



# Potency

---

Calibration and Scoring  
for the PCCL  
September 17, 2003



# Purpose

---

- Examine the distribution of potency values for a set of contaminants that are representative of chemicals likely to be in the CCL Universe
- Utilize the knowledge gained to calibrate one or more approaches to scoring potency for the PCCL



# Learning Set Composition

---

- Regulated chemicals
- Unregulated chemicals with lifetime health advisories
- Nutrients/food additives with toxicity values similar to lifetime health advisories.



# Potency Values Collected

---

- Reference Dose (RfD)
- E-4 risk concentration in water
- NOAEL from the critical study
- LOAEL from the critical study
- Rat Oral LD<sub>50</sub>



# Sources of Information

---

- Integrated Risk Information system (IRIS)
- Office of Water (OW) Health Advisories
- Institute of Medicine (IOM) Tolerable Upper Levels (ULs) for Nutrients



# Data Set Characteristics

---

- 216 chemicals
- 185 RfDs
- 51 E-4 risk concentrations
- 149 Critical NOAELs
- 152 Critical LOAELS
- 171 LD<sub>50</sub>s
- Most potent dioxin
- Least potent dietary phosphorous



# Procedure

---

- Enter the potency values into a spreadsheet
- Divide the range of potency values into tenths and array the potencies using a histogram
- Take the rounded  $\text{Log}_{10}$  for each potency value and array the potencies using a histogram



## Procedure (contd.)

---

- Calibrate scoring equations for each data set that equates the modal  $\text{Log}_{10}$  of the potency value to a score of 5 on a 1-10 potency scale.
  - This was the process used for the March, 2003 algorithm exercise
- Test the scoring equations for each type of potency value and examine whether or not the scores agree.

