

#### AGENCY FOR HEALTHCARE RESEARCH AND QUALITY









### **North American Systematic Review Methods Virtual Research Day**



October 30<sup>th</sup>, 2020 11:30 – 3:30 pm ET / 8:30 – 12:30 pm PT





- Guidelines International Network (GIN) North America
  - <u>https://g-i-n.net/regional-communities/gin-na</u>
- Cochrane United States
  - https://us.cochrane.org/
- Cochrane Canada
  - <u>https://canada.cochrane.org/</u>
- AHRQ Evidence-based Practice Center program
  - <u>https://effectivehealthcare.ahrq.gov/</u>
- Scientific Resource Center
  - <u>https://effectivehealthcare.ahrq.gov/about/src</u>

#### **Non-randomized Studies**

A tool to assess Risk Of Bias In Non-randomised Studies - of Exposures (ROBINS-E)\*

- Julian Higgins (University of Bristol)

• Guideline development using systematic reviews supplemented with internal health system data: The development and application of a conceptual framework\*

- Jennifer Lin & Helen Wu (Kaiser Permanente, USA) GIN NA abstract
- Examining the effect of nutrition interventions to reduce hyperphosphatemia in chronic kidney disease: Is including non-randomized trials a waste of time?\*
  - Mary Rozga (Academy of Nutrition and Dietetics, USA)

\* Slides for this presentation are included in this packet



#### **Strength of Evidence and Bias**

 Meta-analysis as a simultaneous inference problem: a novel approach to assess replicability of evidence

- Orestis Panagiotou (Brown University, USA)
- Methodological review of items for assessing the risk of bias in network meta-analyses provides groundwork for the development of a new risk of bias tool for network metaanalysis\*
  - Carole Lunny (Cochrane Hypertension Group, Canada)
- Ignoring non-significant factors without data may bias the results of meta-analysis of prognostic studies
  - Li Wang (McMaster University, Canada)
- Dealing with retrieval bias for an evidence-informed individual patient data network meta-analysis (GIN abstract)\*
  - Areti A. Veroniki (St. Michael's Hospital, Canada) GIN NA abstract

\* Slides for this presentation are included in this packet



#### **Improving Use of Systematic Reviews**

- depressionscreening100.com/phq: A practice-based perspective to using the Patient Health Questionnaire-9 to screen for depression\*
  - Brooke Levis (McGill University, Canada)
- Using a distribution-based approach and systematic review methods to derive minimum clinically important differences
  - Jennifer Watt (University of Toronto, Canada)
- Progress toward a reporting guideline for overviews of reviews of healthcare interventions: Preferred Reporting Items for Overviews of Reviews (PRIOR)\*
  - Michelle Gates (University of Alberta, Canada)

<sup>\*</sup> Slides for this presentation are included in this packet



#### **Improving Efficiency of Systematic Reviews**

- Accelerating integration of emerging evidence into healthcare delivery: rapid reviews for learning health systems\*
  - Marcy Hager (Oregon Health & Science University, USA)
- Rapid review methods: a systematic scoping review\*
  - Candyce Hamel (Ottawa Hospital Research Institute, Canada)
- A new machine-learning powered tool to aid citation screening for evidence synthesis: PICOPortal\*
  - Eitan Agai (Fast Healthcare Interoperability Resources, Resources for Evidence-Based Medicine [EBMonFHIR], USA)

\* Slides for this presentation are included in this packet

Italicized titles indicate the presenting author selfidentified as an early career investigator



#### A tool to assess Risk Of Bias In Non-randomised Studies - of Exposures (ROBINS-E)

Julian Higgins on behalf of the ROBINS-E development team, led by Jonathan Sterne, Julian Higgins, Rebecca Morgan (*originally due to deliver this*), Kyla Taylor, Andrew Rooney, Holger Schünemann and Kristina Thayer

Morgan R, Taylor K, Higgins J, Rooney A, Thayer K, Schünemann H, Sterne J. A tool to assess Risk Of Bias In Non-randomized Studies – of Exposures (ROBINS-E). In: *Advances in Evidence Synthesis: special issue. Cochrane Database of Systematic Reviews* 2020; (9 Suppl 1): p.321. https://doi.org/10.1002/14651858.CD202001

no conflicts of interest to declare



## A modern family of risk-of-bias assessment tools in health research

	Randomized trials of interventions	Non- randomized/observational studies of interventions	Non- randomized/observational studies of exposures
First	Cochrane RoB	ROBINS-I	
generation	(2008)	(2016)	
Second	RoB 2	ROBINS-I V2	
generation	(2019)	(coming soon)	



#### A plethora of tools

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International Journal of Epidemiology 2007;**36**:666–676 doi:10.1093/ije/dym018

#### Tools for assessing quality and susceptibility to bias in observational studies in epidemiology: a systematic review and annotated bibliography

86 in 2007

Simon Sanderson,<sup>1</sup>\* Iain D Tatt<sup>2,4</sup> and Julian PT Higgins<sup>3</sup>





#### A modern family of risk-of-bias assessment tools in health research

#### **RESEARCH METHODS AND REPORTING** CON OPEN ACCESS ROBINS-I: a tool for assessing risk of bias in non-randomised studies of interventions Jonathan AC Sterne,<sup>1</sup> Miguel A Hernán,<sup>2</sup> Barnaby C Reeves,<sup>3</sup> Jelena Savović,<sup>1,4</sup> Nancy D Berkman,<sup>5</sup> Meera Viswanathan,<sup>6</sup> David Henry,<sup>7</sup> Douglas G Altman,<sup>8</sup> Mohammed T Ansari,<sup>9</sup> Isabelle Boutron,<sup>10</sup> James R Carpenter,<sup>11</sup> An-Wen Chan,<sup>12</sup> Rachel Churchill,<sup>13</sup> Jonathan J Deeks,<sup>14</sup> Asbjørn Hróbjartsson,<sup>15</sup> Jamie Kirkham,16 Peter Jüni,17 Yoon K Loke,18 Theresa D Pigott, 19 Craig R Ramsay,20 Deborah Regidor,21 Hannah R Rothstein,<sup>22</sup> Lakhbir Sandhu,<sup>23</sup> Pasqualina L Santaguida,<sup>24</sup> Holger J Schünemann,<sup>25</sup> Beverly Shea,<sup>26</sup> Ian Shrier,<sup>27</sup> Peter Tugwell,<sup>28</sup> Lucy Turner,<sup>29</sup> Jeffrey C Valentine,<sup>30</sup> Hugh Waddington,<sup>31</sup> Elizabeth Waters, 32 George A Wells, 33 Penny F Whiting, 34 Julian PT Higgins 35 For numbered affiliations see Non-randomised studies of the such as cohort studies and case-control studies in end of article. which intervention groups are allocated during the effects of interventions are critical to Correspondence to: J A C Sterne course of usual treatment decisions, and quasi-ranmany areas of healthcare evaluation, jonathan.steme@bristol.ac.uk domised studies in which the method of allocation but their results may be biased. It is falls short of full randomisation. Non-randomised the journal online. studies can provide evidence additional to that availtherefore important to understand Cite this as: BMJ 2016;355:i4919 able from randomised trials about long term outand appraise their strengths and http://dx.doi.org/10.1136/bmi.i4919 comes, rare events, adverse effects and populations. weaknesses. We developed ROBINS-I that are typical of real world practice.12 The availability of linked databases and compilations of electronic ("Risk Of Bias In Non-randomised health records has enabled NRSI to be conducted in Studies - of Interventions"), a new large representative population cohorts.3 For many tool for evaluating risk of bias in types of organisational or public health interventions, estimates of the comparative NRSI are the main source of evidence about the likely impact of the intervention because randomised trials effectiveness (harm or benefit) of are difficult or impossible to conduct on an area-wide interventions from studies that did basis. Therefore systematic reviews addressing the effects of health related interventions often include not use randomisation to allocate NRSL It is essential that methods are available to evalunits (individuals or clusters of uate these studies, so that clinical, policy, and individindividuals) to comparison groups. ual decisions are transparent and based on a full understanding of the strengths and weaknesses of the The tool will be particularly useful to evidence. those undertaking systematic Many tools to assess the methodological quality of

observational studies in the context of a systematic review have been proposed.45 The Newcastle-Ottawa6 and Downs-Black7 tools have been two of the most popular: both were on a shortlist of methodologically sound tools.5 but each includes items relating to external as well as internal validity and a lack of comprehensive manuals means that instructions may be interpreted differently by different users.5

