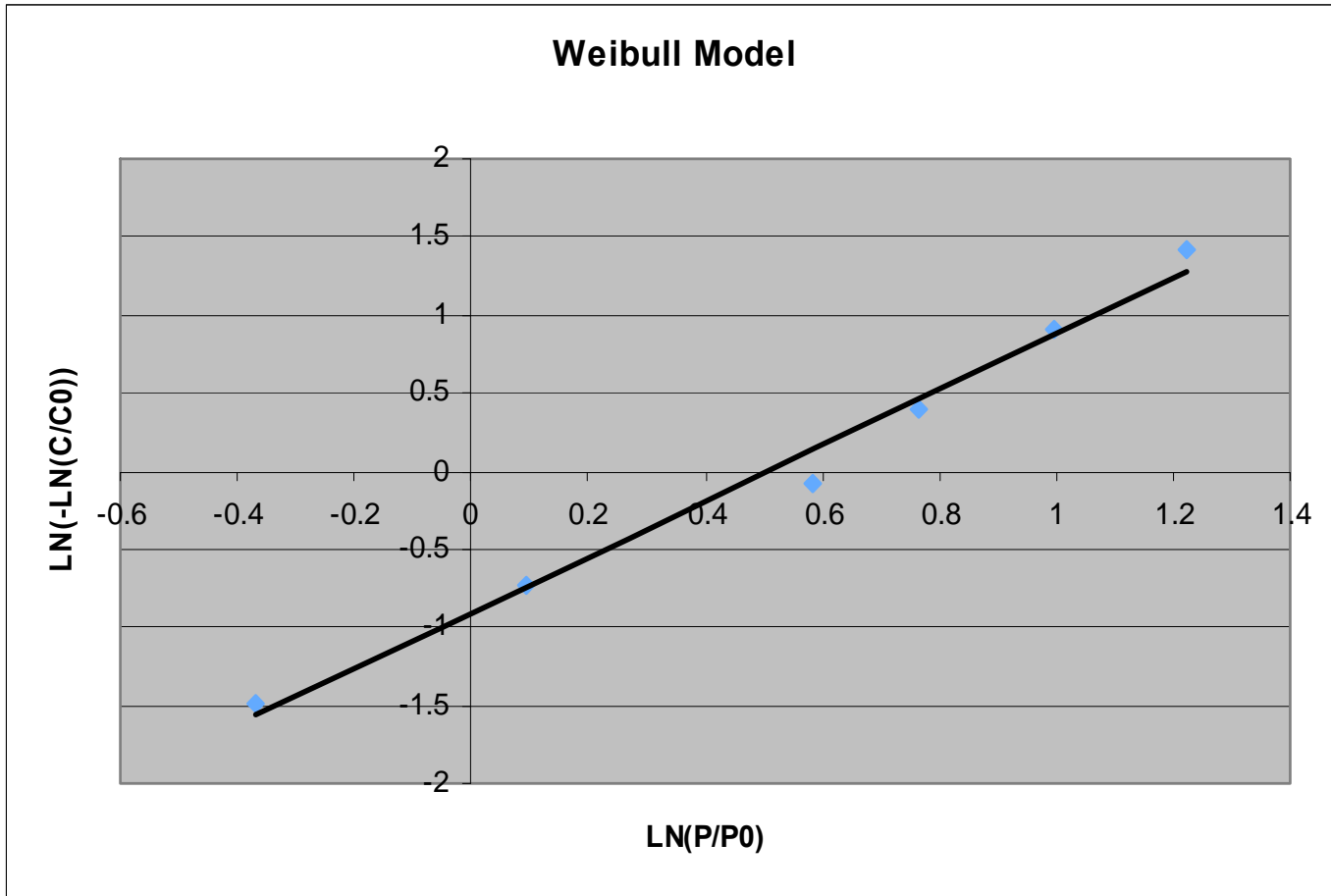
$$\mathbf{z} = \frac{\mathbf{b}_1 - \mathbf{b}_2}{\sqrt{\mathbf{Var}(\mathbf{b}_1) + \mathbf{Var}(\mathbf{b}_2)}}$$

**(Variance of difference is equal sum of variances)**

# NUMERICAL EXAMPLE: RATS DATA

Price	Males							
50	2.0220	1.4820	1.4400	2.2200	1.3800	1.6800	2.0820	1.6560
100	1.6110	1.0800	0.9810	1.4610	1.0110	1.3200	1.7490	0.7710
150	1.2460	0.8200	0.8140	1.0400	0.9000	1.0460	1.3940	0.7200
300	0.8000	0.6130	0.4600	0.6430	0.5870	0.6000	0.6170	0.4600
429	0.4571	0.1589	0.1449	0.4179	0.4809	0.4760	0.1281	0.2751
750	0.1692	0.0588	0.0520	0.2200	0.2200	0.2492	0.0172	0.0960
1500	0.0320	0.0106	0.0140	0.0346	0.0880	0.0834		0.0180
3000				0.0117	0.0290			
6000					0.0057			
12000					0.0022			