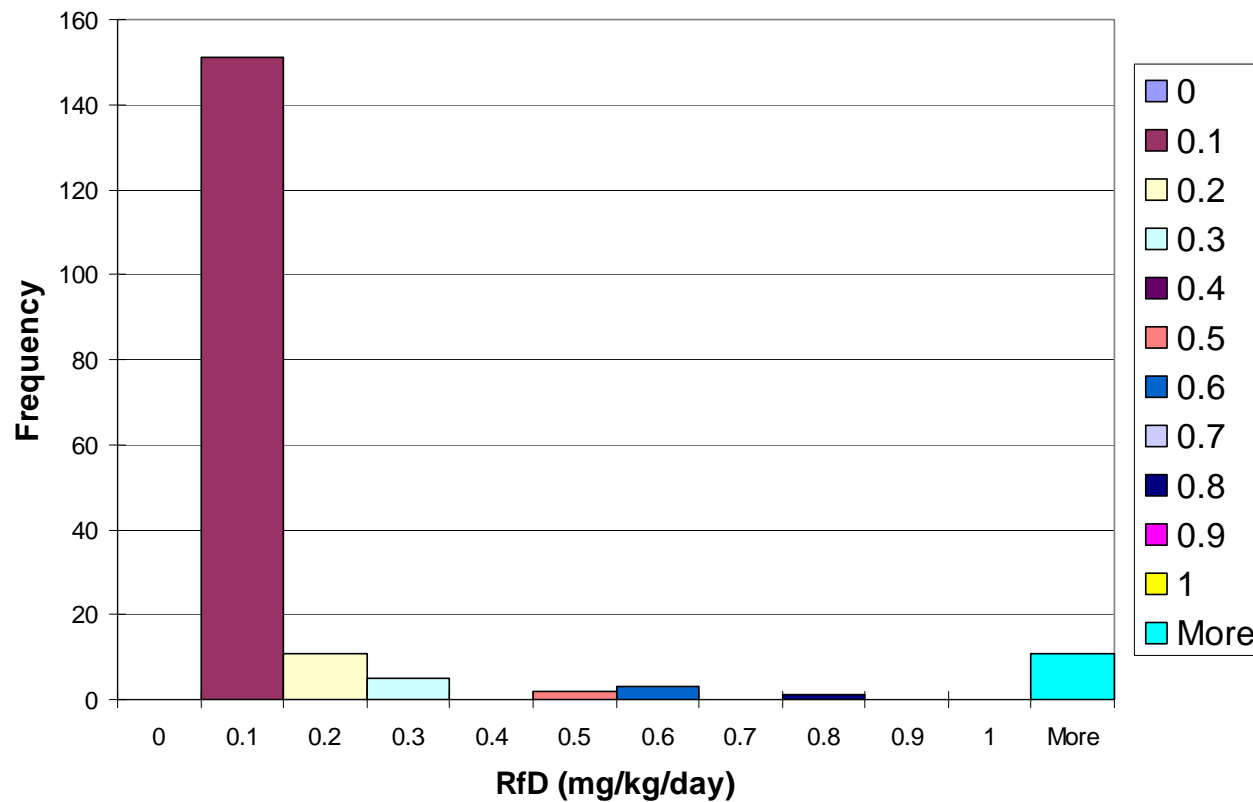




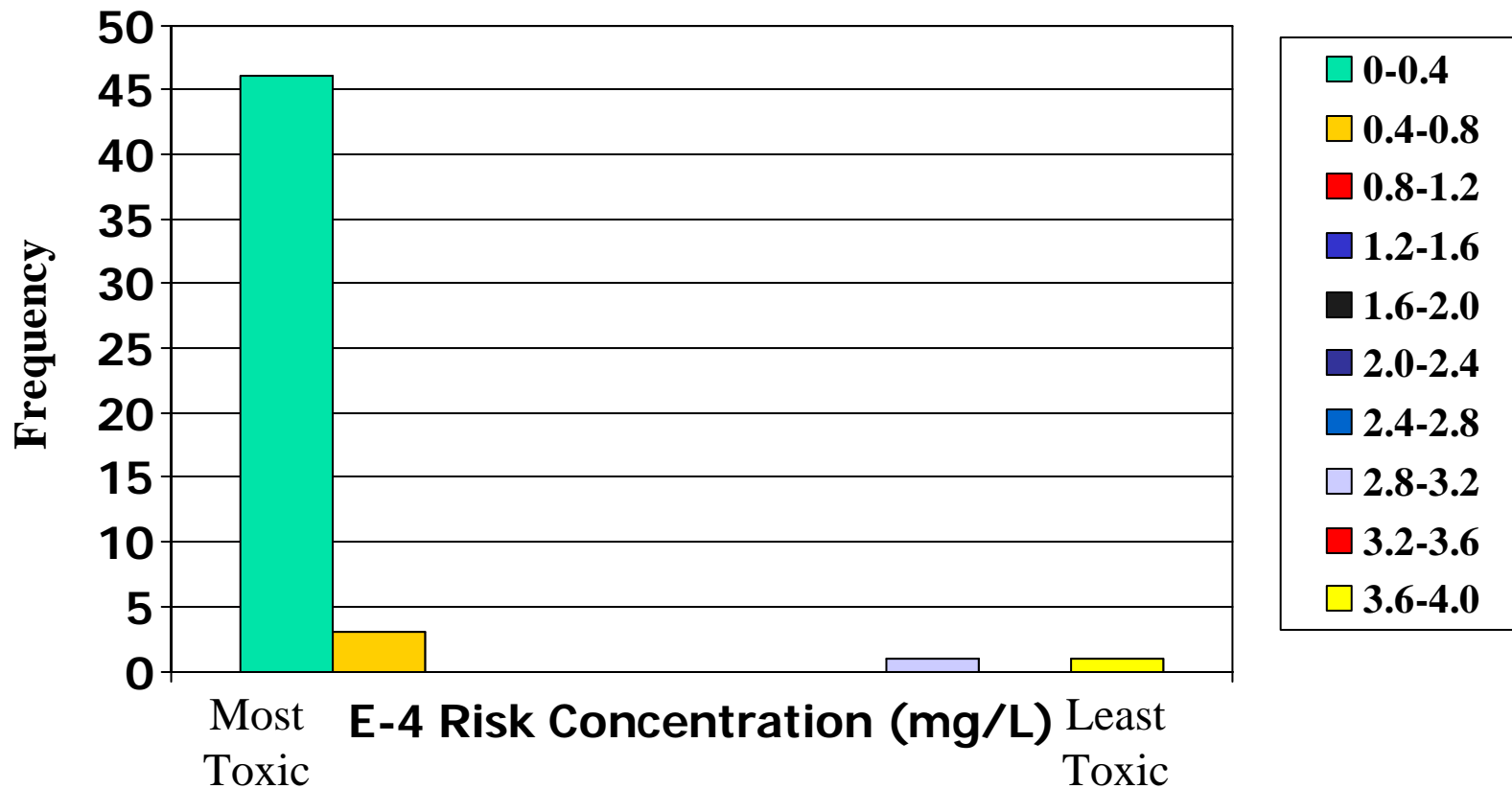
# Results - Histograms

---