In the past decade, major developments have been made in tools to assess study validity. A shift in focus from methodological quality to risk of bias has been accompanied by a move from checklists and numeric scores towards domain-based assessments in which different types of bias are considered in turn. Examples are the Cochrane Risk of Bias tool for randomised trials.8 the OUADAS 2 tool for diagnostic test accuracy studies.9 and the ROBIS tool for systematic reviews.10 However, there is no satisfactory domain-based assessment tool for NRSL<sup>4</sup>

In this paper we describe the development of ROBINS-I ("Risk Of Bias In Non-randomised Studies - of Interventions"), which is concerned with evaluating risk of bias in estimates of the

1

Nonomized/observational dies of interventions

**ROBINS-I V2** (coming soon)

**ROBINS-I** 

(2016)

Nonrandomized/observational studies of exposures

#### **ROBINS-E**

#### bristol.ac.uk

the**bmj** | BMJ 2016;355:i4919 | doi: 10.1136/bmj.i4919

reviews that include non-randomised studies. Non-randomised studies of the effects of interventions

(NRSI) are critical to many areas of healthcare evaluation. Designs of NRSI that can be used to evaluate the effects of interventions include observational studies

#### SUMMARY POINTS

- · Non-randomised studies of the effects of interventions are critical to many areas of healthcare evaluation but are subject to confounding and a range of other potential biases
- We developed, piloted, and refined a new tool, ROBINS-I, to assess "Risk Of Bias" In Non-randomised Studies - of Interventions"
- The tool views each study as an attempt to emulate (mimic) a hypothetical. pragmatic randomised trial, and covers seven distinct domains through which bias might be introduced
- · We use "signalling questions" to help users of ROBINS-I to judge risk of bias within each domain
- The judgements within each domain carry forward to an overall risk of bias judgement across bias domains for the outcome being assessed

Additional material is published online only. To view please visit

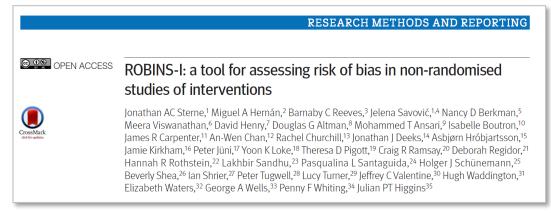
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- Joint initiative between our team in Bristol (UK), McMaster University (Canada), National Toxicology Program (NIH, USA), Environmental Protection Agency (USA) and others
- Fully drafted version piloted in Bristol in October 2019
- Refinements still being made
- Similar to ROBINS-I, but more attention to
  - defining the causal effect of interest
  - exposure measurement





Elie Akl, Carla Ancona, Mohammed Ansari, Bruce Armstrong, Whitney Arroyave, Tom Bateson, Nancy Berkman, Lisa Bero, Aaron Blair, Abee Boyles, Bert Brunekreef, Paul Demers, Tanja Farmer, Francesco Forastiere, Davina Ghersi, Barbara Glenn, Ali Goldstone, Gordon Guyatt, David Henry, Miguel Hernan, Julian Higgins, Ellen Kirrane, Judy LaKind, Juleen Lam, Tom Luben, Ruth Lunn, Alexandra McAleenan, Luke McGuinness, Daniele Mandrioli, Suril Mehta, Joerg Meerpohl, Rebecca Morgan, Rebecca Nachman, Annette O'Connor, Julie Obbagy, Neil Pearce, Beth Radke, Andrew Rooney, Kenneth Rothman, Jelena Savović, Mary Schubauer-Berigan, Holger Schünemann, Pam Schwingl, Beverly Shea, Kyle Steenland, Jonathan Sterne, Patricia Stewart, Kurt Straif, Kyla Taylor, Kris Thayer, Jos Verbeek, Roel Vermeulen, Meera Viswanathan, Shelia Zahm



#### Intervention vs exposure A continuum

- Interventions
  - by a health professional
  - legislation
- Personal choices
  - type of toothbrush
  - taking a vitamin supplement
  - dietary intake
  - lifestyle, e.g. smoking, exercise
- Exposures
  - occupational
  - environmental
- Traits
  - socioeconomic status
  - biomarkers
  - genetic

#### Intended





## Establishing the causal effect being evaluated

- For observational studies we need to define the causal effect estimated by the result under consideration
  - Convenient to use counterfactuals
  - It may help to define a **target experiment** 
    - In the hypothetical target experiment, exposure would be assigned in a planned manner, rather than being observed. An unlimited number of exposure plans can be assigned. The target experiment need not be feasible or ethical.
  - Essential for assessing risk of bias because it defines the result that would be seen (other than due to sampling variation) in the absence of bias



### Establishing the causal effect being evaluated



### Need not be feasible or ethical



#### **ROBINS-E domains**

Risk of bias due to confounding

Risk of bias in measurement of exposure

Risk of bias in selection of participants into the study (or into the analysis)

Risk of bias due to post-exposure interventions

Risk of bias due to missing data

Risk of bias in measurement of outcomes

Risk of bias in analysis and selection of the reported result



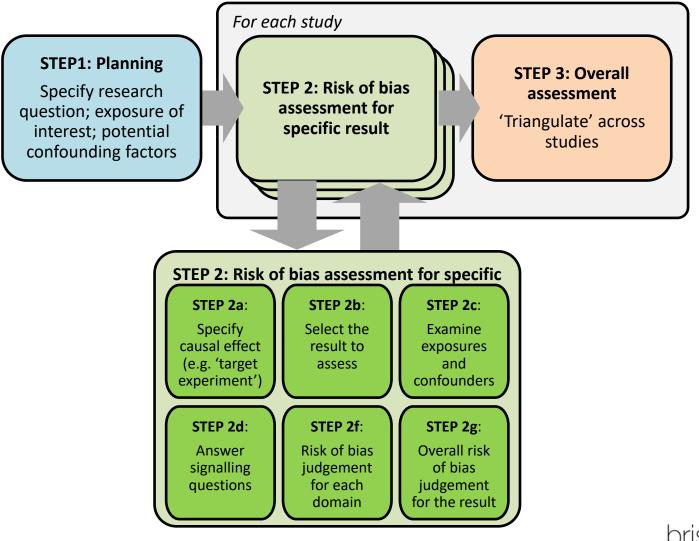


#### **ROBINS-E risk-of-bias judgement**

Judgement	Interpretation
Low risk of bias	there is little or no concern about bias with regard to
	this domain
Some risk of bias	there some concern about bias with regard to this
	domain, although it is not clear that there is an
	important risk of bias
Clear risk of bias	the study has some important problems in this
	domain: characteristics of the study give rise to a
	clear risk of bias
Very high risk of bias	the study is very problematic in this domain:
	characteristics of the study give rise to a very high
	risk of bias



