

# Meta-analysis

## Combining Data

IB/NRES 509 Statistical Modeling

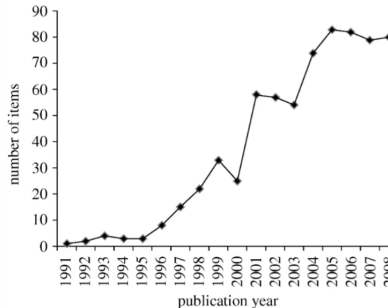
# What is a Meta-analysis?

- ▶ A quantitative synthesis of previous research
- ▶ Studies as individual observations, weighted by  $n$ ,  $\sigma^2$ , quality, etc.
- ▶ Can combine heterogeneous data from heterogeneous studies
- ▶ Evaluate means, responses, sources of heterogeneity

## When is Meta-analysis used?

- ▶ To synthesize results once a large body of work has been conducted
- ▶ To investigate large-scale patterns beyond the scope of individual experiments
- ▶ Today we will look at two examples:
  - ▶ evaluating the hypotheses that N limits NPP
  - ▶ synthesizing trait data to parameterize a DGVM

# Meta-analysis is increasing in Ecology and Evolution



**Figure:** Number of meta-analyses articles published in ISI journals in Ecology and Evolution through 2008. Stewart G Biol. Lett. 2010;6:78-81

# Meta-analysis Overview

- ▶ Develop a research question
- ▶ Study Literature
- ▶ Define Predictor and Response Variables
- ▶ Define Model
- ▶ Collect Data
- ▶ Calculate Response Metric and Weights
- ▶ Assess the effect of unpublished data
- ▶ Interpret, present at conference, publish...

# Literature Search

- ▶ Search methodology:
  - ▶ search terms / phrases
  - ▶ follow references
  - ▶ contact researchers
- ▶ Study Selection
  - ▶ apriori criteria for inclusion:
    - ▶ study design
    - ▶ years of publication
    - ▶ language
    - ▶ sample size
    - ▶ availability of relevant information
  - ▶ block results by quality, conduct analyses in groups
  - ▶ quality based weights

# Choosing Predictor Variables

- ▶ Depends on Question, available data
- ▶ Experimental vs. Observational Data
- ▶ Continuous vs. Categorical Data

## Choosing Response Variables

### 1. Mean

$$\bar{X}$$

### 2. Differences in means

$$d = \bar{X}_{trt} - \bar{X}_{Ctl}$$

### 3. Cohen's g

$$g = \frac{\bar{X}_{trt} - \bar{X}_{Ctl}}{s_p}$$

where  $s_p$  is the pooled sample standard deviation, small n can bias estimate of  $\sigma$  and thus  $g$

### 4. Response Ratio proportionate change

$$\ln R = \log(\bar{X}_T / \bar{X}_C)$$

can be used in competition intensity, relative yield, relative crowding



# Fixed Effects Model

- ▶ Fixed Effects Model:

$$T_i \sim N(\Theta_i, \sigma_i^2)$$

$$\Theta_i = \mu$$

- ▶ where  $\Theta_i$  is the study mean and  $\sigma_i^2$  is the within study variance
- ▶ assumes no between-study variability; all studies have same  $\mu$ 
  - ▶ can be used if RE model shows negligible among-study variance ( $\tau^2 \rightarrow 0$ )
  - ▶ otherwise, inference limited to studies in meta-analysis and studies with identical characteristics
- ▶ Difficult to justify in ecological context.

## Random Effects Model

- ▶ Random Effects Model:

$$T_i \sim N(\Theta_i, \sigma_i^2)$$

$$\Theta_i \sim N(\mu, \tau^2)$$

- ▶ where  $\Theta_i$  is the study mean,  $\mu$  is the grand mean,  $\sigma_i^2$  is the within study variance and  $\tau^2$  is the among-study variance
  - ▶ acknowledges study-specific effects due to location, experimental conditions, etc.
  - ▶ models both within and among study variance
- ▶ As a regression model:

$$\mu = \mu_i = \beta_0 + \beta_1 X_{1i} + \beta_2 X_{2i} + \dots + \beta_n X_{ni}$$

- ▶  $X$ 's can be continuous (linear regression), categorical (e.g. ANOVA) or a combination (mixed effects)

# Study Weights

- ▶ Give greater weight to experiments with smaller standard error:

- ▶ fixed effects model

$$w_i = 1/(\hat{\sigma}_i^2)$$

- ▶ random effects model

$$w_i = 1/(\hat{\sigma}_i^2 + \hat{\tau}^2)$$

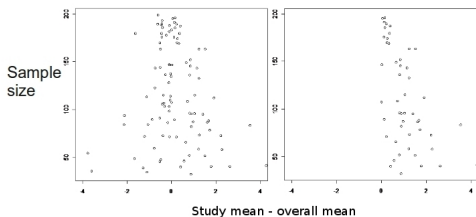
- ▶ Weighted Mean Effect

$$\bar{T} = \frac{\sum w_i T_i}{\sum w_i}$$

# Sources of Bias

- ▶ Publication Bias
  - ▶ studies that show no effect are less likely to be published
  - ▶ “file drawer problem”
  - ▶ Create a funnel plot with effect size versus sample size
- ▶ Unconducted Studies
  - ▶ researchers are less likely to ask a question if they think that the answer is known
  - ▶ e.g. few N fertilization studies have been conducted in the tropics
- ▶ Study Heterogeneity
  - ▶ need to ensure that effect sizes are standardized and comparable across studies
  - ▶ e.g. a large variety of methods may be used, sites and organisms studied may vary

# Assessing Study Bias



**Figure:** Use of a funnel plot to evaluate the file-drawer problem; x axis is deviance from mean effect size ( $T_i - \hat{\mu} = \varepsilon; \varepsilon \sim N(0, \tau^2)$ ) as sample size increases, the variance around the mean effect size (here set to 0 by subtracting) in effect size should decrease. left: no evidence for missing studies, right: missing studies

# Approaches to Inference in Meta-analysis

- ▶ Moment Matching
- ▶ Maximum Likelihood
- ▶ Bayesian

# Moment Matching

## ▶ Moment Matching

- ▶ given data, calculate moments ( $\hat{\mu} = T_i = n \sum x_i$ ,  $\hat{\sigma}^2 = S_i^2$ )
- ▶ estimate global group means
- ▶ partition among vs. within study variance
- ▶ adequate for simple models, but not, e.g. multiple regression

# Analysis of N-fertilization Studies

- ▶ Question: Does Nitrogen limit Terrestrial Net Primary Productivity?
  - ▶ Widely accepted to be true, except for tropical forests
  - ▶ Many previous studies, no global synthesis
- ▶ Approach
  - ▶ Meta-analysis of N-fertilization studies
  - ▶ Regression of N on  $\ln R$



# Analysis of N-fertilization Studies

- ▶ Response Metric

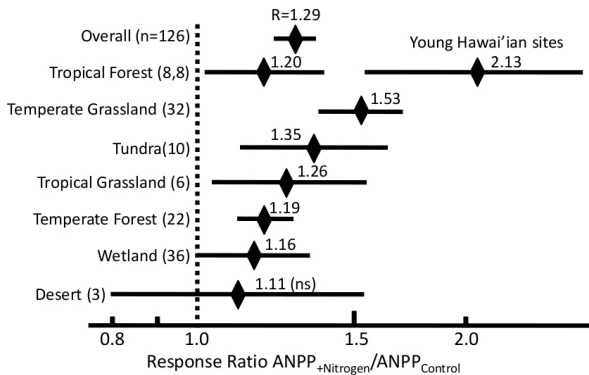
$$\ln R = \frac{ANPP_{+N}}{ANPP_C}$$

- ▶ Measures of ANPP
  - ▶ Grasslands & Wetlands: Peak Biomass, Repeated Harvests
  - ▶ Forests:  $\Delta$ Volume,  $\Delta$ Diameter,  $\Delta$ Basal Area, Litterfall

## Example 1: Does Nitrogen Limit NPP?

## Analysis of N-fertilization Studies

## Biome Level Responses to N



# Maximum Likelihood

- ▶ Maximum Likelihood
  - ▶ choose parameters that maximize  $p(\text{data}|\text{model})$
  - ▶ allows more complex models, non-normal distributions, parameter uncertainty
  - ▶ simultaneous estimation of all parameters, more inference options

