

## **Comparing Interval Estimates**

- □ Statistical interval estimates are constructed to
- Estimate parameters
- Quantify characteristics of population

□ To correctly interpret estimates, it must be clearly defined what each interval is estimating

- Confidence/prediction intervals are well understood
- Definition of a tolerance interval varies among literature sources

# What is a Tolerance Interval?

□ Tolerance intervals are being used with more frequency, thus a consistent definition needs to be established

- **D**efinitions found in literature:
- A bound that covers at least  $(100-\alpha)\%$  of the measurements with  $(100-\gamma)\%$ confidence (*Walpole and Myers*)
- o Focuses on where individual observations fall
- o Equivalent to a  $(100-\gamma)$ % CI on middle  $(100-\alpha)$ % of Normal distribution
- An interval that includes a certain percentage of measurements with a known probability (Mendenhall and Sincich)

o TI is identical to CI, except it attempts to capture a proportion of measurements rather than a population parameter, such as  $\mu$ 

• Computational methods vary depending on author

- Mee's definition uses non-central t distribution
- Owen et al. propose to control the percentage in both tails of a distribution • No more than a specified proportion lies below or above the TI

□ Alternative definition gives interval estimates on lower and upper *percentiles*, not a *percentage*, of a distribution

lower TI = 
$$[\overline{X} - (\alpha/2, n-1, \delta)) \frac{s}{\sqrt{n}}, \overline{X} - t(1-\alpha/2, n-1, \delta) \frac{s}{\sqrt{n}}]$$
 and  
TI =  $[\overline{X} - (\alpha/2, n-1, \delta)) \frac{s}{\sqrt{n}}, \overline{X} - t(1-\alpha/2, n-1, \delta) \frac{s}{\sqrt{n}}]$ 

upper TI = 
$$[\overline{X} + t(\alpha/2, n-1, \delta)) \frac{s}{\sqrt{n}}, \ \overline{X} + t(1-\alpha/2, n-1, \delta) \frac{s}{\sqrt{n}}]$$
, where  $\delta = \Phi(p)$ 

### **Other Interval Estimates**

- □ Simultaneous tolerance interval:
- Tolerance interval computed for more than one population or sample at a time
- **Two 1-sided tolerance interval**:
- 1-sided TI on lower  $\alpha/2\%$  together with 1-sided TI on upper  $\alpha/2\%$ , designed to capture  $(1-\alpha)$ % of distribution
- Confidence/prediction interval on confidence/prediction interval endpoints: Interval that protects the mean/future confidence/prediction interval endpoints (based on asymptotic Normal distribution of interval endpoints)
- $\square$   $\beta$ -expectation tolerance interval (Mee's definition):
- Interval that contains approximately 100β% of the distribution:  $E_{\hat{\mu},\hat{\sigma}_{x}} \{ \Pr_{X}[\hat{\mu} - k\hat{\sigma}_{x} < X < \hat{\mu} + k\hat{\sigma}_{x} | \hat{\mu}, \hat{\sigma}_{x}] \} = \beta$

$$\sigma_x^2 = \sigma_b^2 + \sigma_e^2$$
  $\hat{\sigma}_x^2 = MSB/J + (1-1/J)MSE$ 

I = # of batches, runs, blocks, etc. J = # of reps within runs

# **Evaluating Tolerance Interval Estimates**

Michelle Quinlan, University of Nebraska-Lincoln James Schwenke, Boehringer Ingelheim Pharmaceuticals, Inc. Walt Stroup, University of Nebraska-Lincoln



- Computed using  $\sigma_b^2/\sigma_e^2$  (ratio of between to within batch variance) and central *t*-distribution
- using Satterthwaite approximation
- $\square$   $\beta$ -content tolerance interval:

 $\Pr_{\hat{\mu},\hat{\sigma}_{x}} \{ \Pr_{X} [\hat{\mu} - k\hat{\sigma}_{x} < X < \hat{\mu} + k\hat{\sigma}_{x} | \hat{\mu},\hat{\sigma}_{x} ] \ge \beta \} = \gamma$  $\gamma$  = confidence coefficient