#### **ROBINS-E process**





Key messages

- **1. Risk of bias** is the appropriate way to think about study limitations, and needs to be addressed at multiple stages of a systematic review
- 2. Risk of bias assessments are **detailed and difficult** if you want to do them properly for observational studies
- **3. ROBINS-E is on its way** for [P][EC][O] questions
  - and there is parallel work for addressing multiple other study designs/analyses, including instrumental variables analyses (e.g. Mendelian randomization studies)

Guideline development using systematic reviews supplemented with internal health system data: the development and application of a conceptual framework

> North American Systematic Review Methods Virtual Research Day October 30, 2020

> > **Jennifer S. Lin, MD, MCR** Director, Kaiser Permanente EPC

> > > Helen Wu, PhD

Senior Manager, Kaiser Permanente Care Management Institute Evidence Services

#### Disclosures

Neither presenter has any financial conflicts of interests

The framework presented today was funded by AHRQ 290-2015-00007-I, Task Order 1- "Methods and Dissemination: Collaboration to improve validity, consistency, and utility of systematic reviews"

This presentation was initially submitted to the 2020 GIN Annual Conference

#### Aim:

To articulate a framework for using unpublished health system data alongside systematic reviews to inform guideline development and to explore its application in one health system's, Kaiser Permanente's, guideline program.

#### A Narrative Review and Proposed Framework for Using Health System Data with Systematic Reviews to Support Decision-making



Jennifer S. Lin, MD<sup>1,2</sup>, M. Hassan Murad, MD<sup>3</sup>, Brian Leas, MD<sup>4</sup>, Jonathan R. Treadwell, PhD<sup>4</sup>, Roger Chou, MD<sup>5</sup>, Ilya Ivlev, PhD<sup>1</sup>, and Devan Kansagara, MD<sup>6</sup>

<sup>1</sup>Kaiser Permanente Research Affiliates Evidence-based Practice Center, Portland, OR, USA; <sup>2</sup>The Center for Health Research, Kaiser Permanente Northwest, Portland, OR, USA; <sup>3</sup>Mayo Clinic Evidence-based Practice Center, Rochester, MN, USA; <sup>4</sup>ECRI Institute-Penn Medicine Evidence-based Practice Center, Plymouth Meeting, PA, USA; <sup>5</sup>Pacific Northwest Evidence-based Practice Center, Portland, OR, USA; <sup>6</sup>Veterans Health Administration Health Services Research Department Evidence Synthesis Program, Portland, OR, USA.

Systematic reviews are a necessary, but often insufficient, source of information to address the decision-making needs of health systems. In this paper, we address when and how the use of health system data might make systematic reviews more useful to decision-makers. We describe the different ways in which health system data can be used with systematic reviews, identify scenarios in which the addition of health system data may be most to improve the delivery of care (i.e., what to do and how to do it).<sup>1, 2</sup> Often, findings of systematic reviews are not clinically actionable due to low certainty in the evidence from published research, leaving decision-makers without a clear path forward. Even when an evidence base provides high certainty regarding the effectiveness of an intervention, reviews generally lack key contextual details that inform successful imple-

## Framing the problem

Systematic reviews are often a necessary but not sufficient information source for health system decision making

- often 'insufficient' evidence and a clinical decision needs to be made nonetheless
- often not clear the applicability of findings in the 'research' to local practice
- often evidence around net benefit (= effectiveness – harms) are insufficient for implementing a clinical service

Scenarios when health system data may be incorporated into or used in addition to systematic reviews

using both types of evidence together may:

Improve the strength of evidence, if...

 either data source has important methodological limitations

- either data source is imprecise
- either data source is limited to short-term followup
- either data source does not address important outcomes

Improve the applicability of evidence, if...

If systematic review or health system data are limited,

 systematic review data are indirect (have different population, intervention or setting than those of the health system)

 either data source does not allow for evaluation of effects in important subgroups Improve the implementation of evidence, if...

- systematic review data does not provide details required for replication or adaptation
- either data source lacks contextual information such as patients values and preferences, feasibility and acceptability
- either data source lacks information about cost
   effectiveness or cost

### Limitations and considerations when using unpublished primary data from health systems in systematic reviews

- Formal critical appraisal is a must
  - Biases and limitations for NRS well understood
- Numerous critical appraisal tools available, but may not be robust enough to understand limitations of real-world data (RWD)
  - RWD = data not collected for research purposes
- Vetting information quality and data quality
  - Information quality = the extent to which the data source can answer the question being asked
  - Data quality = integrity of the data (e.g., data accuracy, completeness, interpretability/accessibility, timeliness, mode of data collection)

#### Kaiser Permanente's National Guideline Program

Kaiser Permanente's Care Management Institute maintains a set of national clinical practice guidelines on selected topics

- Methods: Draw from existing external guidelines and systematic reviews when available, with critical appraisal and use of GRADE framework
  - Capacity to conduct internal analysis for high-priority topics that are not addressed well elsewhere
  - Avoid duplication of effort, focus on issues uniquely important for the KP health system
  - Importance of maintaining quality standards for internal analyses, where methods/limitations may not be documented in the explicit, transparent manner of published research
- Expertise: Clinical leaders bring insights about the gaps between external guidelines/systematic reviews and the answers clinicians need

#### Application of the Framework to KP's National Adult Diabetes Guideline

Торіс	Domain(s)	Feasibility*
Cost-effectiveness of treatments	Implementation – Cost for KP as an integrated delivery system is different	Most feasible
Third-step therapy	<b>Applicability</b> – Existing studies are indirect, do not explicitly address third-step therapy or specific combinations of interest	Possibly feasible
Use of SGLT-2 and GLP-1 agonists	<b>Applicability</b> – Existing studies are indirect, do not explicitly evaluate role of HbA1c levels; no head-to-head trials	Possibly feasible
Long-term harms of newer treatments	Strength – Research does not track long-term harms well	Least feasible

\* Key feasibility considerations: data access; formulary differences across the KP health system; timeline needed to measure outcomes of interest; sample size/power

#### Questions / Discussion



Examining Effect of Nutrition Interventions to Reduce Hyperphosphatemia in Chronic Kidney Disease: Is Including Non-Randomized Trials A Waste of Time?

> Mary Rozga, PhD, RDN Nutrition Researcher, Evidence Analysis Center Academy of Nutrition and Dietetics

### Conflicts of Interest



Employee for the Academy of Nutrition and Dietetics



No other conflicts of interest to disclose.

#### Background



#### **Chronic Kidney Disease (CKD)**

Individuals with CKD are at risk of hyperphosphatemia and resulting health outcomes, such as CKD Mineral and Bone Disorder and cardiovascular diseases.

#### **Study Designs for Nutrition Interventions**

- RCTs are the gold standard for examining the efficacy of healthcare interventions
- RCTs of nutrition interventions may not always be feasible
  - long periods of time to affect health outcomes
  - lack of generalizability
- High-quality only vs best evidence to support practice

#### Objectives

1. To examine difference in the effect size and certainty of evidence from RCTs only vs. RCTs + Non-RCTs in a nutrition intervention.

2. To examine the efficacy of phosphate-specific nutrition counseling provided by a dietitian, compared to usual care or an alternative intervention, on serum phosphate levels in individuals with CKD.