Price	Females						
50	3.0000	2.2200	1.6200	3.3600	1.9980	1.8420	2.3220
100	1.4400	0.9690	0.8310	1.8990	1.1700	1.2210	1.3500
150	1.0260	0.9000	0.7340	1.1260	1.0000	1.0940	0.9260
300	0.6830	0.2630	0.2870	0.6830	0.5500	0.7200	0.6700
429	0.1680	0.1470	0.0959	0.5670	0.4571	0.5159	0.4501
750	0.0652	0.0320	0.0428	0.3348	0.2188	0.2732	0.2320
1500			0.0094	0.0586	0.0606	0.1314	0.0400
3000					0.0193	0.0227	0.0157
6000					0.0108	0.0089	
12000							



**Weibull Model fits well**

# RESULTS

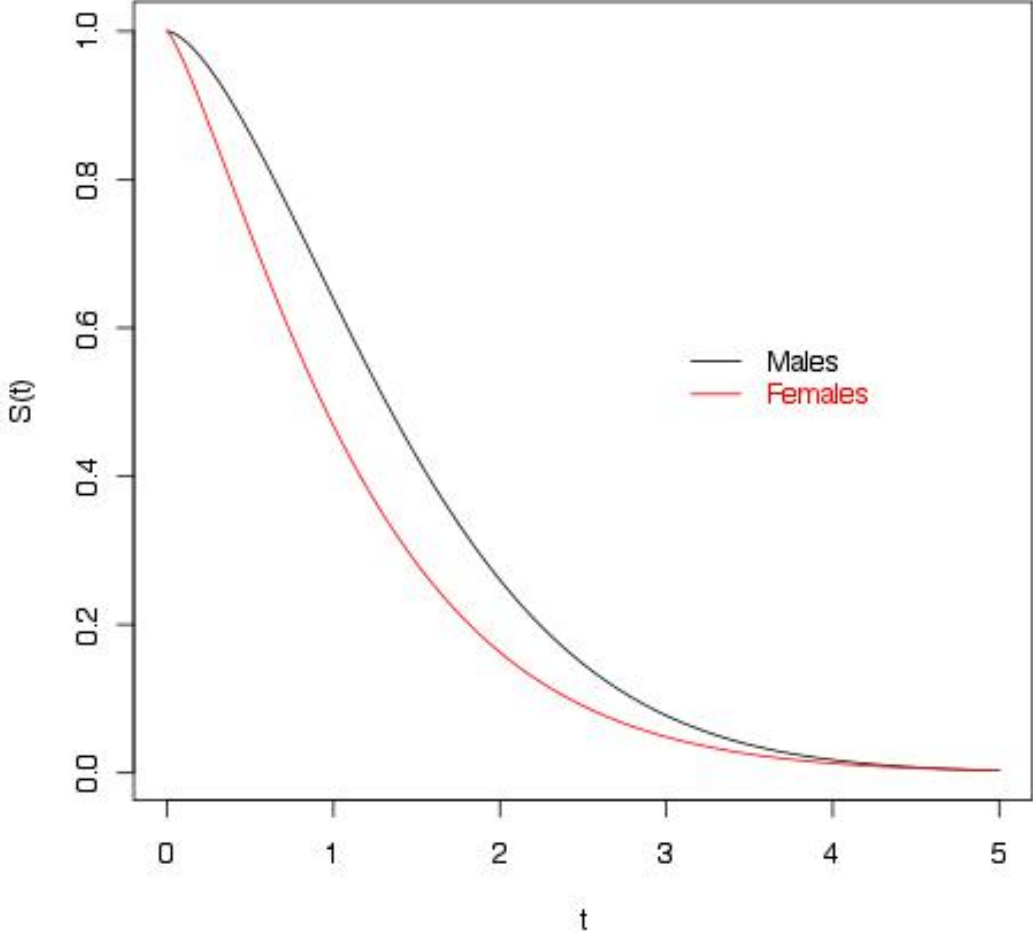
## Males

<b>Intercept</b>	-0.911	-0.667	-0.544	-0.452	-0.891	-0.883	-0.995	-0.123
<b>SE(Intercept)</b>	0.08	0.158	0.122	0.081	0.102	0.034	0.125	0.151
<b>Slope</b>	1.793	1.752	1.651	1.416	1.56	1.558	2.463	1.115
<b>SE(Slope)</b>	0.104	0.205	0.158	0.091	0.093	0.044	0.194	0.197
<b>R<sup>2</sup></b>	0.987	0.948	0.965	0.98	0.976	0.997	0.982	0.889
<b>Global Parameter:</b>								
	<b>Alpha = 0.603</b>							
	<b>Beta = 1.585+/-0.032</b>							
	<b>R<sup>2</sup> = 0.971</b>							

