

# A Hands on Introduction to Systematic Reviews and Meta-Analysis

Part 2: Meta-Analysis (Summarizing the Data)



# Effect Size

- ▶ An *effect size* quantifies the magnitude of the relationship among variables
  - For example, let's say we are comparing 10 boys and 10 girls on reading speed
    - Boys:  $M = 36$ ,  $SD = 7$
    - Girls:  $M = 29$ ,  $SD = 6$
- ▶ Unstandardized Effect Size
  - $M_{\text{diff}} = M_{\text{Boys}} - M_{\text{girls}} = 36 - 29 = 7$
  - This is interpretable if the units are interpretable (e.g., if reading speed was measured in seconds or minutes)

# Effect Size

## ▶ Standardized Effect Size

- E.g., Cohen's  $d$ 
  - Quantifies differences in means in 'standard deviation' units

$$d = \frac{M_{Boys} - M_{Girls}}{\sqrt{\frac{(n_{Boys}-1)SD_{Boys}^2 + (n_{Girls}-1)SD_{Girls}^2}{n_{Boys} + n_{Girls} - 2}}} = \frac{36 - 29}{\sqrt{\frac{(10-1)7^2 + (10-1)6^2}{10+10-2}}} = 1.07$$

- Thus, boys and girls differ by a little more than one SD
- This is interpretable regardless of the units of measurement, and is comparable across studies which use different scales, measures, etc.
  - Which will obviously be useful in meta-analysis

# Effect Size

- ▶ Example 2: Correlation between income and depression
- ▶  $r = .24$ 
  - Increasing income by one standard deviation is, on average, associated with a .24 increase in depression
- ▶ Since correlation values are inherently standardized (range from  $-1$  to  $1$ ), we would almost always adopt a standardized metric to explore the correlation among variables

# Confidence Intervals for Effect Sizes

## ▶ Confidence Interval (CI)

- A range of values over which we expect the true (population) parameter to fall
- E.g., 95% CI
  - If we sampled repeatedly from the population and calculated a CI for each effect size from each sample, 95% of the CIs would contain the population parameter
- Importance of Confidence Intervals
  - CIs provide information regarding measurement precision
    - E.g. 1:  $d = .28$ ; 95% CI =  $\{.27, .29\}$
    - E.g. 2:  $d = .28$ ; 95% CI =  $\{.15, .41\}$
  - The second CI measures the effect of interest much less precisely, even though the effect size is the same
- In general, studies with larger sample sizes have narrower CIs

# Effect Size in Meta-Analysis

- ▶ Effect size is the Outcome/Dependent Variable
  - This will require the computation of effect sizes or transforming from one effect size to another
  - Standardized effect sizes are almost always used in meta-analysis
    - A standardized index must be comparable across studies, represent the magnitude and direction of the relationship of interest, and be independent of sample size
  - It is also possible to use unstandardized effect sizes, but this requires that the exact same scales/variables are used in each study (and that no transformations, modifications, etc. were made to any variables)

