PSYC30013 Research Methods for Human Inquiry Week 11 (Day 1): Meta-Analysis

MANY META-ANALYSIS STUDIES INCLUDE THE PHRASE "WE SEARCHED MEDLINE, EMBASE, AND COCHRANE FOR STUDIES..."

THIS HAS LED TO META-META-ANALYSES COMPARING META-ANALYSIS METHODS.

69 M SAMPSON (2003), PL ROYLE (2005)
E LEE (2011), AR LEMESHOU (2005)

WE PERFORMED A META-META-META-ANALYSIS
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LIFE GOAL #28: GET A PAPER REJECTED
WITH THE COMMENT "TOO META"

A typical research publication

The standard research publication in a psychology journal usually contains the following features:

- 1. Theoretical basis of the research.
- 2. *Narrative review* of the relevant literature.
- 3. Rationale for undertaking the research.
- 4. Set of research questions (or hypotheses).
- 5. Description of the participant recruitment methods.
- 6. Explanation of the research design and measurement methods.
- 7. Justification of the statistical analysis to be used.
- 8. Summary of the findings.
- 9. Discussion of findings in relation to theory, previous literature, the rationale for the research, and limitations.

What's a narrative review?

Borenstein et al. (2021) explain:

"An expert in a given field would read the studies that addressed a question, summarize the findings, and then arrive at a conclusion – for example, that the treatment in question was, or was not, effective."

Key features:

- Expert subjectively decides which studies should be included in the review, and how they should be weighted
- Strategy of search for studies generally not disclosed
- Focus on qualitative description of prior results, rather than on quantification.
- Expert subjectively researches a conclusion about the overall trend of prior results

Problems with narrative reviews

Shadish, Cook, & Campbell (2002) list four problems with narrative reviews:

- 1. The greater the number of studies under review, the greater the difficulty in forming a coherent narrative.
- 2. They encourage a comparison between the number of statistically significant vs non-significant studies ('vote counting').
- 3. Narrative reviews tend to describe results in an imprecise way
- 4. Narrative reviews struggle to account for more studies involving complex relationships between variables.

What is a systematic review?

Cochrane Handbook (2023): "A systematic review attempts to collate all the empirical evidence that fits pre-specified eligibility criteria in order to answer a specific research question"

Systematic reviews adopt transparent and predefined

- Inclusion/exclusion criteria
- Search strategies
- Coding of studies

Most systematic reviews include *statistical* synthesis of past research (i.e. they include a meta-analysis)

What is meta-analysis?

Glass (1976): "[T]he statistical analysis of a large collection of analysis results from individual studies for the purpose of integrating the findings."

Cochrane Handbook (2023): "Meta-analysis is the statistical combination of results from two or more separate studies."

Unlike systematic review, meta-analysis is inherently statistical. Typically, the key statistic will be a summary estimate of the true effect of some variable.

- Example: a meta-analysis combines 50 studies on drug X, and finds that:
 - Heart attack patients randomly assigned to take drug X live on average 70 days longer than patients randomly assigned to take a placebo. 95% Confidence Interval: 60-80 days.

Meta-analysis and systematic review

They often go together. Lots of papers have titles like:

 Van Nostrand, M. (2024). Uromycitisis poisoning: a systematic review and meta-analysis.

To do a meta-analysis, you pretty much need to do a systematic review first

Not true in reverse - people often do systematic reviews and don't go on to a meta-analysis

Meta-analysis was initially controversial

In Part III I briefly examine the issue of the relevance of statistical inference, of whatever kind, to research in social sciences. I devote a section to a new bête noire, meta-analysis. It should be stifled at birth.

