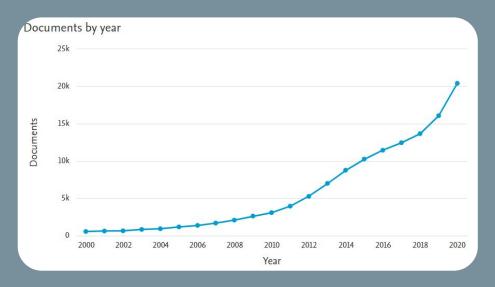
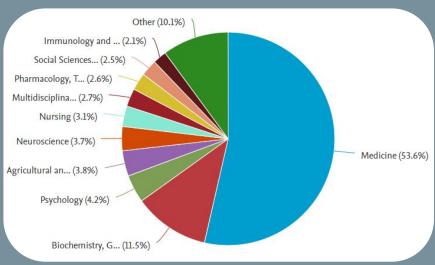
# Intro to meta-analysis using R

Tengku Muhd Hanis Mokhtar PhD student, USM August 7, 2021

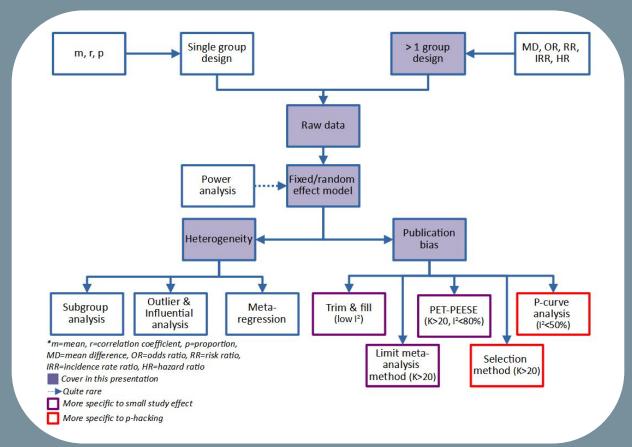
### **Background**

- Meta-analysis:
  - Statistical methods used to combine individual results into pooled result
- From Scopus database (20-07-2021): 144, 904 documents





#### **General framework**



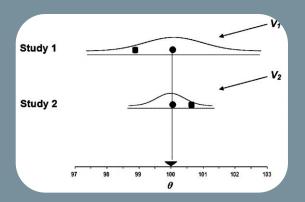
## **Basic jargons**

- Fixed vs random effect model
- Between-study heterogeneity
- Publication bias
- Forest plot
- Funnel plot



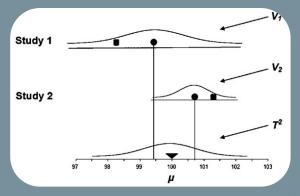
#### Fixed effect:

- One true effect size
- Estimate one true effect size



#### Random effect:

- True effects varies (ie; distribution of true effect sizes)
- Estimates mean of the distribution of true effects



(Borenstein et al., 2010)

Heterogeneity (almost always refer to between study heterogeneity):

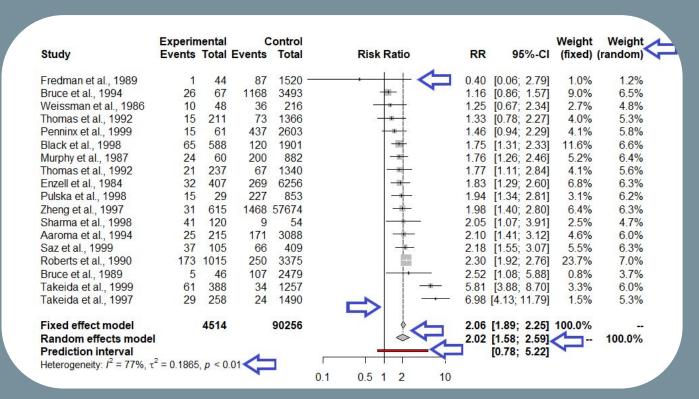
- Variation in study outcomes between studies (statistical heterogeneity)
- Measurement: Q-statistics, T<sup>2</sup>, I<sup>2</sup>, H<sup>2</sup>
- Other types of heterogeneity refer to Rucker at al., 2008

#### Publication bias:

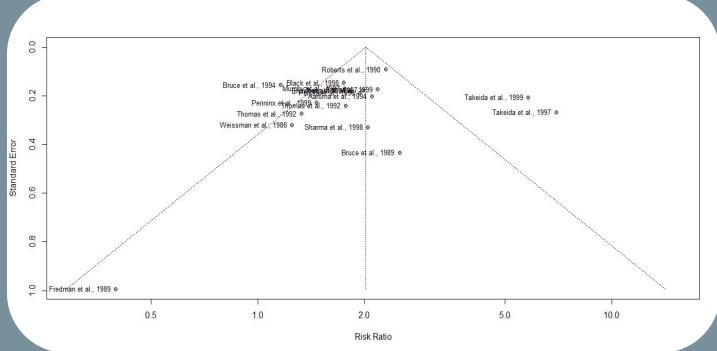
- Studies being published depends on the its result
- Consequences:
  - Overestimate the effect size
  - Overlook negative effect size