# Effect Size in Meta-Analysis

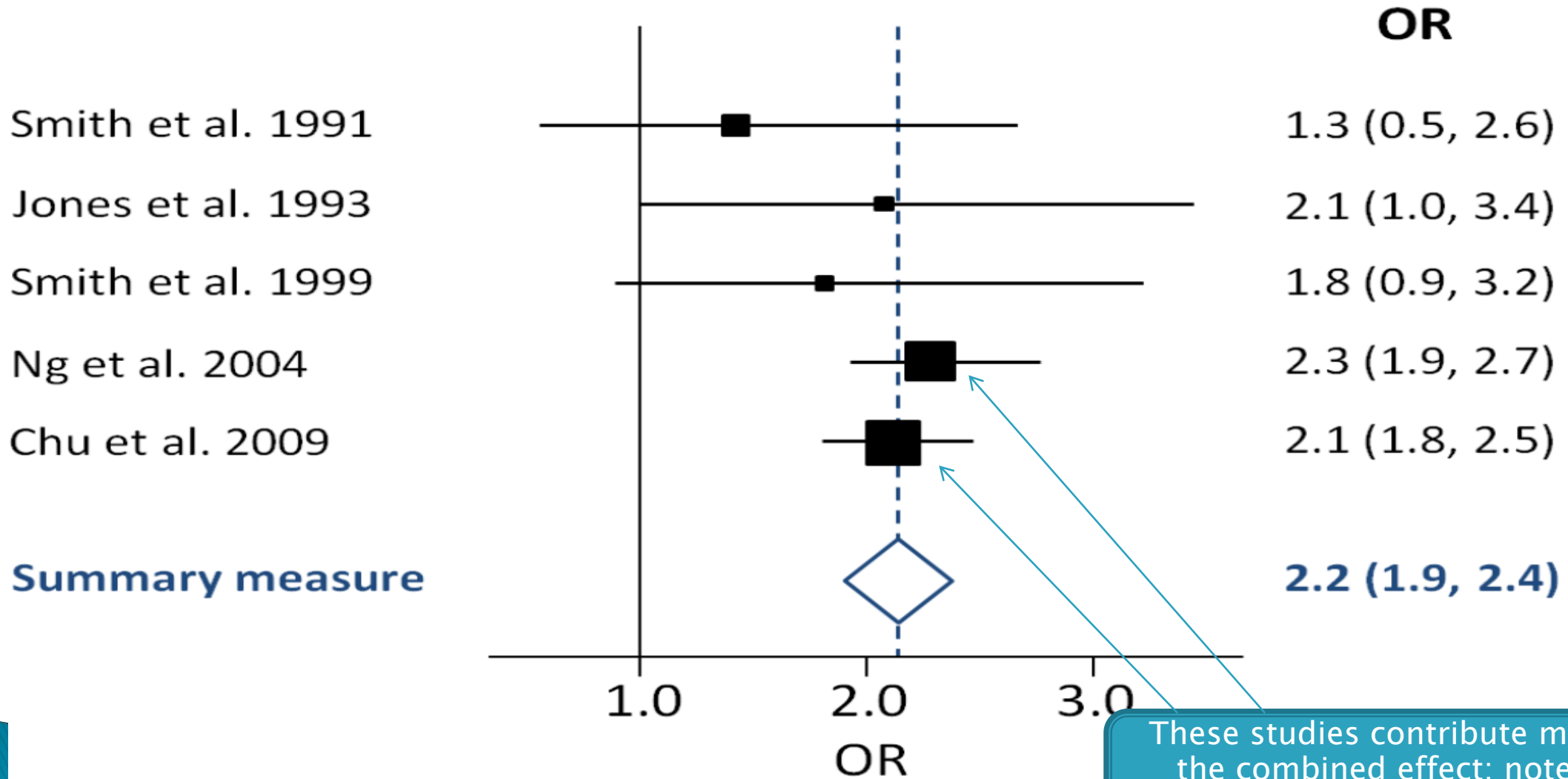
- ▶ Note that in some testing situations it might be tricky to obtain a proper effect size estimate
  - Take, for example, a study that looks at the difference between boys and girls in vocabulary development over 6 months from 18 months to 24 months using a repeated measures analysis
- ▶ Typical methods that convert  $t/z$  statistics to  $d$  will be incorrect for repeated measures studies, and corrections need to be applied in order to minimize bias

# Forest Plot

- ▶ A visual representation of effect sizes (and confidence intervals for the effect sizes) from multiple studies included in a meta-analysis
  - Recall: all effects must be measured in the same metric (e.g.,  $d$ , correlation)
- ▶ The area of the effect size icons (usually squares) on the plot indicates the “weight” of the study to the combined effect
  - E.g., larger N studies have a higher weight
- ▶ The plot also shows the *combined* effect size, and confidence interval for the effect size, across studies