AMERICAN PSYCHOLOGIST • MAY 1978

H. J. EYSENCK
Institute of Psychiatry
London, England

An Exercise in Mega-Silliness

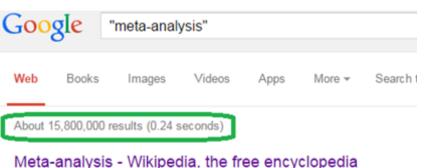
The article by Smith and Glass (September 1977) begins promisingly by referring to my "tendentious diatribes" (p. 752) on the outcome problem in psychotherapy, inviting



Meta-analysis/Shmeta-analysis

Samuel Shapiro

Meta-analysis is alive and well



In statistics, meta-analysis comprises statistical methods for contrasting and

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en.wikipedia.org/wiki/Meta-analysis - Wikipedia -

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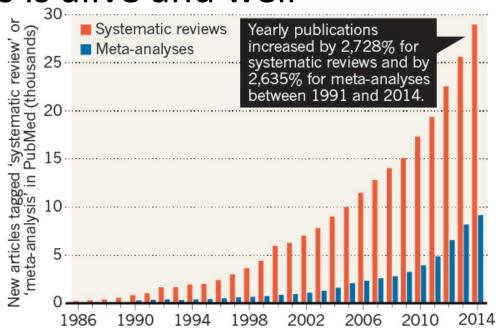
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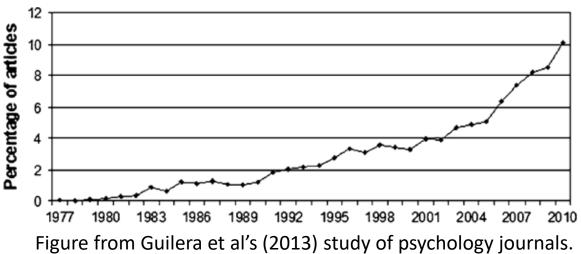
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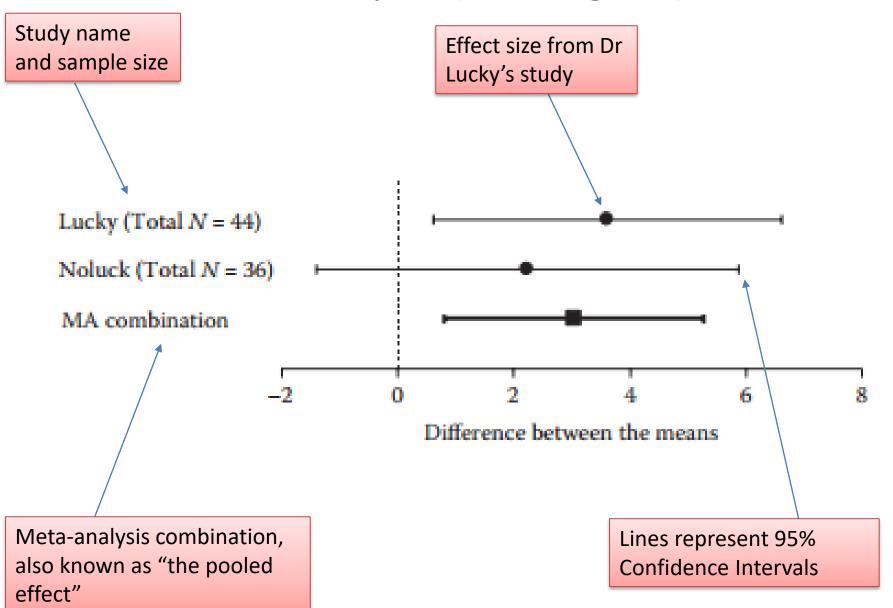
LIFE GOAL #28: GET A PAPER REJECTED

