# **ITC Tutorial: Design of Experiments and Analysis of Data**

by

Joel Tellinghuisen Department of Chemistry Vanderbilt University Nashville, TN 37235

## The Plan of Attack

- "5-minute statistics":
- Linear least squares:
- Nonlinear least squares:
- Application to ITC:

- Systematic Errors:
- Beyond the "black box":

Means, variances, and probability distributions The basics

What's new?

Optimizing parameters by least squares in "experiment design" mode Is the fit model right? Changing things Reporting results

### **5-Minute Statistics** Sampling Theory The mean $\overline{x} = \langle x \rangle = \frac{1}{n} \sum_{i=1}^{n} x_i$ $\langle x \rangle \equiv \mu = \int_{x_{\min}}^{x_{\max}} x P(x) dx$ The variance $s_x^2 = \frac{1}{n-1} \sum_{i=1}^n \delta_i^2 \quad (\delta_i = x_i - \overline{x}) \qquad \sigma_x^2 = \int_{x_{\min}}^{x_{\max}} (x - \mu)^2 P(x) dx$ The standard deviation $\mathcal{O}_{\mathbf{x}}$ $S_{\chi}$ The standard error (standard deviation in the mean) $\frac{S_{\chi}}{\sqrt{n}}$ $\frac{O_x}{\sqrt{n}}$

### **Probability Distributions**

**Uniform:**  $P(x) = \text{constant} (a \le x \le b); 0 \text{ otherwise}$ 

Normal: 
$$P_G(\mu,\sigma;x) = \frac{1}{\sigma\sqrt{2\pi}} \exp\left[-\frac{(x-\mu)^2}{2\sigma^2}\right]$$

**Poisson:** governs counting —  $\sigma^2 = \mu$  (= # counts).

Chi-square  $(\chi^2)$ :sampling estimates of variances.*t*-distribution:confidence limits for sampling<br/>estimates of parameters.

**NOTE:** Poisson,  $\chi^2$ , and *t*-distributions all become Gaussian in the limit of large  $\mu$  (Poisson) or *v* (degrees of freedom, *n*-*p*).

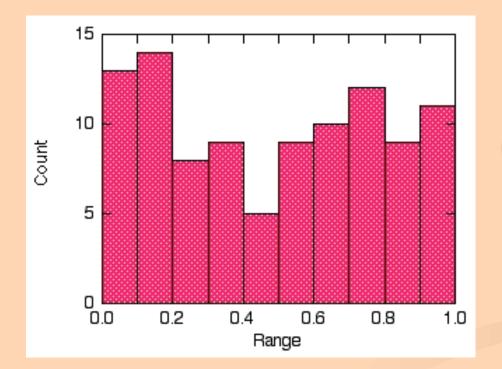
#### **Important probabilities**

$$\int_{\mu-\sigma}^{\mu+\sigma} P_G(\mu,\sigma;x) \, dx = 0.683$$

$$\int_{\mu-2\sigma}^{\mu+2\sigma} P_G(\mu,\sigma;x) \, dx = 0.954$$
 
$$\mu-2\sigma$$

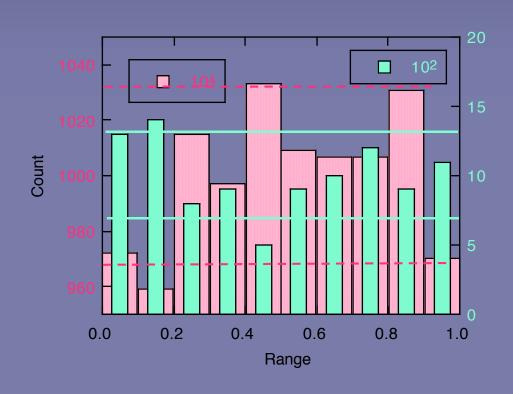
### Illustrations

The uniform distribution is the basis of computer random number generators. By default, the range is 0 < x < 1, for which  $\mu = \frac{1}{2}$  and  $\sigma^2 = \frac{1}{12}$ . Let's check ...



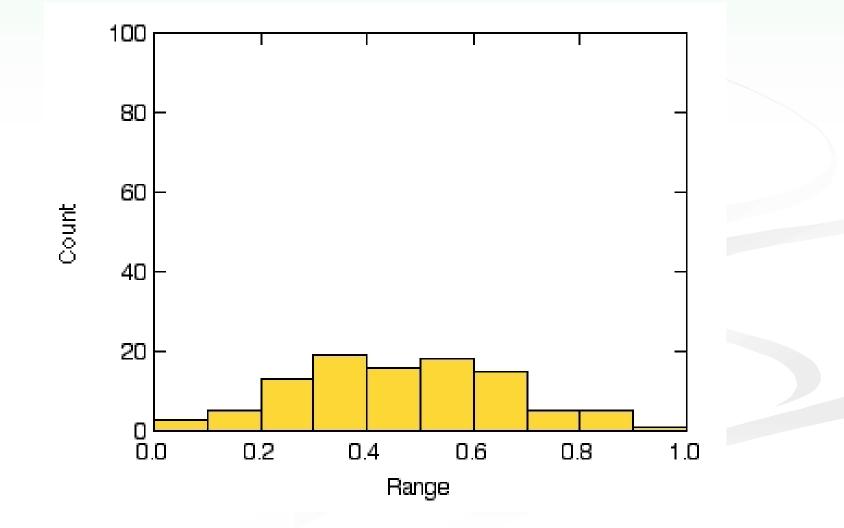
Minimum	9.36e-05
Maximum	0.99998742
Sum	5022.7734
Points	10000
Mean	0.50227734
Median	0.50219405
RMS	0.57845803
Std Deviation	0.28694843
Variance	0.082339402
Std Error	0.0028694843
Skewness	-0.0081906446
Kurtosis	-1.1866255

#### **Closer look at the binning statistics**

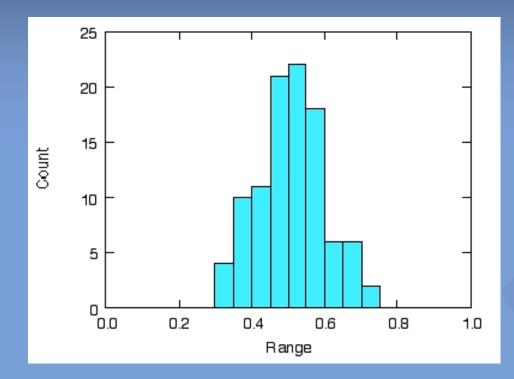


- Compare results for  $N = 10^4$  and  $10^2$ .
- Recall  $\sigma \approx \sqrt{n}$  (Poisson).
- Thus about 2/3 of bin counts should fall within  $\pm \sigma$  (32,3.2) of the expected values (Gaussian approx.).

### Next, bin average of 2 random numbers



#### **And now 12 ...**



#### Stats for sum of 12

Minimum	2.6777
Points	10000
Mean	6.0038
Median	5.9946
RMS	6.0863
Std Deviation	0.9989
Variance	
Std Error	0.009988

Results the very important Central Limit Theorem: Distributions of sums become normal, no matter what the parent distribution, as long as it has finite variance.