#### Systematic Review Methods



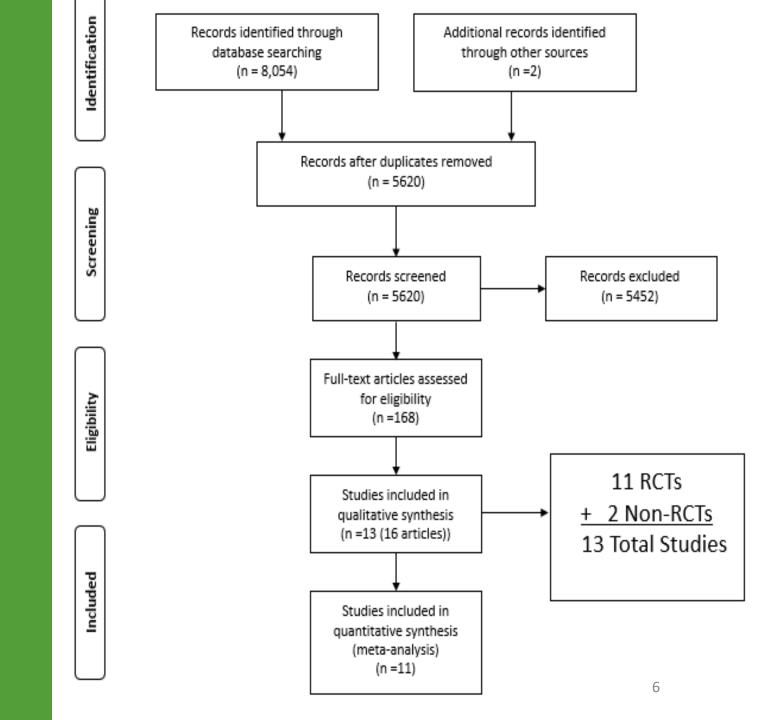
#### • Eligibility Criteria

- Individuals with CKD (P)
- Phosphate-specific nutrition therapy from a dietitian (I)
- Usual Care/Controlled Trials (C)
- Phosphate Levels (O)
- Literature Search
  - 2000-2019
  - MEDLINE, Embase, CINAHL, Web of Science and other databases

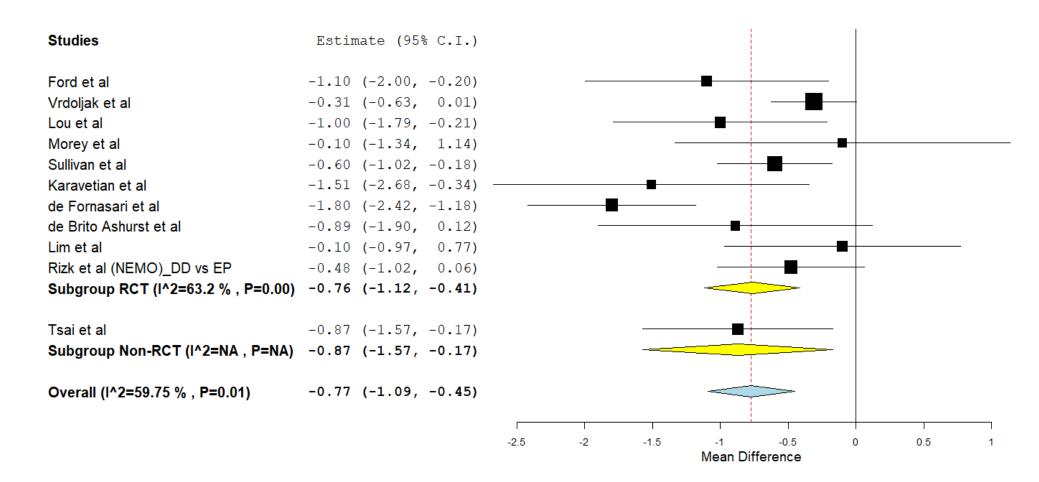
#### Risk of Bias Assessment

- ROB 2.0 for RCTs
- ROBINS-I for Non-RCTs
- Meta-analysis
  - Stratified by Study Design
- Quality of Evidence
  - Stratified by Study Design

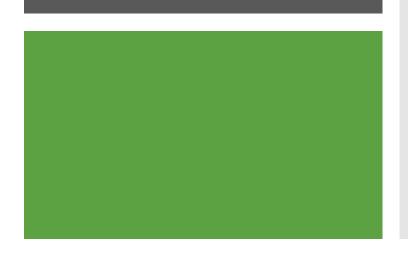
### Study Inclusion



# Effect of Phosphate-Specific Nutrition Counseling Provided by a Dietitian compared to Control on Serum Phosphate Levels (mg/dL) by Study Design (N=11/13 Studies)



### Results of Studies Not Included in Meta-Analysis



Two studies did not include data that could be pooled in metaanalysis.

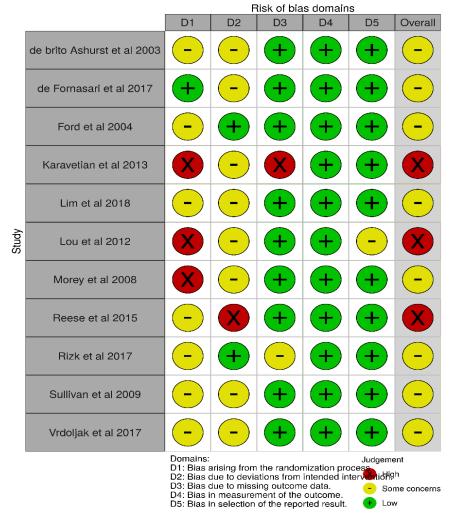
RCT

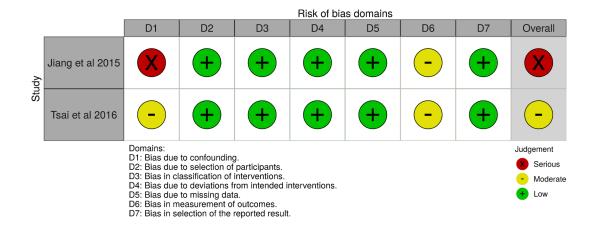
- Reese et al 2015
- Intervention group had a non-significant greater decrease in median change in serum phosphate at 10-weeks compared to the control group.

### Non-RCT

- Jiang et al 2015
- Intervention group had significantly reduced serum phosphate levels at 9 and 12 months compared to control group.
- Peritoneal Dialysis

## Risk of Bias: RCTs vs NRCTs





Serum Phosphate Levels: Summary of Findings

Outcomes	Anticipated absolute effects <sup>*</sup> (95% CI)			Cortainty of the	
	Risk with usual care	Risk with phosphate- focused diet therapy	№ of participants (studies)	Certainty of the evidence (GRADE)	Comments
Serum Phosphate Levels follow up: range 2.5 months to 12 months	Reference	MD <b>0.76 mg/dl<sup>c</sup></b> <b>lower</b> (1.12 lower to 0.41 lower)	Total: 1168 (11 RCTs) In Meta-analysis: 1144 (10 RCTs)	⊕⊕⊖⊖ Low ª	The evidence suggests phosphate-focused diet therapy from a dietitian reduces serum phosphate levels.
Serum phosphate levels follow up: 4 months to 12 months	Reference	MD <b>0.87 mg/dl<sup>c</sup> lower</b> (1.57 lower to 0.17 lower)	Total: 158 (2 Non-RCTs) In Meta-analysis: 61 (1 Non-RCT) <sup>h</sup>	⊕⊕⊖⊖ Low <sup>b</sup>	The evidence suggests phosphate-focused diet therapy from a dietitian reduces serum phosphate levels.