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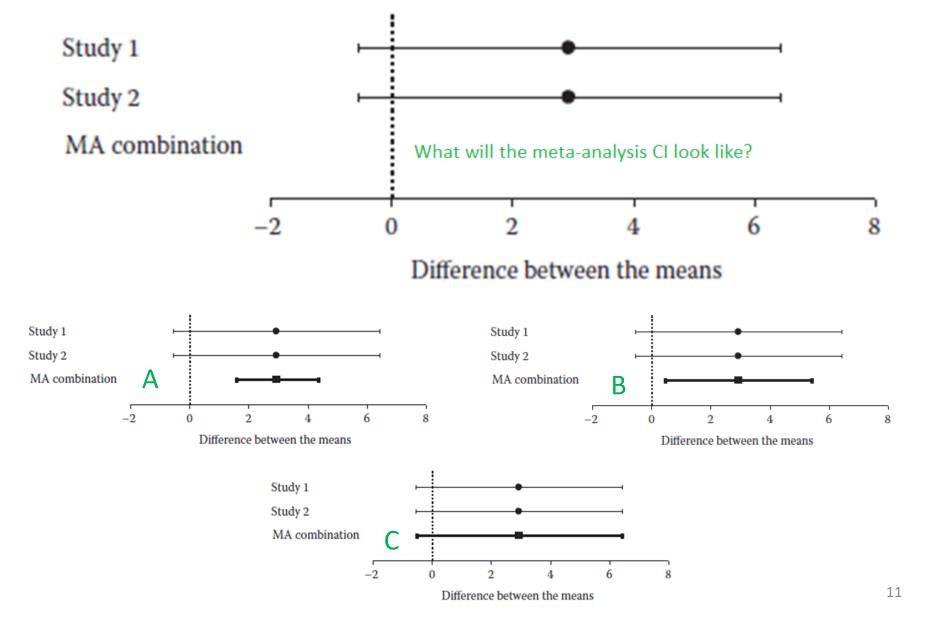




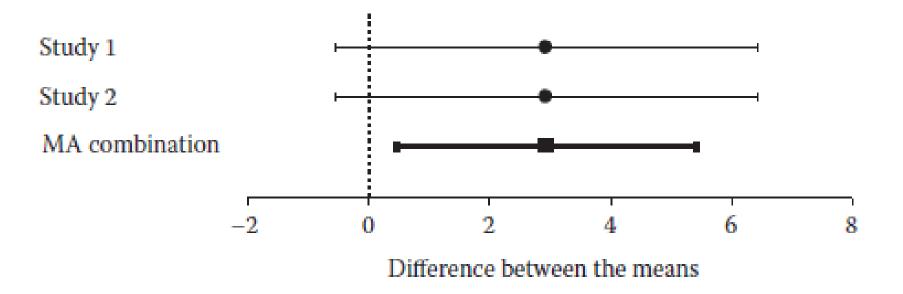
Forest plot (blobbogram)



Example: combining two studies in which p = .10



Example: combining two studies in which p = .10



The meta-analysis confidence interval is about 30% shorter than those of the two original studies

• Exact reduction in length depends on the sample sizes of the original studies, and on the similarity of the confidence intervals in the original studies.

Risk Ratio (also known as "Relative Risk")

	Outcome	No outcome	N
Exposed	Α	В	$n_1 = A + B$
Unexposed	С	D	$n_2 = C + D$

$$Risk \ Ratio = \frac{\binom{A}{n_1}}{\binom{C}{n_2}}$$

Risk ratios tell us about the strength of association between an exposure and an outcome. To get the risk ratio, take the risk (i.e. probability) of the outcome in the exposed group divided by the risk of the outcome in the unexposed group.

Risk Ratio (also known as "Relative Risk")

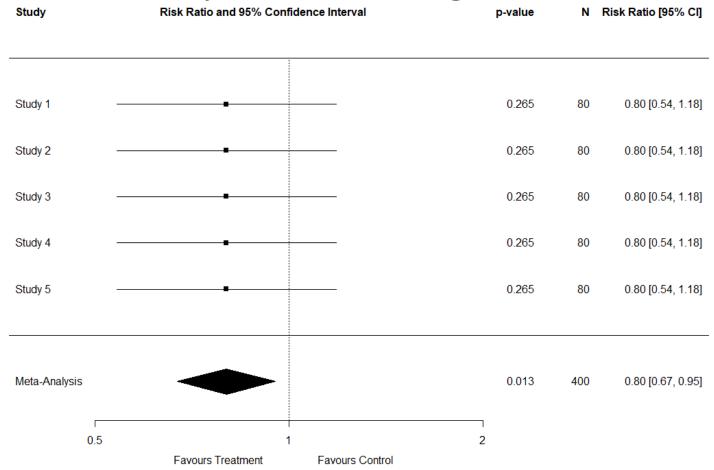
	Infected	Not infected	N
Treatment	A = 20	B = 20	n ₁ = 40
Control	C = 25	D = 15	$n_2 = 40$