# Forest Plot Example - Odds Ratios



These studies contribute more to the combined effect; note the narrow CIs (and likely large Ns)

# Statistical Models

- ▶ There are two popular models available for conducting a meta-analysis
  - In other words, two models available for arriving at a “combined” measure of effect size
  - Fixed Effects Model
    - Assumes that all the studies investigated the same population, and therefore estimate the same population effect size
      - Highly questionable
  - Random Effects Model
    - Allows for the possibility that the studies investigated somewhat different populations, and therefore estimate different population effect sizes
      - Another way to say this is that we expect some “true” variability in effect sizes

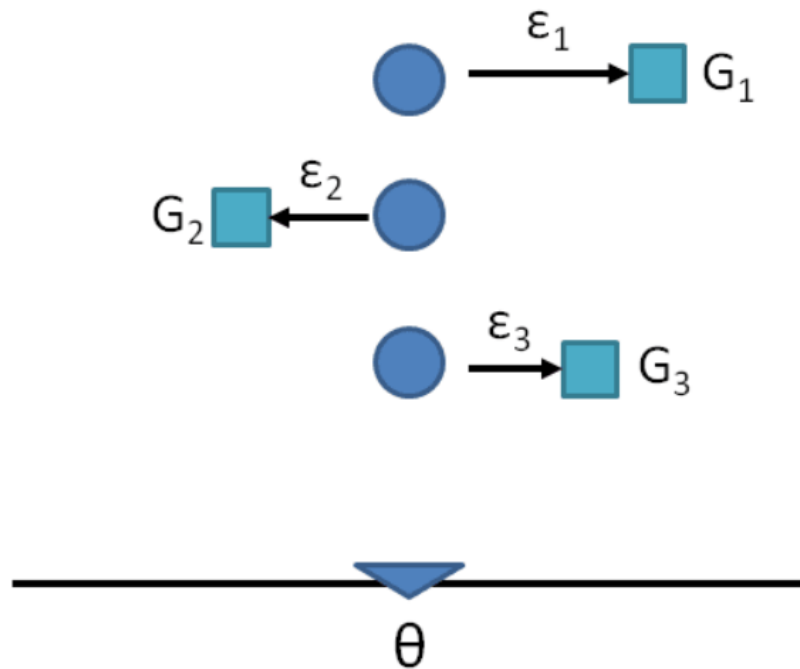
# Fixed Effects vs Random Effects

- ▶ It is difficult to imagine a setting in which multiple studies conducted in different locations, with different samples, and with potentially different measures are all studying the same population (and thus are after a single population effect size)
- ▶ The random effects model is more realistic and provides a basis for understanding the heterogeneity of effect sizes
  - Further, the models give the same answer if there is only a single population, so it is hard to find a reason for a researcher to prefer a fixed effects model