# RfD Distribution by Deciles

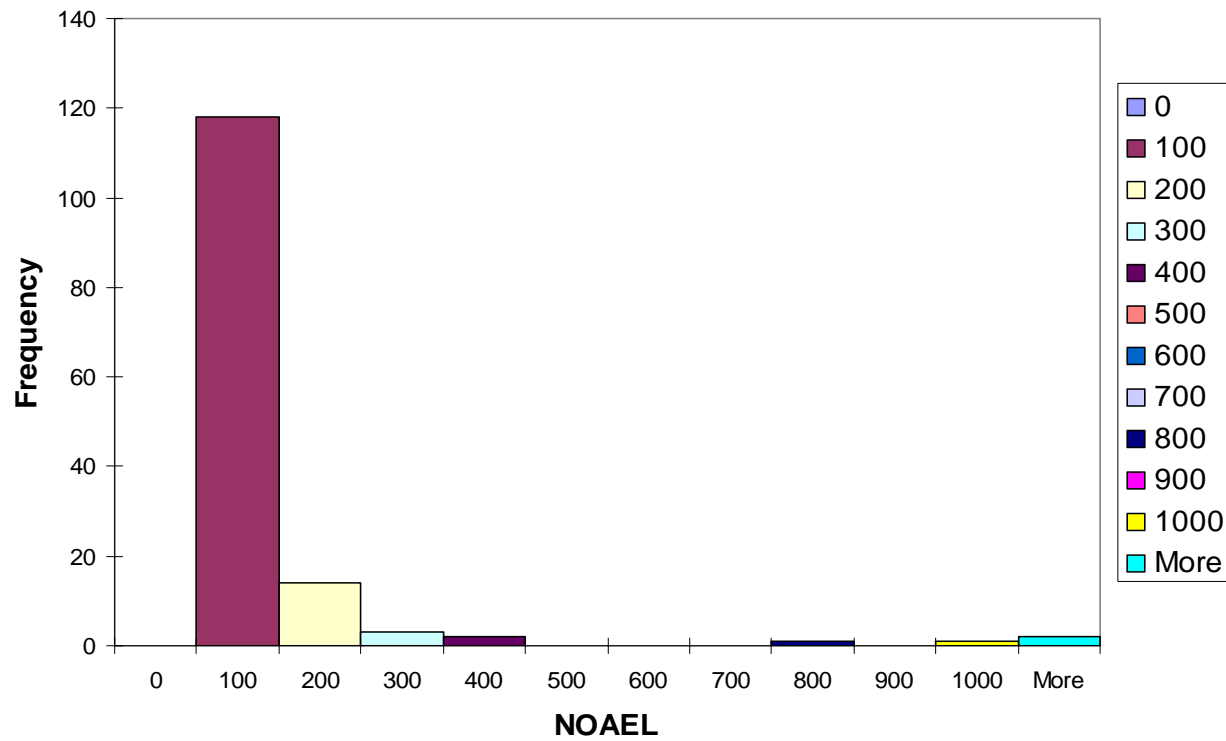


# Distribution of Concentration for E<sup>-4</sup> risk