Risk Ratio =
$$\frac{\binom{A}{n_1}}{\binom{C}{n_2}} = \frac{\binom{20}{40}}{\binom{25}{40}} = 0.8$$

The risk of infection was 62.5% (25/40) in the control group The risk of infection was 50.0% (20/40) in the treatment group

People taking the treatment had 0.8 times the risk of infection of those in the control group.

Why vote counting is bad



According to vote counting: "All five previous studies failed to achieve a statistically significant result."

According to meta-analysis: "The summary p-value is statistically significant, p = .013. The estimate of the effect size is..."

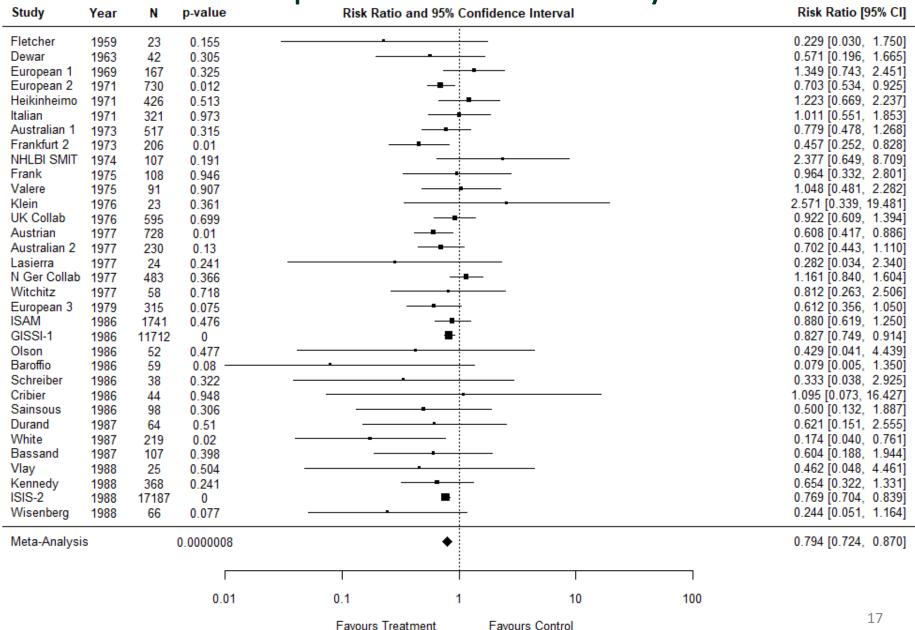
Study	Year	# of patients	p-value < .05?
Fletcher	1959	23	NO
Dewar	1963	42	NO
European 1	1969	167	NO
European 2	1971	730	YES
Heikinheimo	1971	426	NO
Italian	1971	321	NO
Australian 1	1973	517	NO
Frankfurt 2	1973	206	YES
NHLBI SMIT	1974	107	NO
Frank	1975	108	NO
Valere	1975	91	NO
Klein	1976	23	NO
UK Collab	1976	595	NO
Austrian	1977		YES
Australian 2	1977	230	NO
Lasierra	1977	24	NO
N Ger Collab	1977	483	NO
Witchitz	1977	58	NO
European 3	1979	315	NO
ISAM	1986	1,741	NO
GISSI-1	1986	11,712	YES
Olson	1986	52	NO
Baroffio	1986	59	NO
Schreiber	1986	38	NO
Criber	1986	44	NO
Sainsous	1986	98	NO
Durand	1987		NO
White	1987	219	YES
Bassand	1987		NO
Vlay	1988	25	NO
Kennedy	1988	368	NO
ISIS-2	1988	17,187	YES
Wisenberg	1988	66	NO
TOTAL		36,974	6 vs 27