CI= Confidence Interval; MD= Mean Difference; RCT= Randomized Controlled Trial

<sup>a</sup> Risk of bias, moderate heterogeneity

<sup>b</sup>Risk of bias, small sample size, large confidence interval

## Limitations



## Lack of Non-RCTs included for comparison to RCTs



Only one study, a Non-RCT, included individuals on peritoneal- vs hemodialysis

## Conclusions

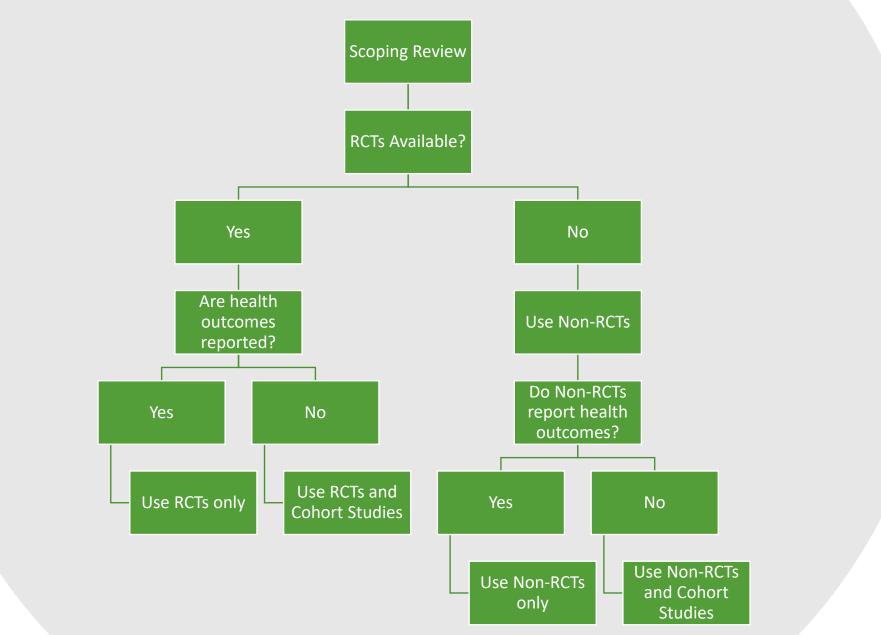
Overall effect size and certainty of evidence were not notably affected by including or excluding Non-RCTs when examining the effect of phosphate-focused nutrition therapy on phosphate levels in individuals with CKD on dialysis.

Suggest conducting a scoping review. When RCTs are identified, it may save time and effort to consider RCTs without Non-RCTs.

### Future Research

Does inclusion of long-term cohort studies improve understanding of the long-term feasibility and effects of nutrition interventions on patientcentered outcomes when assessed in tandem with RCTs?

### Suggested Process for Determining Study Design Inclusion



14

## Questions?

Mary Rozga, PhD, RDN mrozga@eatright.org

### EAC Evidence Analysis Center

right. Academy of Nutrition and Dietetics

Rozga M, Cheng F, Moloney L, Handu D. **Examining the effect of nutrition interventions to reduce hyperphosphatemia in chronic kidney disease: is including non-randomized trials a waste of time?** Advances in Evidence Synthesis: special issue *Cochrane Database of Systematic Reviews* 2020;(9 Suppl 1). https://doi.org/10.1002/14651858.CD202001

## References

1. Ashurst Ide B, Dobbie H: A randomized controlled trial of an educational intervention to improve phosphate levels in hemodialysis patients. J Ren Nut 13:267-274, 2003

2. de Fornasari ML, Dos Santos Sens YA: Replacing Phosphorus-Containing Food Additives With Foods Without Additives Reduces Phosphatemia in End-Stage Renal Disease Patients: a Randomized Clinical Trial. J Ren Nut 27:97-105, 2017

3. Ford JC, Pope JF, Hunt AE, Gerald B: The effect of diet education on the laboratory values and knowledge of hemodialysis patients with hyperphosphatemia. J Ren Nutr 14(1):36-44, 2004

4. Jiang N, Fang W, Gu AP, Yuan JZ, Yang XX, Lin AW, Ni ZH, Qian JQ: Improving diet recipe and cooking methods attenuates hyperphosphatemia in patients undergoing peritoneal dialysis. Nutr Metab Cardiovasc Dis 25:846-852, 2015.

5. Karavetian M, Ghaddar S: Nutritional education for the management of osteodystrophy (nemo) in patients on haemodialysis: a randomised controlled trial. J Ren Care 39:19-30, 2013

6. Lim E, Hyun S, Lee JM, Kim S, Lee MJ, Lee SM, Oh YS, Park I, Shin GY, Kim H, Morisky D, Jeong JC: Effects of education on low-phosphate diet and phosphate binder intake to control serum phosphate among maintenance hemodialysis patients: a randomized controlled trial. *Kidney Res Clin Prac* 37:69-76, 2018

7. Lou LM, Caverni A, Gimeno JA, Moreno R, Perez J, Alvarez R, B Campos, M García, A Gutiérrez, S Bielsa, J Castilla, A Sanz, F Martin, Aragon CKD Research Group: Dietary intervention focused on phosphate intake in hemodialysis patients with hyperphosphoremia. *Clin Nephrol* 77:476-483, 2012

8. Morey B, Walker R, Davenport A: More dietetic time, better outcome? A randomized prospective study investigating the effect of more dietetic time on phosphate control in end-stage kidney failure haemodialysis patients. *Nephron* 109:c173-180, 2008

9. Reese PP, Mgbako O, Mussell A, Potluri V, Yekta Z, Levsky S, Bellamy S, Parikh CR, Shults J, Glanz K, Feldman HI, Volpp K: A Pilot Randomized Trial of Financial Incentives or Coaching to Lower Serum Phosphorus in Dialysis Patients. *J Ren* Nutr 25:510-517, 2015

10. Rizk R, Hiligsmann M, Karavetian M, Evers S: Cost-effectiveness of dedicated dietitians for hyperphosphatemia management among hemodialysis patients in Lebanon: results from the Nutrition Education for Management of Osteodystrophy trial. J Med Econ 20:1024-1038, 2017

Sullivan C, Sayre SS, Leon JB, Machekano R, Love TE, Porter D, Marbury M, Sehgal AR: Effect of food additives on hyperphosphatemia among patients with end-stage renal disease: a randomized controlled trial. *JAMA* 301:629-635, 2009
 Tsai WC, Yang JY, Luan CC, Wang YJ, Lai YC, Liu LC, Peng YS: Additional benefit of dietitian involvement in dialysis staffs-led diet education on uncontrolled hyperphosphatemia in hemodialysis patients. *Clin Exp Nephrol* 20:815-821, 2016
 Vrdoljak I, Panjkota Krbavcic I, Bituh M, Leko N, Pavlovic D, Vrdoljak Margeta T: The impact of education and cooking methods on serum phosphate levels in patients on hemodialysis: 1-year study. *Hemodial Int* 21:256-264, 2017

METHODOLOGICAL REVIEW OF ITEMS FOR ASSESSING BIAS IN NETWORK META-ANALYSES PROVIDES GROUNDWORK FOR THE DEVELOPMENT OF A NEW RISK OF BIAS TOOL FOR NETWORK META-ANALYSIS (ROB NMA TOOL)

Lunny C, Tricco AC, Veroniki AA, Dias S, Hutton B, Salanti G, Wright J, Higgins J, White IR, Whiting P.

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### **ACKNOWLEDGEMENT OF TRADITIONAL LAND**

We wish to acknowledge the land in Vancouver, BC, Canada, on which the research is conducted.

For thousands of years it has been the traditional, ancestral, and unceded territory of the xwməθkwəyəm (Musqueam), Skwxwú7mesh (Squamish), Stó:lō and Səlĭlwəta?/Selilwitulh (Tsleil- Waututh) Nations.