# Fixed Effects vs Random Effects

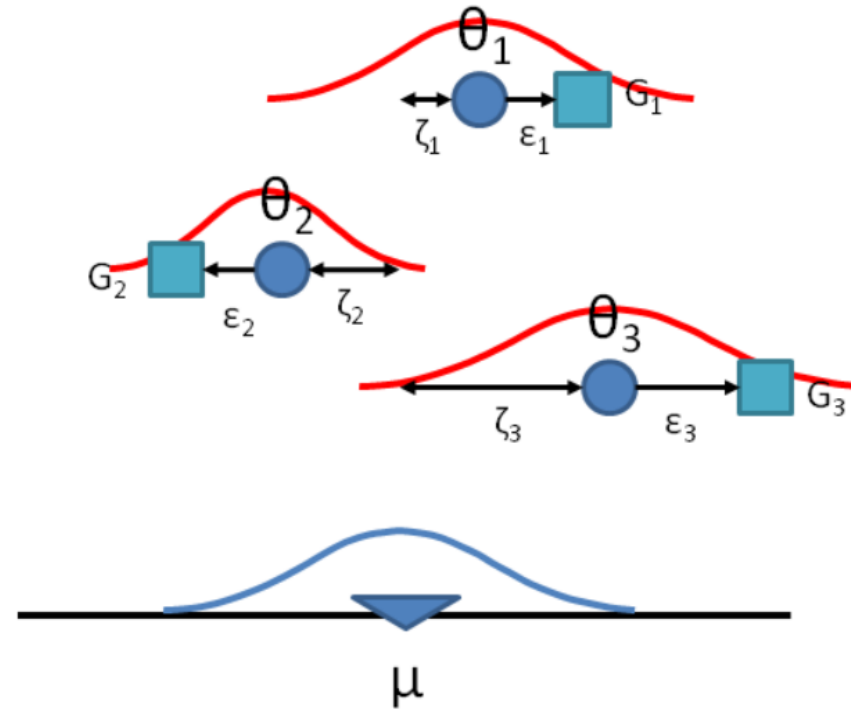
Fixed Effects Model

$$G_i = \theta + \varepsilon_i$$



Random Effects Model

$$G_i = \mu + \zeta_i + \varepsilon_i$$



# Fixed Effects Meta-Analysis

- ▶ For a set of  $S$  effect size measures ( $\gamma$ )

- $\hat{Y}_F = \frac{\sum_{i=1}^S w_i \hat{Y}_i}{\sum_{i=1}^S w_i}$

- $w_i = \frac{1}{s^2(\hat{Y}_i)}$

- $s^2(\hat{Y}_F) = \frac{1}{\sum_{i=1}^S w_i}$

This info is used to generate a mean effect size and a CI around the mean effect size

# Fixed Effects Meta-Analysis Example

- ▶ Study 1:  $M_1 = 12$ ,  $M_2 = 14$ ,  $SD_1 = 3$ ,  $SD_2 = 3$ ,  $n_1 = 22$ ,  $n_2 = 32$
- ▶ Study 2:  $M_1 = 14$ ,  $M_2 = 16$ ,  $SD_1 = 2$ ,  $SD_2 = 2$ ,  $n_1 = 25$ ,  $n_2 = 52$
- ▶ Study 3:  $M_1 = 11$ ,  $M_2 = 13$ ,  $SD_1 = 4$ ,  $SD_2 = 4$ ,  $n_1 = 142$ ,  $n_2 = 128$

## ▶ Cohen's $d$ Values

$$d = \frac{M_1 - M_2}{\sqrt{\frac{(n_1 - 1)SD_1^2 + (n_2 - 1)SD_2^2}{n_1 + n_2 - 2}}}$$

$$d_1 = \frac{M_1 - M_2}{\sqrt{\frac{(n_1 - 1)SD_1^2 + (n_2 - 1)SD_2^2}{n_1 + n_2 - 2}}} = \frac{12 - 14}{\sqrt{\frac{(22 - 1)3^2 + (32 - 1)3^2}{22 + 32 - 2}}} = -.67$$

$$d_2 = \frac{M_1 - M_2}{\sqrt{\frac{(n_1 - 1)SD_1^2 + (n_2 - 1)SD_2^2}{n_1 + n_2 - 2}}} = \frac{14 - 16}{\sqrt{\frac{(25 - 1)2^2 + (52 - 1)2^2}{25 + 52 - 2}}} = -1.00$$

$$d_3 = \frac{M_1 - M_2}{\sqrt{\frac{(n_1 - 1)SD_1^2 + (n_2 - 1)SD_2^2}{n_1 + n_2 - 2}}} = \frac{11 - 13}{\sqrt{\frac{(142 - 1)4^2 + (128 - 1)4^2}{142 + 128 - 2}}} = -.50$$