Streptokinase studies

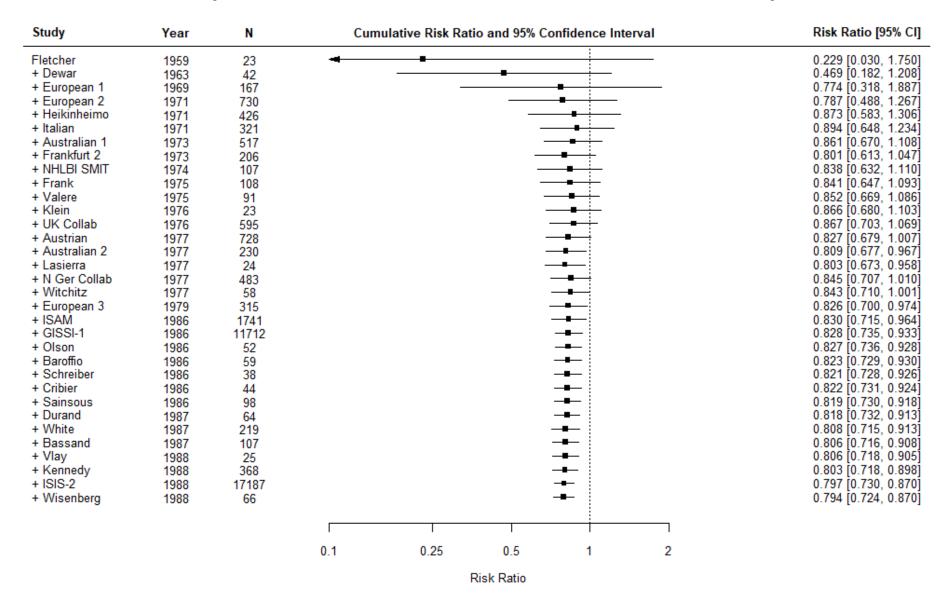
- Streptokinase is a medicine that dissolves blood clots
- Particularly useful after myocardial infarction (heart attack) has occurred
- Extremely lengthy period of clinical trials from late 1950s to late 1980s.



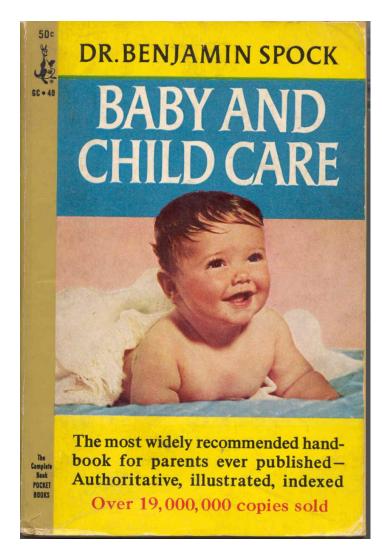
Streptokinase meta-analysis



Streptokinase cumulative meta-analysis



Meta-analysis and Sudden Infant Death Syndrome



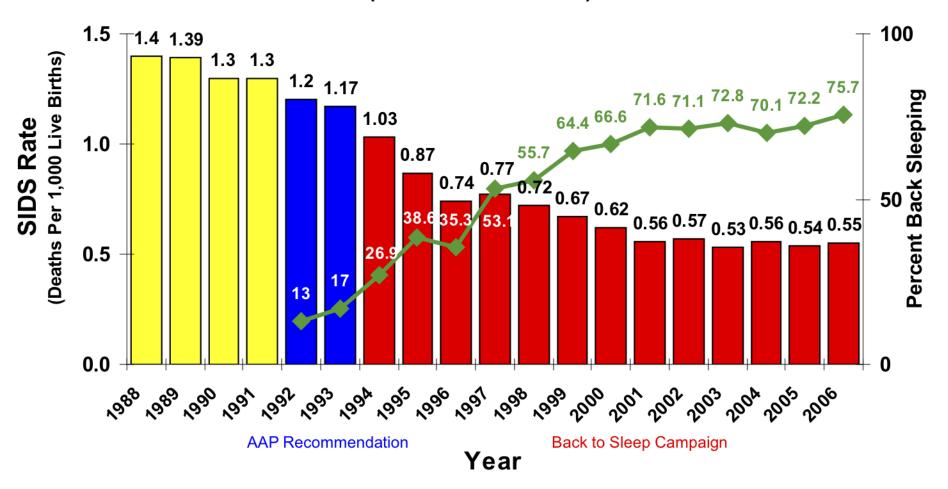
This well-known book contained advice now believed to be incorrect