- Publication bias tested using:
  - Visual: Funnel plot
  - Statistical (min k=10):
    - Classical: Begg, Egger (default), Thompson
    - Binary outcome: Peters, Harbord (default for OR), Schwarzer, Deeks, etc.
    - SMD (for Hedges' g): Pustejovsky

Forest plot



Funnel plot



#### **Advanced** method

- Variation of visualization:
  - Forest plot equivalent: Drapery plot
  - Variant of funnel plot: Contour-enhanced funnel plot
- Meta-regression
- Subgroup analysis
- Outlier and influential diagnostic (rule of thumb;  $I^2 > 50\%$ )
- Publication bias related method
- etc

#### Type of meta-analysis

- 1. "General" meta-analysis (Intervention/observational study)
  - Single group design: Pool mean, correlation coefficient, prevalence/proportion
  - >1 group design: Pool mean difference, OR, RR, IRR, HR
- 2. "Multilevel" meta-analysis
  - There is 3rd level
- 3. Network meta-analysis
  - Compare several treatment effect directly and indirectly
- 4. Dose response meta-analysis
  - Quantify level of exposure effect to response

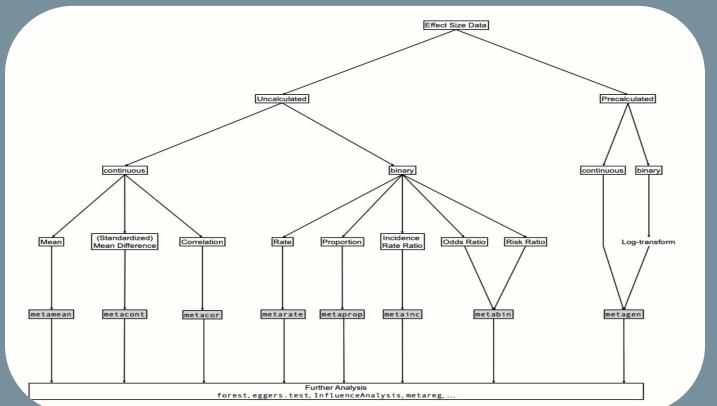
#### Type of meta-analysis (cont.)

- 5. Diagnostic test accuracy meta-analysis
  - Pool sensitivity, specificity, AUC
- 6. Multivariate/Anova/SEM meta-analysis
- 7. Bayesian approach
- 8. Genome meta-analysis

## Packages in R

Package Characteristics													
Package	Version	Title	Effect Size	Power	Missing Data	Dependent Effects		Random Effects	Moderator Analyses	Publication Bias	Sensitivity Analysis	Creates Plots	Primary Function
General meta-ana	llysis												
CAMAN	0.7	Finite mixture models and meta- analysis tools						$\checkmark$	$\checkmark$				bivariate
epiR	0.9-62	Tools for the analysis of epidemiological data	$\checkmark$		$\checkmark$		$\checkmark$	$\checkmark$					epi.dsl
gmeta	2.2-3	Meta-analysis via a unified framework under confidence distribution	$\checkmark$			$\checkmark$	$\checkmark$	$\checkmark$				$\checkmark$	gmeta
Mac	1.1	Meta-analysis with correlations	$\checkmark$		$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	mareg
Mad	0.8-2	Meta-analysis with mean differences	$\checkmark$		$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$		$\checkmark$	mareg
Meta	4.2-0	General package for meta- analysis	$\checkmark$		$\checkmark$		$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$	metacont
metacor	1.0-2	Meta-analysis of correlation coefficients	$\sqrt{}$				$\sqrt{}$	$\checkmark$					metacor.DSL
metafor	1.9-5	Meta-analysis package for R	V		V	<b>√</b>	V		$\checkmark$	$\checkmark$		V	rma
metaplus	0.7-1	Robust meta-analysis and meta- regression	V			$\sqrt{}$		$\checkmark$	$\checkmark$		$\checkmark$	$\checkmark$	metaplus
psychometric	2.2	Applied psychometric theory					$\checkmark$		$\checkmark$	$\checkmark$		$\checkmark$	MetaTable
rmeta	2.16	Meta-analysis	$\checkmark$				V	$\checkmark$	2.4%	V	$\checkmark$	V	meta.MH

## Main functions in meta packages



(Harrer et al., 2021)

#### References

- Borenstein, M., Hedges, L. V., Higgins, J. P. T. & Rothstein, H. R. A basic introduction to fixed-effect and random-effects models for meta-analysis. Res. Synth. Methods 1, 97–111 (2010).
- Harrer, M., Cuijpers, P., Furukawa, T.A., & Ebert, D.D. (2021). <u>Doing Meta-Analysis with R: A Hands-On Guide</u>. Boca Raton, FL and London: Chapman & Hall/CRC Press. ISBN 978-0-367-61007-4.
- Polanin, J. R., Hennessy, E. A. & Tanner-Smith, E. E. A Review of Meta-Analysis Packages in R. J. Educ. Behav. Stat. 42, 206–242 (2017).
- Rücker, G., Schwarzer, G., Carpenter, J. R. & Schumacher, M. Undue reliance on I2 in assessing heterogeneity may mislead. BMC Med. Res. Methodol. 8, 1–9 (2008).

# Hands-on in



# Thank you!



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