by Deciles

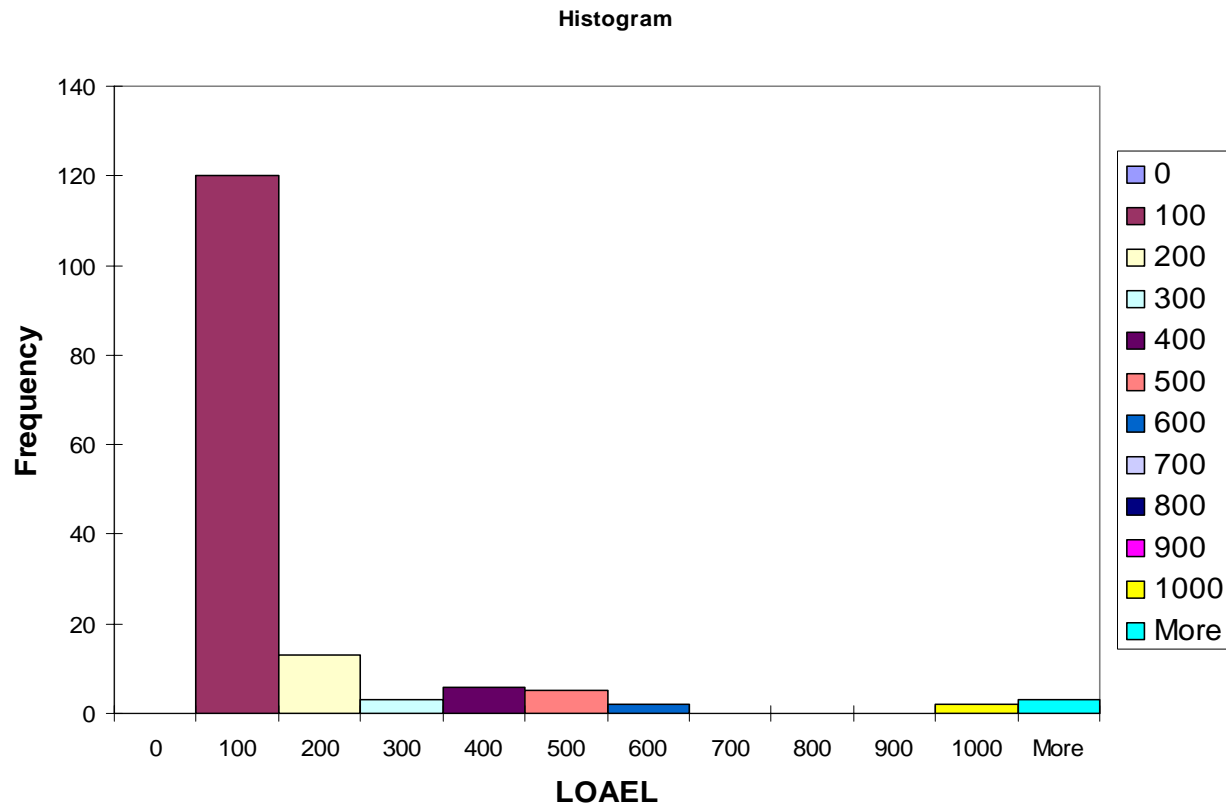


# NOAEL Distribution by Deciles

Histogram