- SIDS refers to the sudden unexplained death of a child < 1 year old
- 100,000+ deaths from 1950s-1990s
- Gilbert et al. (2005) write that
 - "Advice to put infants to sleep on the front for nearly half a century was contrary to evidence available from 1970 that this was likely to be harmful. Systematic review of preventable risk factors for SIDS from 1970 would have led to earlier recognition of the risks of sleeping on the front and might have prevented over 10,000 infant deaths in the UK and at least 50,000 in the Europe, the USA and Australasia."

Note: nothing in this lecture should be taken as medical advice. If any of the lecture examples become relevant to you in real life, seek advice from health professionals with relevant expertise

SIDS Rate and Back Sleeping

(1988 - 2006)



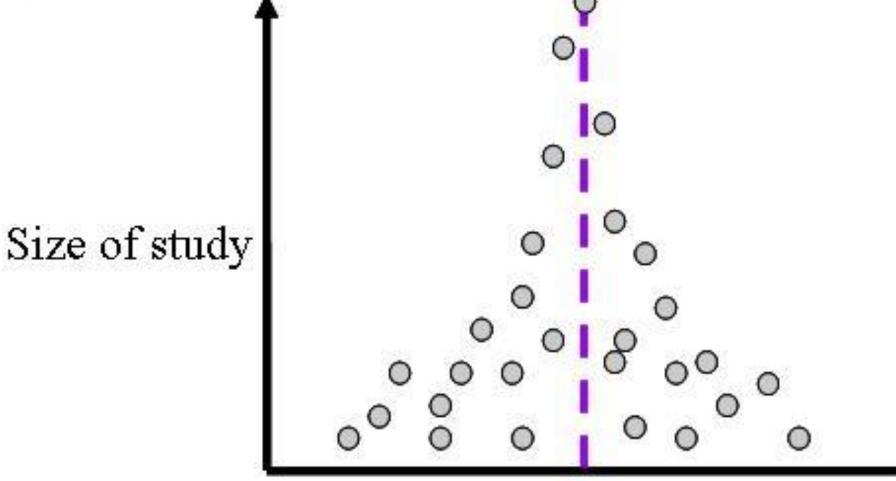
SIDS Rate Source: CDC, National Center for Health Statistics, Sleep Position Data: NICHD, National Infant Sleep Position Study.

- 1. The file drawer problem invalidates meta-analysis
- 2. One number cannot summarize a research field
- 3. Mixing apples and oranges
- 4. Garbage in, garbage out
- 5. Meta-analyses are done poorly
- 6. Meta-analyses are prone to conflicts of interest