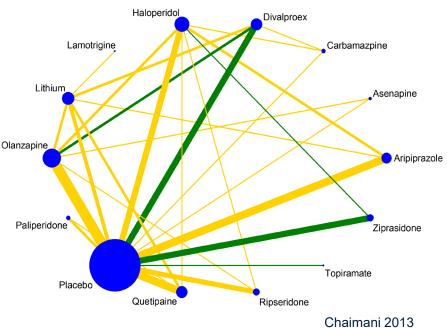
### WE HAVE NO ACTUAL OR POTENTIAL CONFLICTS OF INTEREST IN RELATION TO THIS PRESENTATION





### **SESSION OUTLINE**

- Background
  - Project team
  - Quality assessment tools
  - Rationale for a risk of bias tool for reviews Olanzapine with NMA
- Stages of development of our risk of bias tool
- Methodological review of bias items for NMAs
- Future steps







#### **Steering group:**

Carole Lunny, Andrea Tricco, Brian Hutton, Argie Veroniki, Georgia Salanti, Julian Higgins, Ian White, Sofia Dias, Penny Whiting

**Stakeholders**: PHAC, NICE, WHO, CADTH, G-BA, BC Support Unit, HTAi, Patient Voices Network, SPOR Evidence Alliance, Health Canada, Cochrane UK, Cochrane Consumer Network, Cochrane Canada, Cochrane Hypertension Group; Universities : Bristol, Ottawa Hospital Research Institute, University of British Columbia, University of Bern, University College London, University of Toronto, University of York



### QUALITY ASSESSMENT FOR REVIEWS WITH PAIRWISE META ANALYSIS

- A structured quality assessment tool provides a standardised providing consistency across reviews
- Many tools and checklists can be used for systematic reviews analysis:

Bias occurs if systematic flaws or limitations in the design, conduct or analysis distort the review conclusions

Number of checklists, instruments and tools				
Reporting	Quality of conduct	Risk of Bias		
60	40	1		

#### TOOLS AND CHECKLISTS TO AID IN SYSTEMATIC REVIEW CONDUCT, OR TO ASSESS THE REPORTING OR METHODOLOGICAL QUALITY OF A REVIEW

Tool purpose	Examples of tools or checklists for reviews with narrative summary or pairwise MA	Tools or checklists for reviews with NMA
Guidance for conducting systematic reviews	MECIR	Νο
Assess the quality of conduct of reviews	AMSTAR-2, OQAQ	ISPOR (International Society for Pharmacoeconomics and Outcomes Research)
Guidelines for the complete reporting published reviews	PRISMA	PRISMA-NMA , NICE-DSU
Assess the risk of bias of published reviews	ROBIS	Νο
Assess the certainty in evidence and the strength of recommendations	GRADE	GRADE-NMA, CINeMA, Threshold method



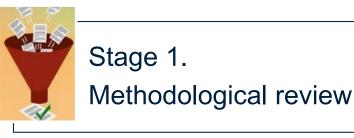
## RATIONALE FOR THE DEVELOPMENT OF A RISK OF BIAS TOOL FOR NMAS

- 1. Quality subpar. In a survey of 438 NMAs:
  - 75% considered moderate to low methodological quality using AMSTAR
  - ~50% inadequately reported 6 ISPOR network meta-analysis items
- 2. Novel elements in reviews with NMA require a bias assessment tool tailored to identifying NMA biases
  - Example, choice of node-making method (e.g. splitting, lumping) can drastically alter the network and subsequent results
- 3. Many reviews with NMA are published on the same topic with conflicting conclusions. For example, one study identified 28 NMAs on treatments for rheumatoid arthritis
  - Choosing a high quality NMA from multiple conflicting NMAs is difficult without a tailored risk of bias tool

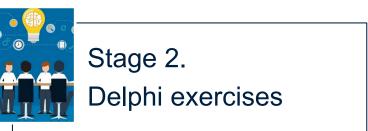


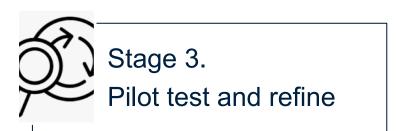
### STAGES OF TOOL DEVELOPMENT

- Our RoB NMA tool will address the degree to which the methods lead to a risk of bias in the review conclusions
- Objectives in 3 stages
- Methodology for developing a comprehensive and systematic risk of bias tool described in Whiting et al.'s "Framework for Developing Quality Assessment Tools"

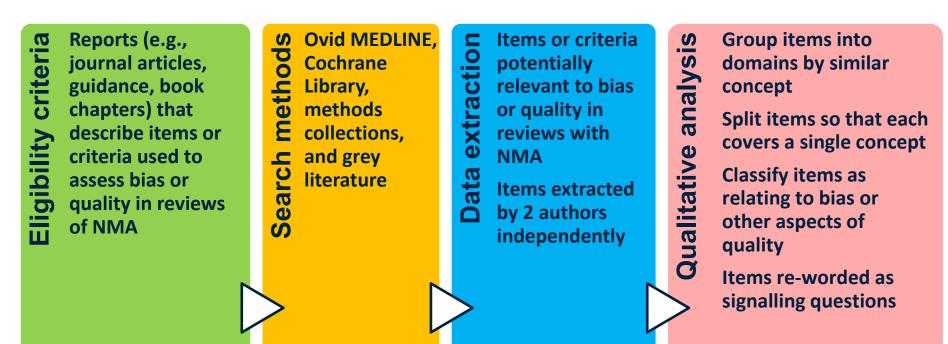








### **METHODS: METHODOLOGICAL REVIEW OF ITEMS**



### PRELIMINARY RESULTS: METHODOLOGICAL REVIEW OF ITEMS

• 56 studies included (2003-2020), which fall under these categories:

Tools	Methods	Guidance	Quality assessments of NMAs
16	21	12	7

- Included the updated PRISMA guidance for SRs with pairwise meta-analysis
- · Data extraction of items underway
- List of retained items will be refined by the steering group
- Final list of unique items will be compiled into a domain-based risk of bias tool



### **NEXT TWO STAGES**

- Conduct a multi-round Delphi process to solicit expert opinion on what items should be included
  - Sample of 50 experts will be invited
  - Consensus defined as 70% agreement
- Pilot test and refine the tool
- Knowledge translation strategy including training knowledge users in how to use the tool







#### THE UNIVERSITY OF BRITISH COLUMBIA

Abstract: Lunny C, Andrea T, Veroniki A, Wright J, White I, Dias S, Salanti G, Hutton B, Higgins J, Whiting P. *Methodological review of items for assessing the risk of bias in network meta-analyses provides groundwork for the development of a new risk of bias tool for network meta-analysis.* Advances in Evidence Synthesis: special issue *Cochrane Database of Systematic Reviews* 2020;(9 Suppl 1).