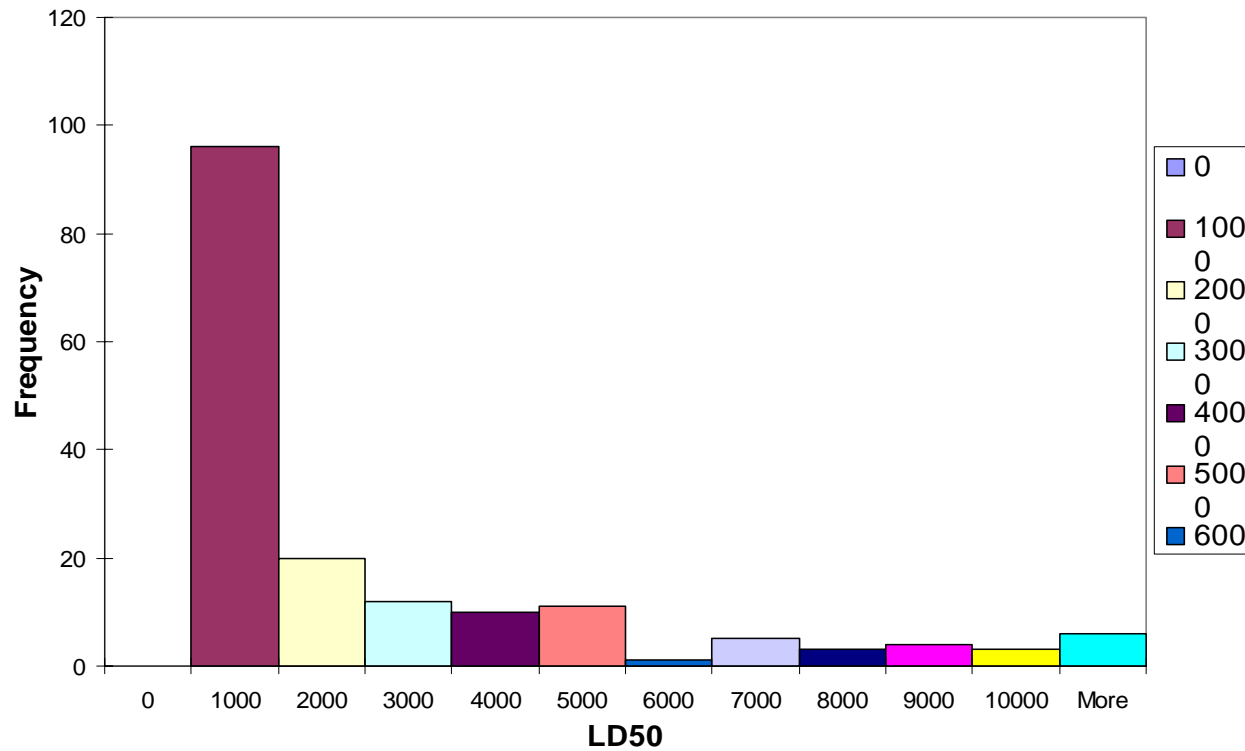


# LOAEL Distribution by Deciles

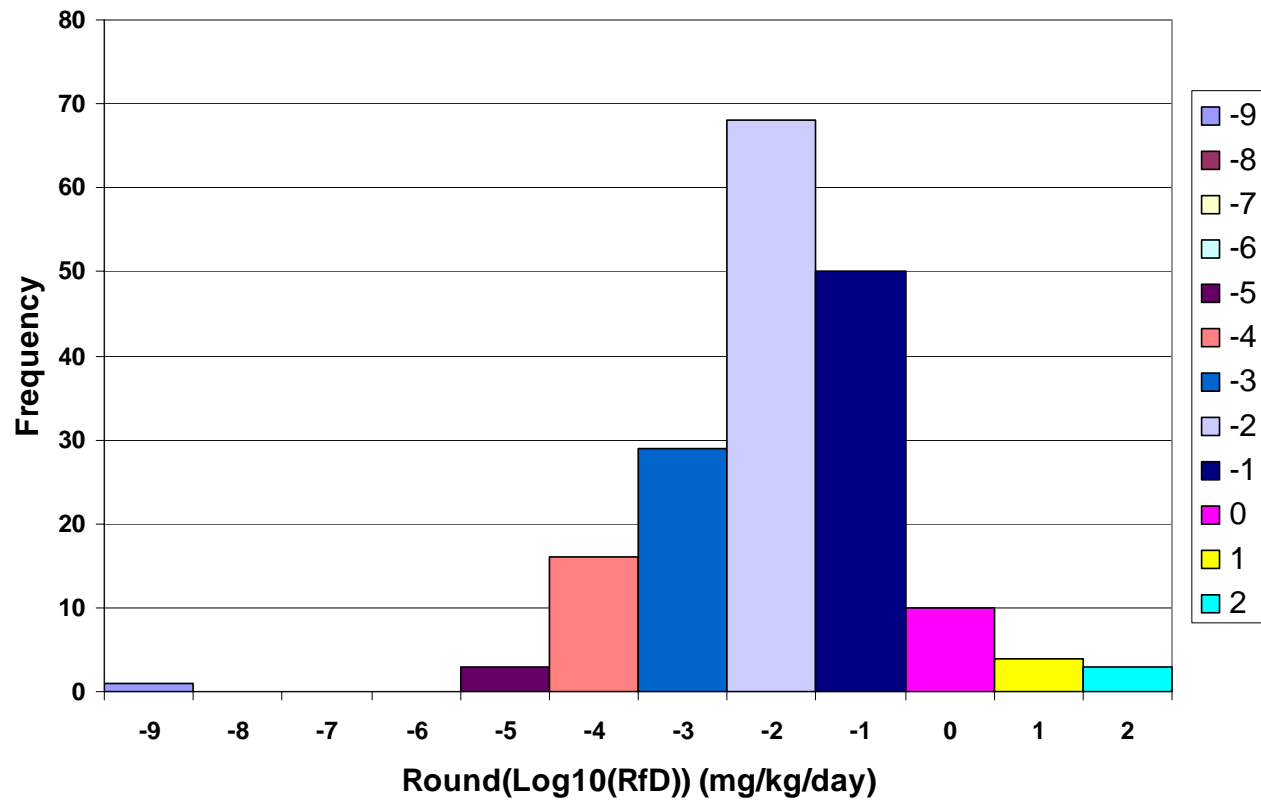