# Fixed Effects Meta-Analysis Example

- ▶ Study 1:  $M_1 = 12$ ,  $M_2 = 14$ ,  $SD_1 = 3$ ,  $SD_2 = 3$ ,  $n_1 = 22$ ,  $n_2 = 32$
- ▶ Study 2:  $M_1 = 14$ ,  $M_2 = 16$ ,  $SD_1 = 2$ ,  $SD_2 = 2$ ,  $n_1 = 25$ ,  $n_2 = 52$
- ▶ Study 3:  $M_1 = 11$ ,  $M_2 = 13$ ,  $SD_1 = 4$ ,  $SD_2 = 4$ ,  $n_1 = 142$ ,  $n_2 = 128$

## ▶ Variances of the $d$ values

$$\circ s^2(d) = \frac{n_1+n_2}{n_1n_2} + \frac{d^2}{2(n_1+n_2-2)}$$

$$\circ s^2(d_1) = \frac{n_1+n_2}{n_1n_2} + \frac{d^2}{2(n_1+n_2-2)} = \frac{22+32}{(22)(32)} + \frac{-.67^2}{2(22+32-2)} = .085$$

$$\circ s^2(d_2) = \frac{n_1+n_2}{n_1n_2} + \frac{d^2}{2(n_1+n_2-2)} = \frac{25+52}{(25)(52)} + \frac{-1.00^2}{2(25+52-2)} = .073$$

$$\circ s^2(d_3) = \frac{n_1+n_2}{n_1n_2} + \frac{d^2}{2(n_1+n_2-2)} = \frac{142+128}{(142)(128)} + \frac{-.50^2}{2(142+128-2)} = .016$$

Notice that the study with the smallest variance for its associated effect size has the largest N

# Fixed Effects Meta-Analysis Example

- ▶ Study 1:  $M_1 = 12$ ,  $M_2 = 14$ ,  $SD_1 = 3$ ,  $SD_2 = 3$ ,  $n_1 = 22$ ,  $n_2 = 32$
- ▶ Study 2:  $M_1 = 14$ ,  $M_2 = 16$ ,  $SD_1 = 2$ ,  $SD_2 = 2$ ,  $n_1 = 25$ ,  $n_2 = 52$
- ▶ Study 3:  $M_1 = 11$ ,  $M_2 = 13$ ,  $SD_1 = 4$ ,  $SD_2 = 4$ ,  $n_1 = 142$ ,  $n_2 = 128$

## ▶ Weights

- $w = \frac{1}{s^2(d)}$
- $w_1 = \frac{1}{s^2(d)} = \frac{1}{.085} = 11.73$
- $w_2 = \frac{1}{s^2(d)} = \frac{1}{.073} = 13.78$
- $w_3 = \frac{1}{s^2(d)} = \frac{1}{.016} = 63.34$

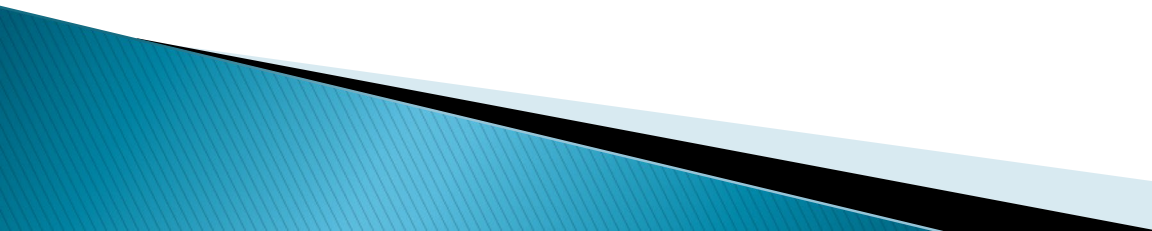
Notice that the study with the largest weight for its associated effect size has the smallest variance/largest N