- level  $\gamma$  (*Mee*)
- using noncentral *t*-distribution
- *Wald and Wolfowitz* use same definition to define tolerance intervals but instead use the formula:

 $\overline{X} \pm \sqrt{\frac{\pi}{\gamma^2}} rs$ 

□ SAS<sup>®</sup> Proc Capabilities Method 3 computes an approximate statistical tolerance interval that contains at least *p* proportion of the population with formula given by:

$$\overline{\mathbf{X}} \pm \mathbf{z}_{\frac{1+\mathbf{p}}{2}} (1+1/2n) \sqrt{\frac{n}{\chi_{\alpha}^2}}$$

- limits on percentiles
- Formulas are provided for constructing TI when one or both  $\mu$  and  $\sigma$  are unknown

### **Evaluation of Interval Estimates**

- $\Box$  The relationship among prediction,  $\beta$ -expectation, and  $\beta$ -content intervals is investigated
- with 1 variance component (all factors fixed)
- $\beta$ -expectation intervals are compared with  $\beta$ -content intervals

# **Prediction vs. β-Expectation Tolerance Intervals**

- □ 1 variance component
- treatment means (regardless of the number of factors in model)
- Here  $\sigma_b^2 / \sigma_e^2 = 0$  because = 0
- No df adjustment needed

# **Comparing Interval Estimates**

• To understand tolerance intervals and their relationship among other interval estimates for one sample with one variance component, a computer simulation was conducted

50 iterations with data sets of size 50 • Observations were randomly generated, various interval estimates were constructed and compared

percentile) $\sqrt{n}$ 

#### PQRI Stability Shelf Life Working Group

• Variance is a linear combination of independent mean squares, df calculated

Interval that contains at least 100β% of population with given confidence

• Computed using factors from Normal and Chi-squared distributions  $\circ$   $\beta$ -content interval in models with only 1 source of variation are computed

□ *Patel* states 1-sided tolerance limits are directly related to 1-sided confidence

• Prediction intervals are compared with  $\beta$ -expectation intervals for models

•  $\beta$ -expectation interval equals prediction interval for both overall mean and

Formula for β-expectation interval reduces to formula for prediction interval

# □ The mean of each interval estimate across iterations is computed • Comparisons are made among the interval estimates





# **Characteristics of Interval Estimates**

- □ Interval estimates can be characterized into 3 groups:
- Group 1: Bounds on mean or on a CI endpoint o lowerupper PICI, lowerupper CICI, lower 1-sided CI, lower 2-sided CI, lowerlower CICI. lowerlower PICI
- Group 2: Bounds on individual observations, PI endpoints, or upper bound on lower  $\alpha/2$  percentile
- o lowerupper TI, lowerupper PIPI, lowerupper CIPI, lower 1 sided PI
- Group 3: Combination of bounds on individual observations, PI endpoints, or lower bound on lower  $\alpha/2$  percentile
- $\circ$  lower  $\beta$ -expectation (i.e. lower 2-sided PI), 2 1-sided lower TI,
- lowerlower TI, lower SAS TI, lowerlower CIPI, lower 1-sided TI, lower 1 sided TI with Bonferroni correction, lowerlower PIPI

#### Conclusions

- intervals
- $\square$  Prediction intervals equal  $\beta$ -expectation tolerance intervals for models with 1 variance component
- $\square$   $\beta$ -content intervals TI are wider than  $\beta$ -expectation TI
- □ 1-sided TI is directly related to a 2-sided TI on percentile
- □ Simulation results indicate
- Interval estimates can be characterized into 3 groups
- It must be determined the correct parameter of interest to be captured to determine which estimate to use



#### **University of Nebraska Department of Statistics**

• Current literature sources offer a wide variety of definitions for tolerance