# LD50 Distribution by Deciles

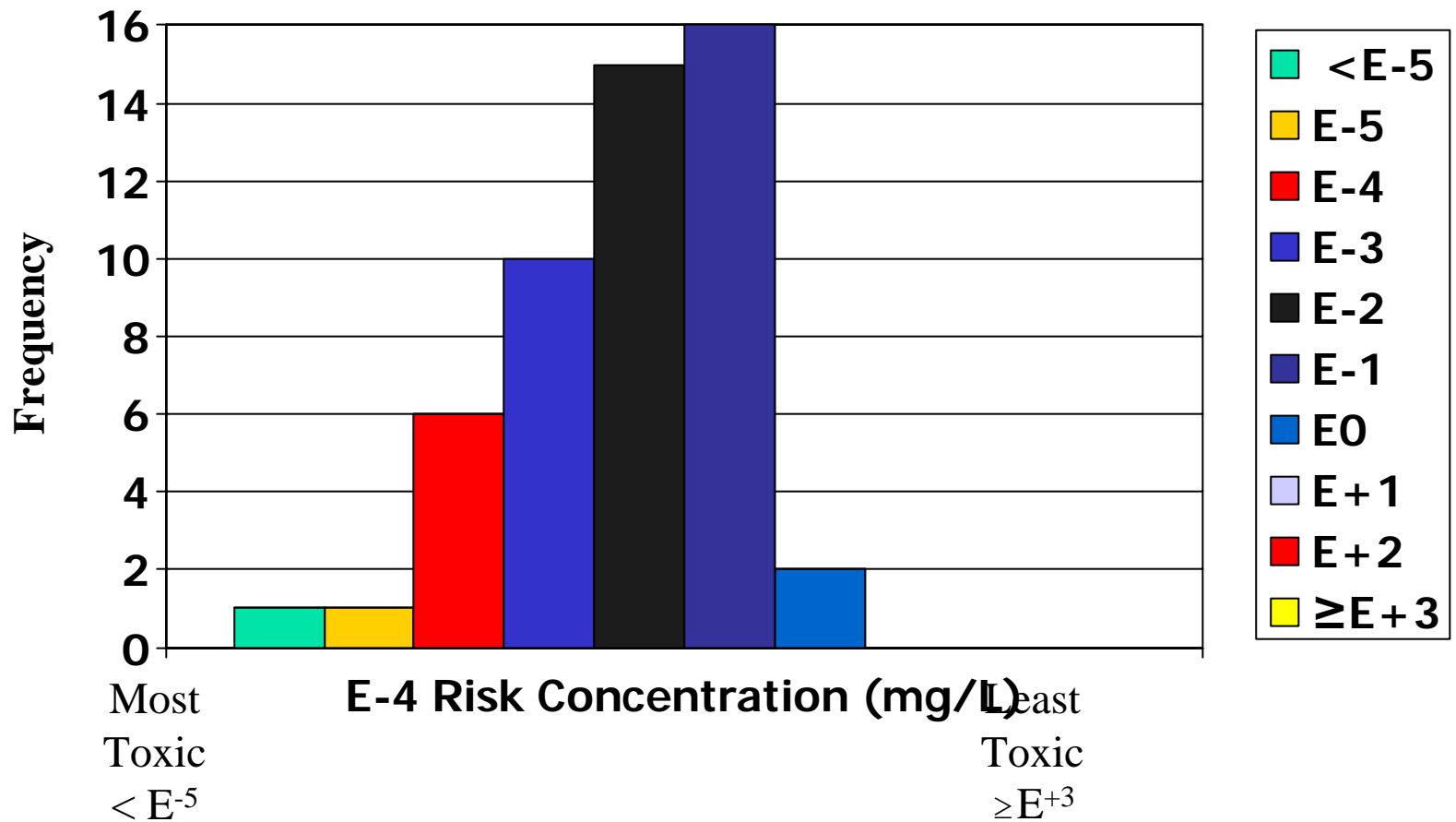
Histogram



# RfD Distribution – Rounded $\log_{10}$