# Fixed Effects Meta-Analysis Example

- ▶  $\hat{Y}_F = \frac{\sum_{i=1}^S w_i \hat{Y}_i}{\sum_{i=1}^S w_i} = \frac{(11.73)(-.67) + (13.78)(-1.00) + (63.34)(-.5)}{11.73 + 13.78 + 63.34} = -.60$
- ▶  $s^2(\hat{Y}_F) = \frac{1}{\sum_{i=1}^S w_i} = \frac{1}{11.73 + 13.78 + 63.34} = .011$
- ▶  $SE(\hat{Y}_F) = \sqrt{s^2(\hat{Y}_F)} = \sqrt{.011} = .10$
- ▶  $95\%CI(\hat{Y}_F) = \hat{Y}_F \pm (1.96)SE(\hat{Y}_F) =$   
 $\{(-.60 - 1.96 * .10), (-.60 + 1.96 * .10)\} = \{-.80, -.40\}$

# Fixed Effects Meta-Analysis Example

- ▶ Note: You are not going to be doing any of these “hand calculations” yourself
    - All of the calculations will be done using software
  - ▶ The example was simply to provide you with an idea of how the process of meta-analysis is carried out “behind-the-scenes”
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# Random Effects Meta-Analysis

- ▶ For a set of  $S$  effect size measures ( $\gamma$ )

- $\hat{\gamma}_R = \frac{\sum_{i=1}^S w_i^* \hat{\gamma}_i}{\sum_{i=1}^S w_i^*}$

- $w_i^* = \frac{1}{s^2(\hat{\gamma}_i) + \tau^2}$

- $\tau^2 = \frac{Q - (S-1)}{\sum_{i=1}^S w_i - \frac{\sum_{i=1}^S w_i^2}{\sum_{i=1}^S w_i}}$  for  $Q > S-1$

- $Q = \sum_{i=1}^S w_i (\hat{\gamma}_i - \hat{\gamma}_F)^2$

- $s^2(\hat{\gamma}_R) = \frac{1}{\sum_{i=1}^S w_i^*}$

Weights are more similar across studies given the addition of the constant  $\tau^2$

# Heterogeneity of Effect Sizes

- ▶ A simple goodness-of-fit test can be used to test for excessive heterogeneity
  - $Q \sim \chi^2_{df=S-1}$ 
    - We computed  $Q$  on the previous slide
    - We reject the null that there is no population heterogeneity if  $Q \geq \chi^2_{\alpha, df=S-1}$
- ▶ The problem with this approach is that the test has low-power when  $S$  is small

# Heterogeneity of Effect Sizes

- ▶ A better approach to quantifying heterogeneity is to use an effect size measure