Funnel plot with no file drawer problem



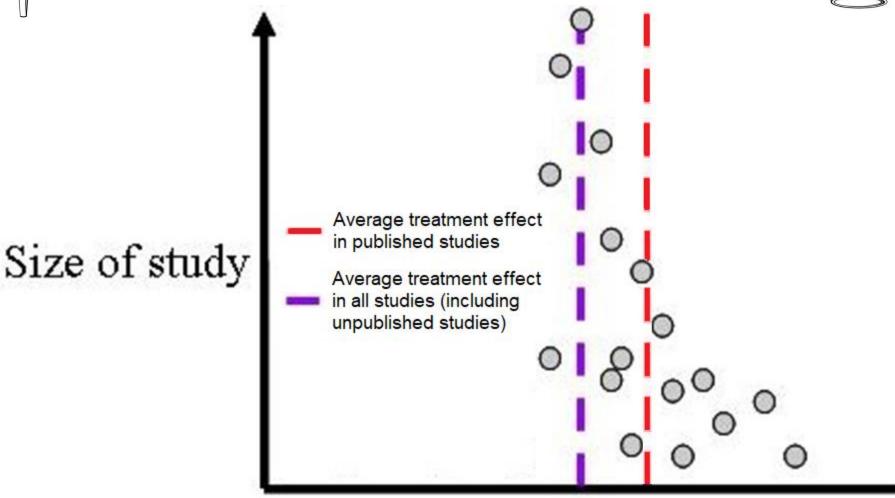


Treatment effect



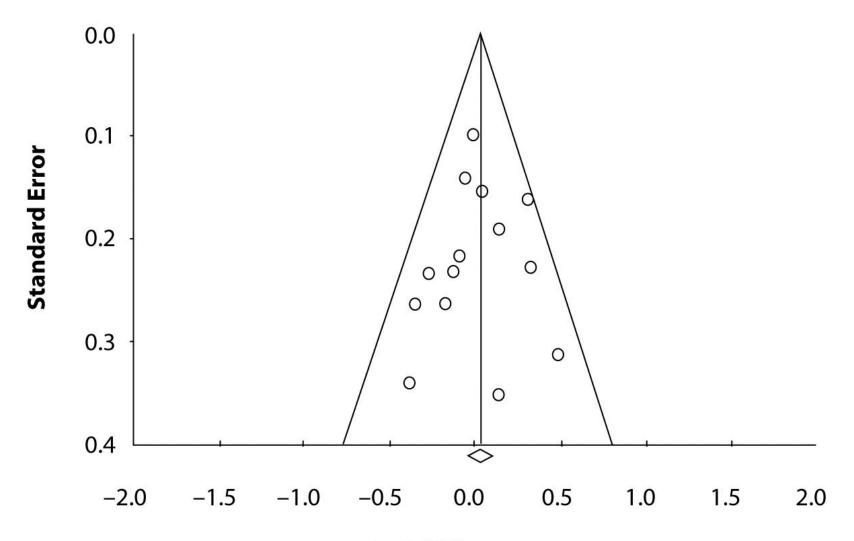
Funnel plot with file drawer problem



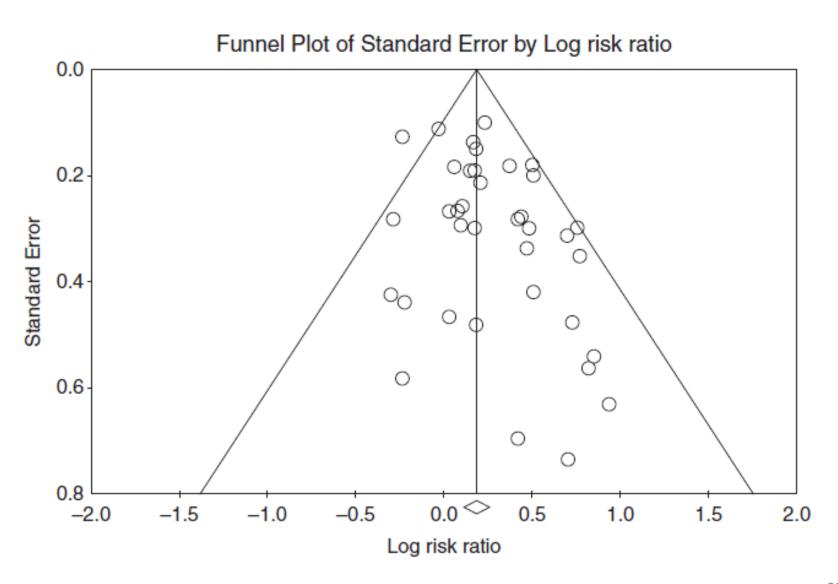


Treatment effect

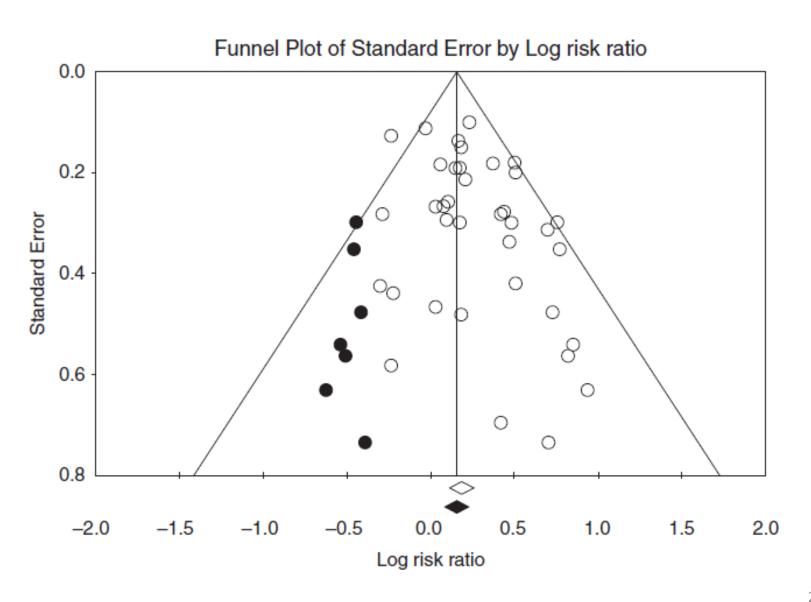
Funnel Plot: std error and std mean differences Funnel Plot of Standard Error by Std diff in means



Funnel Plot: std error and log risk ratio



Studies imputed with the 'trim and fill' method



Criticism 1: The file drawer problem invalidates meta-analysis.

- "Trim and fill" method can help
- Problem also arises in narrative review
- Unpublished studies can be sought out
- Preregistration of studies can also help

Criticism 2: One number cannot summarize a research field.

Response: Meta-analysis seeks synthesis and not (necessarily) a summary

Criticism 3: Meta-analysis may ignore important differences across studies, and hence be 'mixing apples and oranges'.

- Meta-analysis allows us to address the differences formally
- Mixing apples and oranges is good if you want to make a fruit salad

Criticism 4: Garbage in, garbage out. By including many studies meta-analyses include low quality studies.

- The problem is addressed by adopting appropriate inclusion criteria.
- Meta-analysis allows a better understanding of the properties of (allegedly) garbage studies

Criticism 5: Meta-analyses are done poorly.

- The problem is addressed by improving standards of meta-analysis
- No evidence narrative reviews are done better
- The APA Publication Manual publishes guidelines to help improve the quality of meta-analysis reporting, as does the Cochrane handbook.

Criticism 6: Meta-analyses are prone to conflicts of interest

Case study of anti-depressants:

- 185 meta-analyses of antidepressants published during 2007-2014
- 54 (29%) had authors who were employees of the assessed drug's manufacturer
- 147 (79%) had some other industry-related conflict of interest

Some suggestions from Ioaniddes (2016):

- More transparency re: author conflicts of interest
- More widespread sharing of data
- Better enforcement of preregistration

Meta-analysis and the APA Publication Manual

Recommends providing detailed statistical information, and the complete set of raw data where possible.

- "Mention all relevant results ... be sure to include small effect sizes (or statistically nonsignificant findings)" (p. 32).
- "Your work will more easily become a part of the cumulative knowledge of the field if you include enough statistical information to allow its inclusion in future meta-analyses" (p. 34).

Sets out Meta-Analysis Reporting Standards

Provides a sample paper to illustrate a meta-analysis write-up.

References

Optional reading, not required for this course but potentially useful for your future career:

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