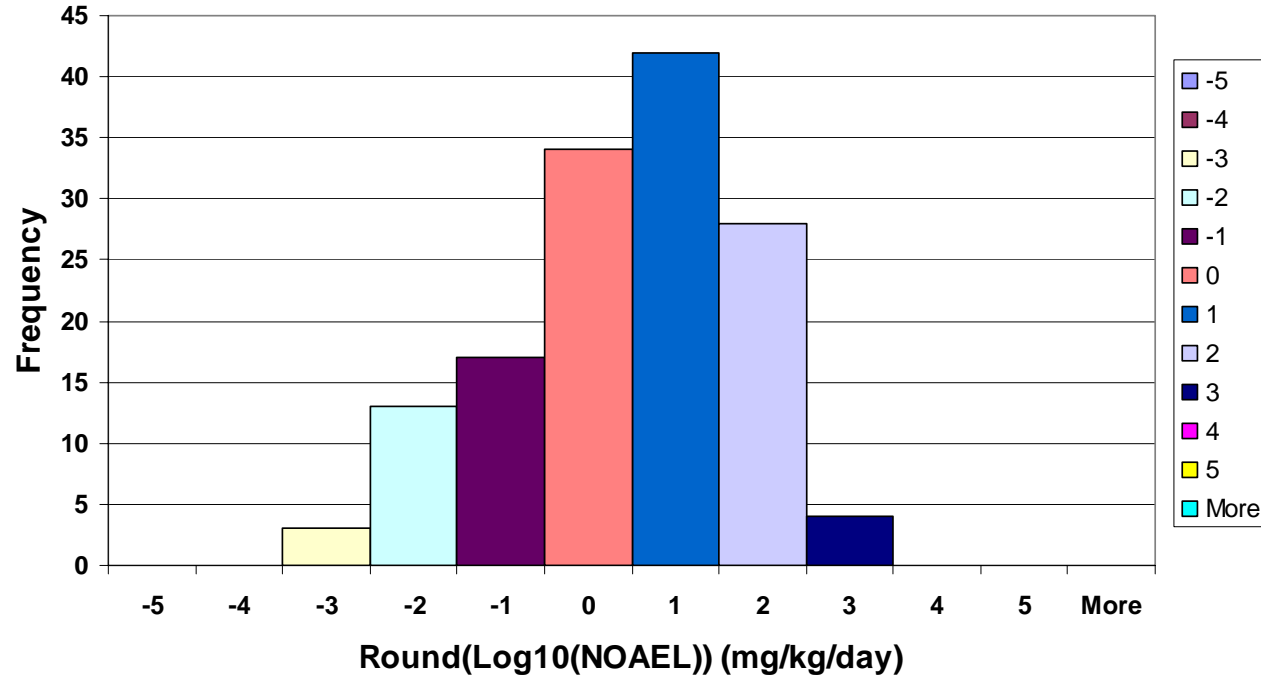
# E-4 Risk Concentration Distribution - Rounded Log<sub>10</sub> Scale