Carole Lunny, MPH, PhD, Cochrane Hypertension Group and Therapeutics Initiative, University of British Columbia, Vancouver, BC, Canada <u>carole.lunny@ti.ubc.ca</u>

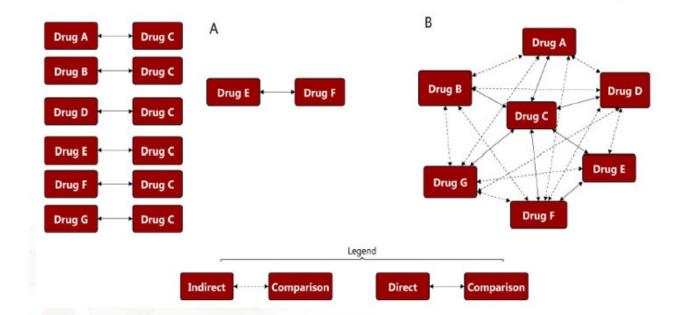
Twitter: @carole\_lunny and @Drug\_Evidence

### EXTRA SLIDES FOR QUESTIONS



#### WHAT IS AN NMA?

- Review with NMA aims to, or intends to, simultaneously synthesise more than two heath care interventions of interest
- Moving from individual pairs of comparisons to a network/unified meta-analysis





### **EXAMPLES OF BIASES IN NMA SYNTHESIS**

Node splitting bias. Methods can include:

- 1. Broad lumping approach that groups similar interventions at a broad level and is useful to estimate effects of intervention groups,
- 2. Grouping interventions with similar PICO elements together, taking account of clinically important variables,
- 3. Lumping-and-dismantling approach informed by meta-regression to investigate effects attributed to different components
- Class-effect model approach that lumps similar interventions together as a class but assumes effect variations between these interventions, using modeling to estimate effects of specific interventions
- Rank/ probabilities can be biased if the uncertainty of some treatment effects is larger than on others
- Distortions of summary estimates in cases with between-study heterogeneity that are not accounted for (e.g. with meta-regression). These can impact the ordering of treatments and treatment effects
- Pre-specification of methods is essential and should be based on objectives



October 30, 2020

### DEALING WITH RETRIEVAL BIAS FOR AN EVIDENCE-INFORMED INDIVIDUAL PATIENT DATA NETWORK META-ANALYSIS

Areti Angeliki Veroniki, MSc, PhD

Co-authors: Ashoor H, Rios P, Seitidis G, Mavridis D, Straus S, Tricco A

<u>Prepared for</u>: Systematic Review Methods Research Day



## CONFLICTS OF INTEREST

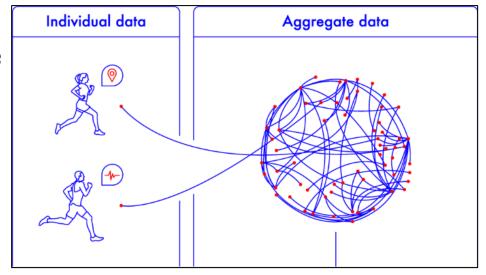
# I have no actual or potential conflict of interest in relation to this presentation

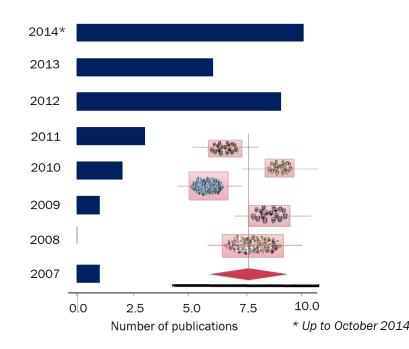
This project was partly funded by the Canadian Institutes of Health Research Drug Safety and Effectiveness Network and the Canadian Institutes of Health Research Knowledge Synthesis

## INDIVIDUAL PATIENT DATA NETWORK META-ANALYSIS

Personalized medicine is required to optimize health care IPD meta-analyses: use data from each individual patient enrolled in each included trial

 Gold standard for synthesising evidence across clinical trials





IPD indirect comparisons are published with increasing frequency in health care literature

Veroniki et al BMC Med Res Methodol 2016

## SHARING INDIVIDUAL PATIENT DATA (IPD) FROM RCTS

- Network meta-analyses modelling IPD usually include non-sponsored or publicly sponsored RCTs
- Evidence suggests that IPD sharing may depend on study characteristics, such as funding type, RCT size, RCT risk of bias, and treatment effect
- Retrieval bias in IPD network meta-analysis of sponsored RCTs has not been assessed before
- What are the challenges and barriers?



Veroniki et al J Clin Epidemiol. 2019

## SHARING INDIVIDUAL PATIENT DATA (IPD) FROM RCTS

- Missing data can distort the medical literature and harm patients when erroneous decisions are made
  - IPD meta-analyses based only on a portion of the trials can affect the results (selection bias)!

- 33% of the negative responses ٠ were due to:
  - lack of resources or time,
  - lack of ownership or IPD, • and
  - old IPD that could not be retrieved



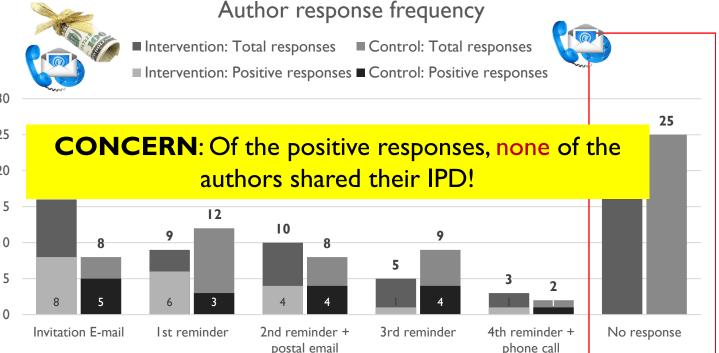
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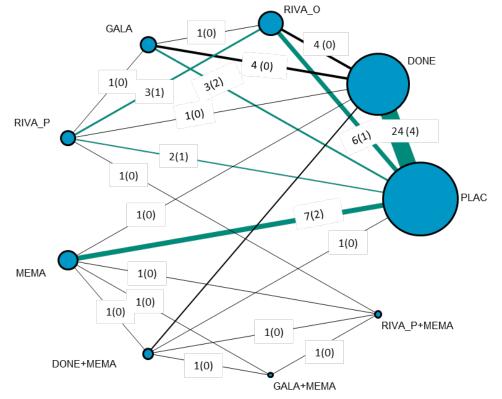
Veroniki et al | Clin Epidemiol. 2019

## SYSTEMATIC REVIEW AND NETWORK META-ANALYSIS



To individualize the management of patients with Alzheimer's dementia To fill an important knowledge gap in health care, and to inform decision making

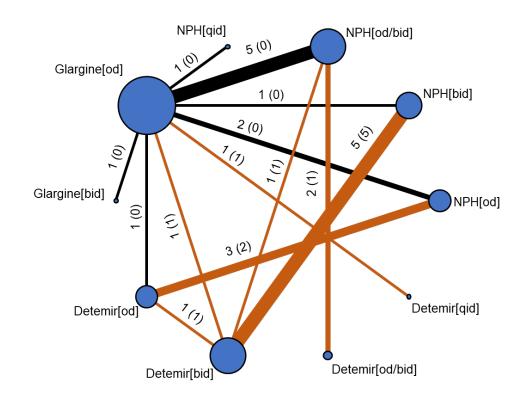
Veroniki et al BMJ Open 2016





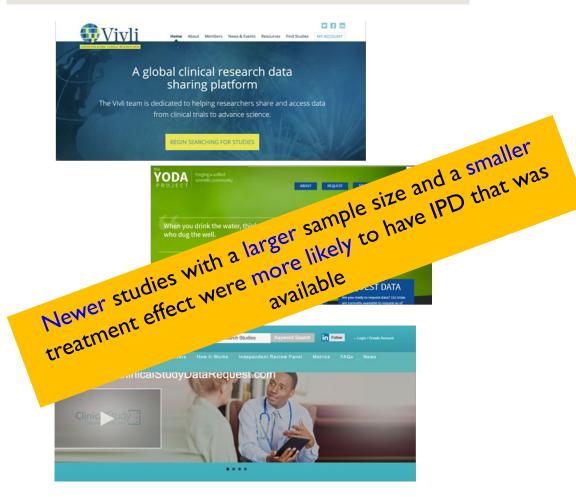
To inform clinical practice guidelines in the development of tailored management recommendations for patients with type I diabetes

Veroniki et al BMJ Open 2015



## INDIVIDUAL PATIENT DATA NETWORK META-ANALYSIS

Data sharing was only possible though proprietary sponsor-specific platforms!