$$I^2 = \frac{Q - S + 1}{Q}$$

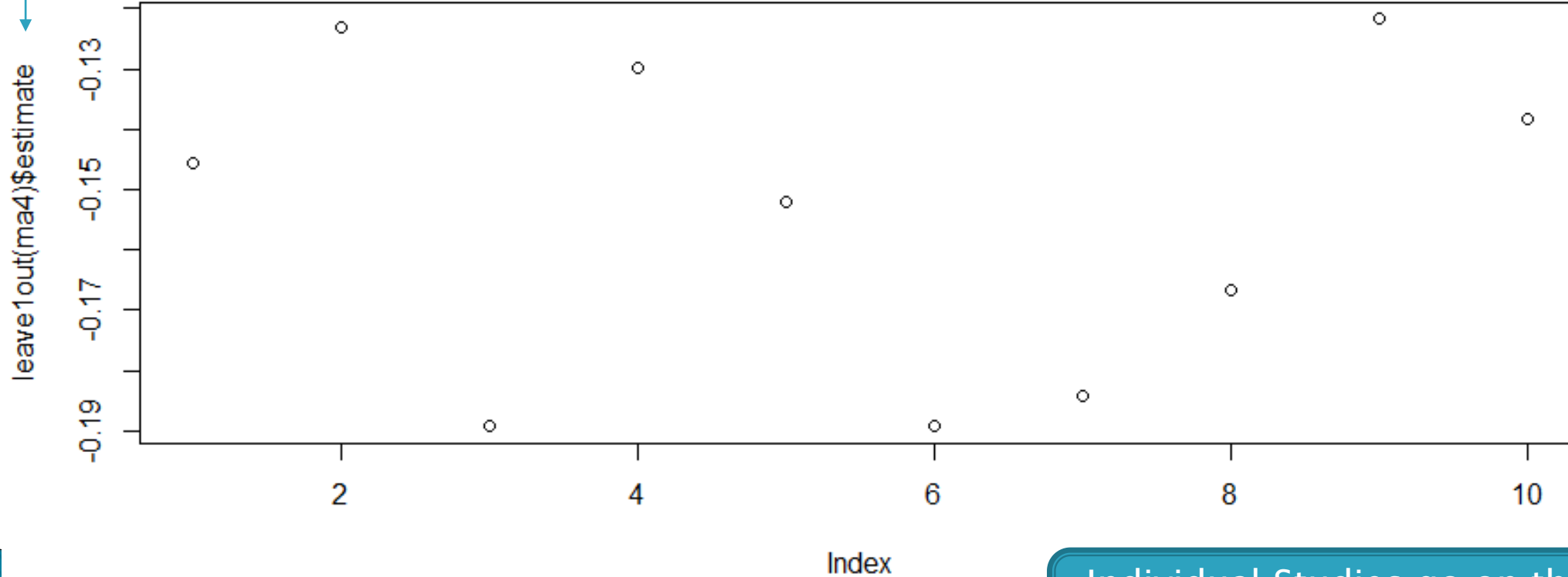
- ▶ Tells us what proportion of the observed variance in effect sizes is due to true differences in effect sizes, rather than sampling error
- ▶  $I^2$  ranges from 0 to 1, with larger values indicating more heterogeneity

# Outliers (Influential Cases)

- ▶ There are different ways of assessing the effect of outliers, but the main issue relates to what effect each study has on the combined effect size
- ▶ The easiest way to observe the effect of outliers is through “leave-one-out” analyses
  - Cook’s distance
    - A measure of the influence of individual cases on the combined effect
      - Popular cutoff is  $4/(\text{Number of Studies})$
  - Plot the combined effect, as a function of which study is left out