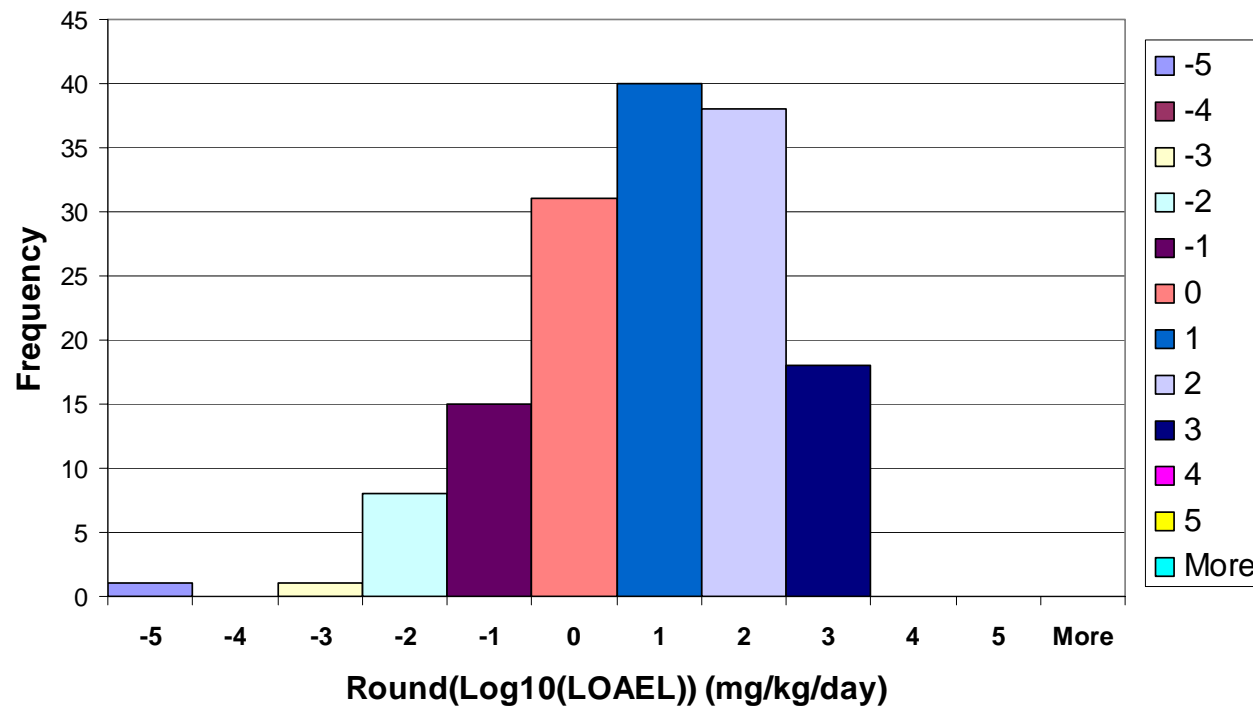
# NOAEL Distribution – Rounded

$\text{Log}_{10}$



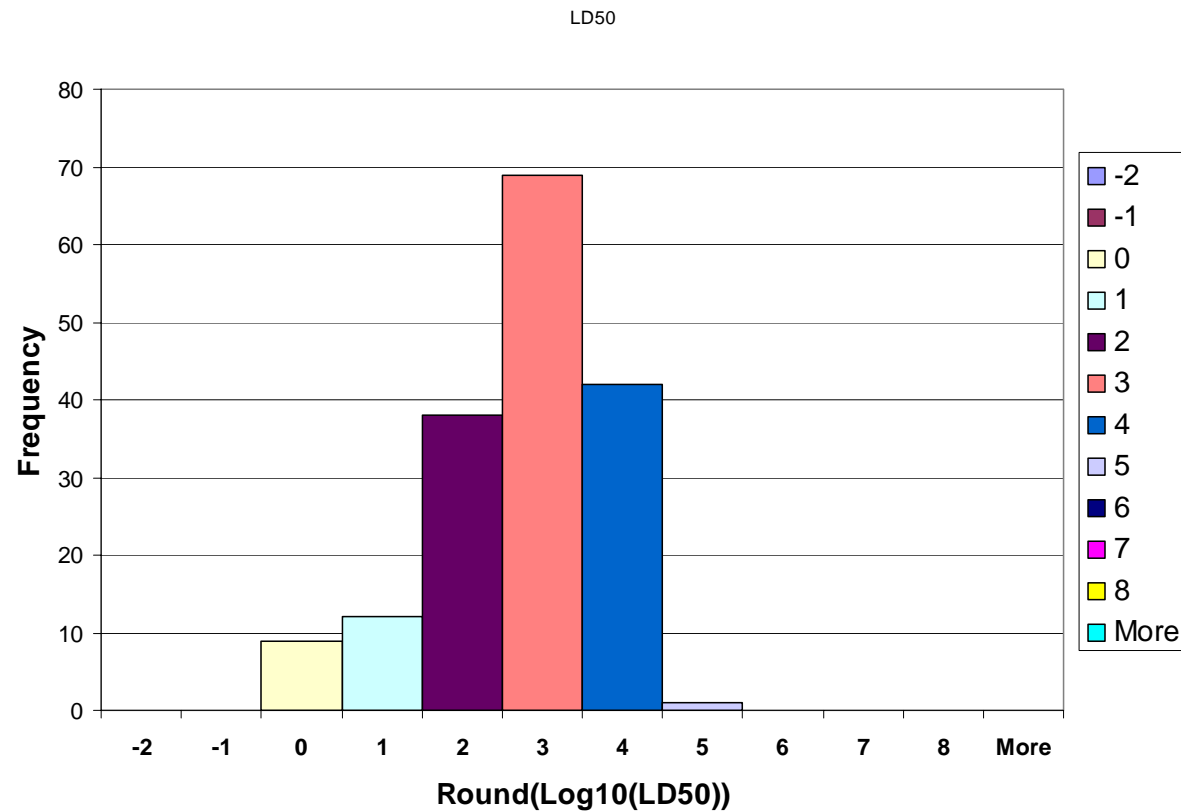
# LOAEL Distribution – Rounded

$\text{Log}_{10}$



# LD50 Distribution – Rounded

$\text{Log}_{10}$





# Scoring Equations

---



# Method

---

- Base Equation
  - $5 = 10 - (\text{modal } \log_{10} \text{ of potency value} + X)$
- Derivation of RfD equation
  - $5 = 10 - (-2 + X) \quad X = +7$
  - RfD-based Score =  $10 - (\text{Log}_{10} \text{ of RfD} + 7)$
- Other values of X
  - E-4 risk concentration:  $X = +6$
  - NOAEL:  $X = +4$
  - LOAEL:  $X = +4$
  - LD50:  $X = +2$



# Scoring Results

---



## Examples of Scoring (contd)

Chemical	RfD	NOAEL	LOAEL	LD50
Dioxin	10		10	4
Phosphorous	1	4		7
Methyl ethyl ketone	3	3	3	5
Phenol	4	4	4	5
Hexazinone	4	5	4	5
Iodine	5	8	8	4



# Examples of Scoring

Chemical	RfD	NOAEL	LOAEL	LD50
Baygon	5	-	6	6
Dacthal	5	6	5	5
Ethylene Glycol	3	4	3	4
Silver	5	-	8	4
Paraquat	5	6	6	6
Calcium	1	-	4	-





# Conclusions

---

- Scores are fairly consistent for a given chemical.
- Low uncertainty factors increase the spread of scores between RfD, NOAEL and LOAEL
- LD50s for inorganics must be for a relevant form of the chemical
- Options exist for refining the process



# Options for Refinement

---

- Looking at other distributions of the learning set data.
- Expanding the learning set
- Centering the scoring scale on the median unrounded  $\text{Log}_{10}$  value.
- Examining other approaches to using the learning set distributions to calibrate scoring and comparing the results.