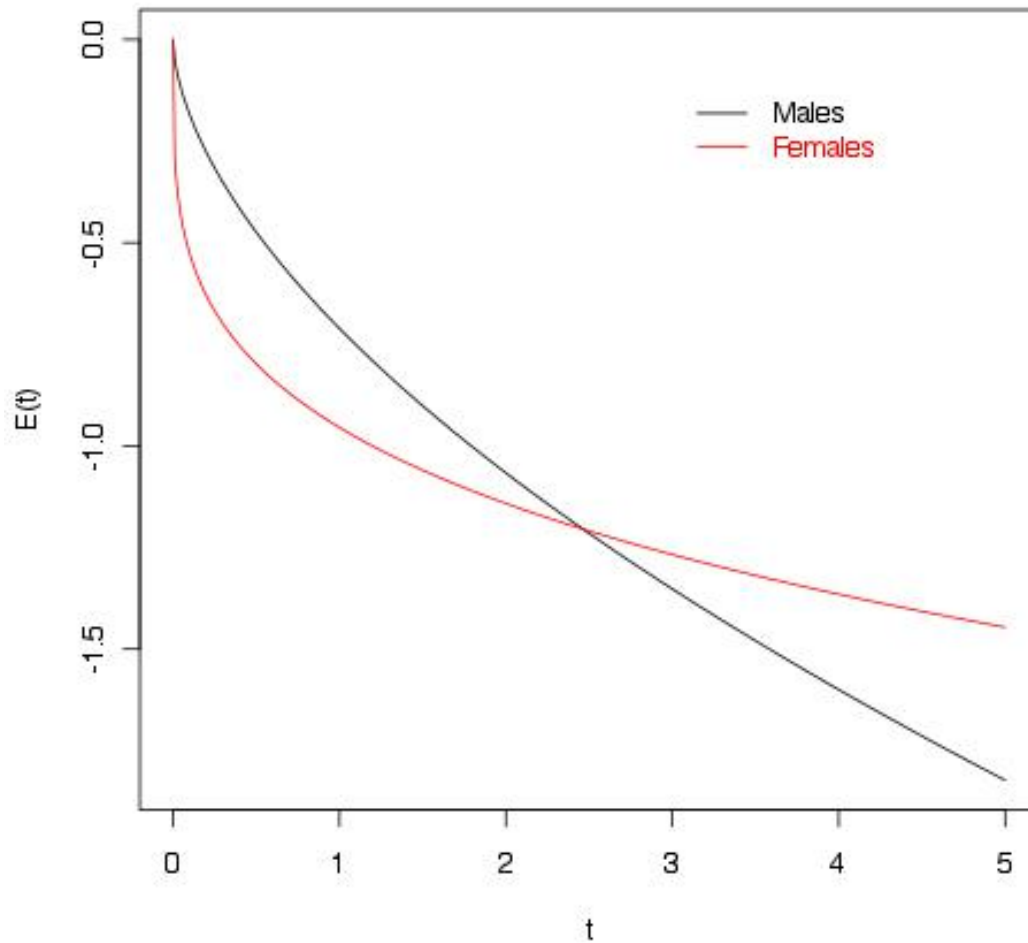
## Females

<b>Intercept</b>								
<b>SE(Intercept)</b>								
<b>Slope</b>								
<b>SE(Slope)</b>								
<b>R<sup>2</sup></b>								
<b>Global Parameters:</b>								
	<b>Alpha = 0.803</b>							
	<b>Beta = 1.258+/-0.047</b>							
	<b>R<sup>2</sup> = 0.956</b>							

# STANDARDIZED DEMAND CURVES



# ELASTICITY CURVES





- (1) For both males and females, we have “decreasing elasticity”; consumption reduction accelerates as prices increases;**
- (2) What’s interesting is the two curves are crossing at a very high price; for lower prices the consumption for females drops faster first but it becomes slower at higher prices.**
- (3) One possible explanation is that female rats are weaker (larger  $\alpha$ , early reduction) but more addicted (smaller  $\beta$ , more resistant to reduction, difference narrows down).**

# Suggested Exercises

- #3** Refer to the Demand Curve data and focus on the group of female rats:
- (a) Compare the slopes and, if the difference is not significant, calculate the weighted average and its standard error.
  - (b) Compare the slopes and, if the difference is not significant, calculate the weighted average and its standard error.
  - (c) Show how to obtain the global parameters  $\alpha$  and  $\beta$ .
- #4** For the Kidney activities, we have a “control” data set for which investigator lowered pressure in order to perturb the kidney equilibrium point so as to provide a range of values for regression; there were 5 observations for each of 10 kidneys, data are given on next page). Compare the slopes and, if the difference is not significant, calculate the weighted average and its standard error.

<b>Kidney i</b>	<b>Slope, <math>b_i</math></b>	<b><math>SSX_i</math></b>
<b>1</b>	<b>0.00967</b>	<b>1384</b>
<b>2</b>	<b>0.04784</b>	<b>360</b>
<b>3</b>	<b>0.03134</b>	<b>753</b>
<b>4</b>	<b>0.01928</b>	<b>3153</b>
<b>5</b>	<b>0.01928</b>	<b>3050</b>
<b>6</b>	<b>0.01747</b>	<b>4575</b>
<b>7</b>	<b>0.04817</b>	<b>1570</b>
<b>8</b>	<b>0.01893</b>	<b>4175</b>
<b>9</b>	<b>0.04233</b>	<b>719</b>
<b>10</b>	<b>0.02706</b>	<b>885</b>

Only # 7.1(a) and #7.2 are required