# Plot of “Leave-One-Out” Analyses

Combined  
Effect Size



Individual Studies go on the  
X-axis, here studies 1–10

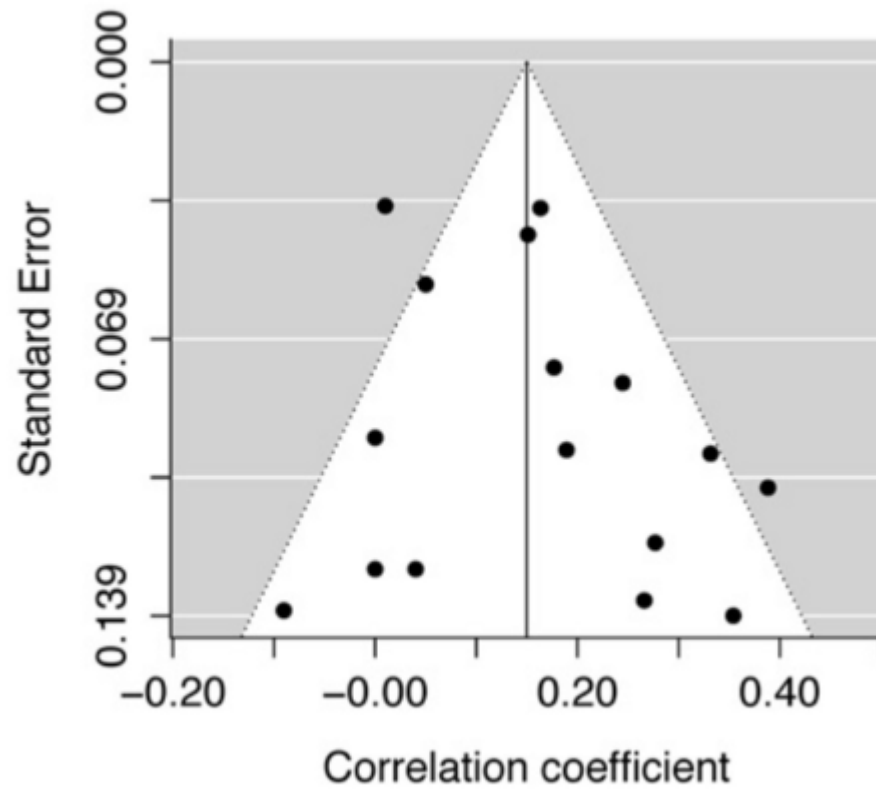
# Publication Bias

- ▶ When there is publication bias (e.g., studies with statistically significant effects getting published), studies with small sample sizes tend to have large effects
  - A large effect size is needed for an effect based on a small N to be statistically significant
- ▶ If we plot effect sizes against sample size/standard error, publication bias would show up in terms of “asymmetry”
  - Small N studies would all tend to have large effects
- ▶ This plot is called a *funnel plot*

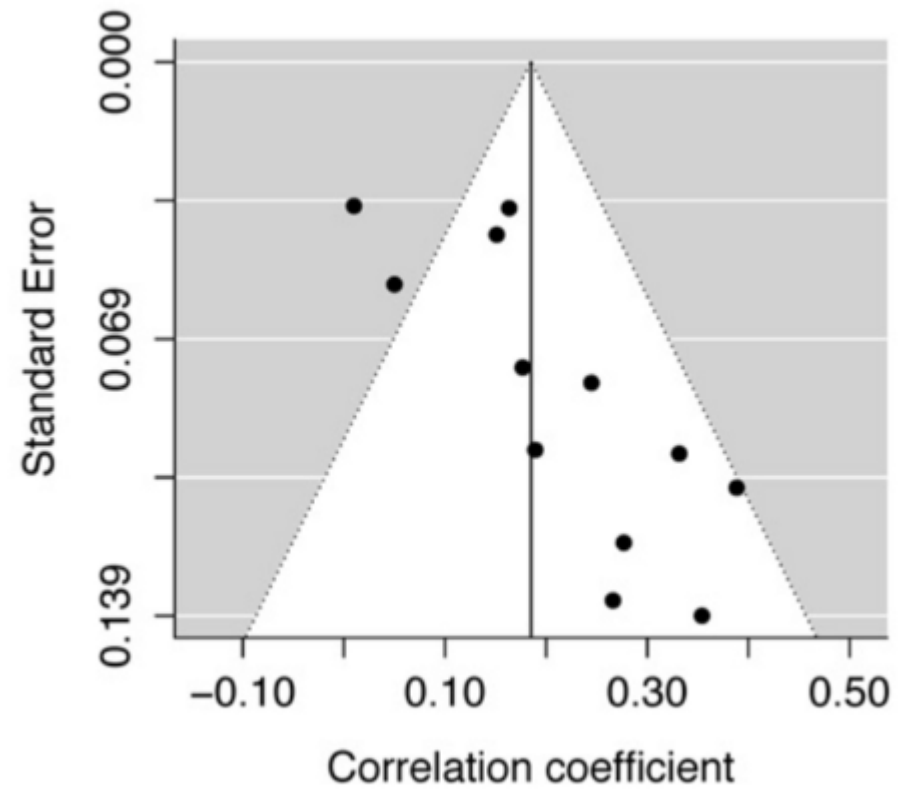


# Funnel Plot: Which Shows Evidence of Publication Bias?

**A**



**B**



# Conclusion: Why Meta-Analysis?

- ▶ Focuses on effect sizes, not statistical significance
  - ▶ Combines multiple studies for a more precise estimate of the effect size
  - ▶ Provides a rationale for small-N research
    - I.e., the results will be combined with other studies for a more precise estimate of the effect size
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