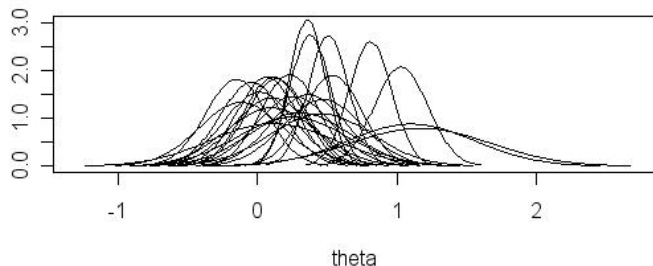
## Maximum Likelihood Example

- ▶ Given data, what is the pdf of different parameter values?
- ▶ equivalent to MM for simple models, but more parameters can be estimated, e.g. each  $\Theta_i$
- ▶ Random Effects Model:

$$T_i \sim N(\Theta_i, \sigma_i^2)$$

$$\Theta_i \sim N(\mu, \tau^2)$$

## Maximum Likelihood Example



**Figure:** Likelihood of observing the data ( $T_i$  and  $S_i^2(\hat{\sigma}^2)$ ) from each of 25 studies, given different values of  $\Theta$

# Bayesian Meta-analysis

- ▶ posterior is the Likelihood of observing parameters given the data
- ▶ like MLE, can estimate all parameters simultaneously
- ▶ Much more flexible

# What Values Might A Plant Trait Hold?

## A Bayesian Approach to DGVM Model Parameterization

### ► Overview

- Plant traits are used to parameterize models of vegetation, ie: ED2
- Std. Practice - select single values from the literature
- Our goal - parameterize model with meta-analysis posterior distributions

### ► Response Metric:

- Trait Means of 24 (and counting) traits

# Plant Trait Meta-analysis Model

$$T_i \sim N(\Theta_i, \sigma_i^2)$$

$$\Theta_i \sim N(\mu, \tau^2)$$

## Informed Priors

$$\mu \sim \text{distn}(a, b)$$

$$s_i^2 \sim IG\left(\frac{n_i}{2}, \frac{n_i}{2} \sigma_i^2\right)$$

## Priors

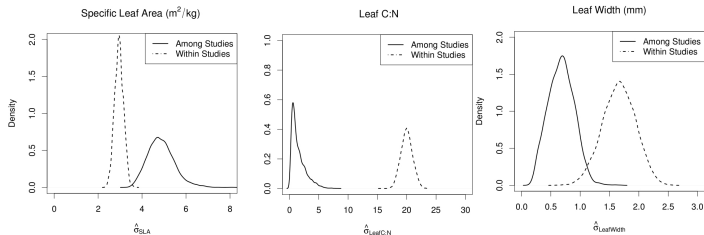
$$\tau^2 \sim IG(0.1, 0.1)$$

$$\sigma^2 \sim IG(0.1, 0.1)$$



## Example 3: Synthesis of Plant Traits

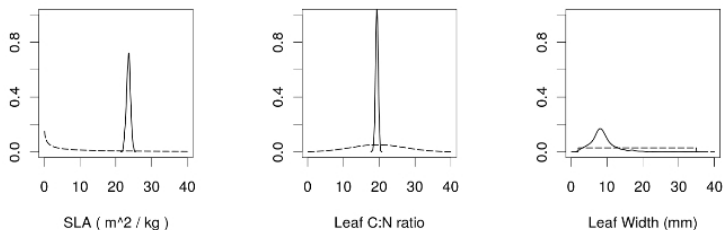
## Within vs. Among Study Variance



**Figure:** Distribution of among ( $\tau^2$ ) and within ( $\sigma^2$ ) study variance of three plant traits.

## Example 3: Synthesis of Plant Traits

# Distribution of Priors and Posterior Means



**Figure:** Distribution of prior (dashed line) and posterior (solid line) means of three plant traits.



Example 3: Synthesis of Plant Traits

# Model Forecast With Uncertainty

## A Bayesian Approach to Model Parameterization

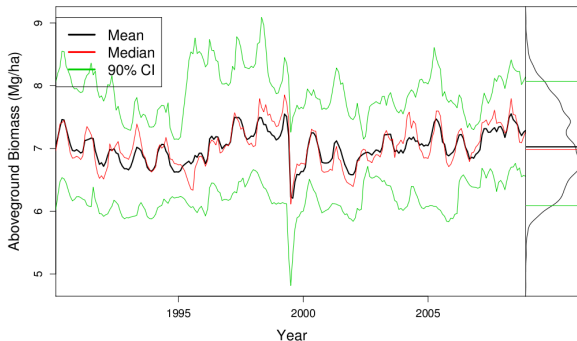


Figure: Model prediction with uncertainty estimate.