### **Alzheimer's Dementia**

- I5 sponsors were contacted for 82 RCTs
- 6 (40%) sponsors shared their data through proprietary sponsor-specific platforms
- 6 sponsors were contacted for 46 RCTs (14,580 participants)
- We obtained IPD for 14 RCTs (8,007 participants)
  - 1,058 total waiting days up to March 9,2020

### TIDM

- 2 sponsors were contacted for 25 RCTs
- I (50%) sponsor shared their data through a proprietary sponsor-specific platform
- We obtained 12 RCTs (4,877 participants)

## INDIVIDUAL PATIENT DATA NETWORK META-ANALYSIS

### **Alzheimer's Dementia**

- We were able to include 12 RCTs in our NMA due to incompleteness of provided data
   A study included only IPD for the placebo arm
  - A study did not include outcome data
- A big challenge in the IPD was the high dropout rate from the RCTs
- Two studies did not report an outcome of interest in the final publication, but in the retrieved IPD we were able to use data for this outcome

### TIDM

• We were able to include 12 RCTs in our NMA



## PRIMARY CHALLENGES ENCOUNTERED



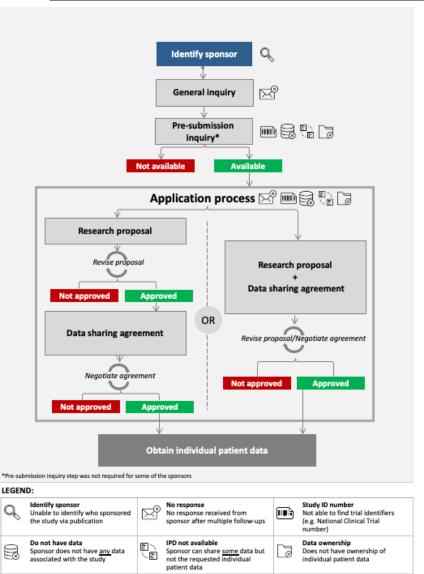


**Challenges** encountered included:

- Identification of trial data set when certain details were not available (e.g. NCT number)
- Data ownership

- Sponsors **switched platforms**, while we were navigating the data
- **Software availability**: Required R packages (e.g., mice) were not available/provided
  - we were not allowed to install any new R packages; R packages were older versions (e.g. Ime4)
- IPD available through proprietary sponsor-specific platforms did not allow a **one-stage analysis** as planned in our protocol

## PRIMARY CHALLENGES ENCOUNTERED



- **Time** that the platform permitted access to the individual patient data was often **limited**
- **Cost** associated with **obtaining access** to the data for certain time
- Cost associated with the WHO Drug Dictionary license to obtain access to the additional medications used for each patient
- Available individual patient data did not include the **full information** as shown in the publication:
  - Only data for placebo were available, or
  - Did not give information about a reported outcome (e.g. only baseline MMSE values were available).
  - Date of follow-up was coded in some studies and it was impossible to make a judgement on first and last date

Veroniki et al JCE 2019, DOI: 10.1016/j.jclinepi.2019.05.031

### HOW THESE FINDINGS WILL INCREASE SCIENTIFIC KNOWLEDGE

- Well-conducted individual patient data network meta-analyses facilitate tailored decision making
- We were able to obtain data for studies that did not report outcome data in the original publication
- Retrieval bias can severely impact the knowledge synthesis findings and decisionmaking
- We retrieved individual patient data for 15% (12/80) of the eligible RCTs in Alzheimer's dementia and 46% (12/26) of RCTs in type I diabetes
- IPD sharing is not yet well-established in the field of Alzheimer's dementia and type I diabetes, and more efforts are required to achieve this goal

## **QUESTIONS?**



@AVeroniki







# depressionscreening100.com/phq: A practice-based perspective to using the Patient Health Questionnaire-9 to screen for depression

October 30, 2020

## **Brooke Levis**

Co-authors: Yuliia Dehtiarova, Ying Sun, Yin Wu, Andrea Benedetti, & Brett D. Thombs



• I have no personal or financial conflicts of interest to disclose

• The abstract for this presentation was originally submitted to the 2020 Cochrane Colloquium

Levis B, Dehtiarova Y, Sun Y, Wu Y, Benedetti A, Thombs BD, and the DEPRESSD PHQ Collaboration. depressionscreening100.com/phq: a practice-based perspective to using the Patient Health Questionnaire-9 to screen for depression. In: Advances in Evidence Synthesis: special issue. *Cochrane Database of Systematic Reviews* 2020;(9 Suppl 1):53.

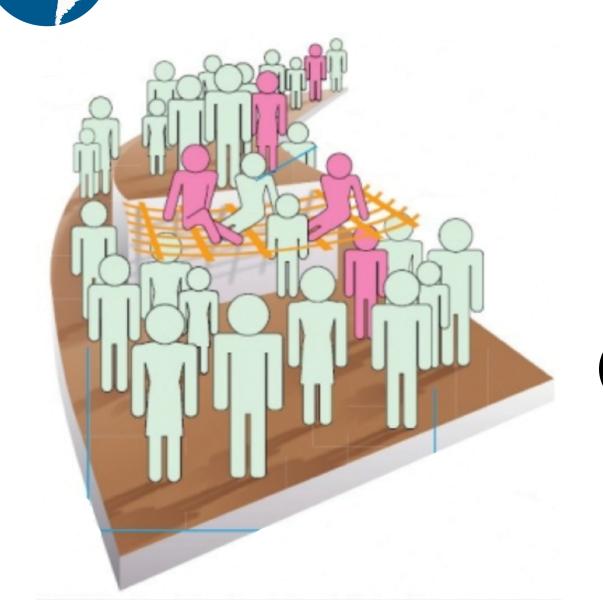


# Depression in medical settings

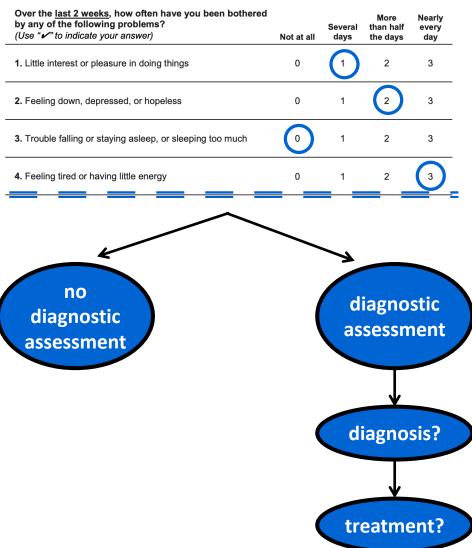
- Common and disabling condition
- Highly prevalent
  - General population: ~5%
  - Primary care: 10%
  - Specialty care: 10-20%
- Associated with poor prognosis
- One possible solution: routine depression screening



**DEPRESSD** Depression screening



#### PATIENT HEALTH QUESTIONNAIRE-9 (PHQ-9)



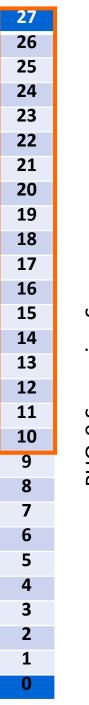


# Is the PHQ-9 accurate? **Diagnostic test accuracy**

### **Diagnostic Interview**

			<b>Depression</b> +	<b>Depression</b> –
PHQ-9	Screening	Screen +	a	b
	ΤοοΙ	Screen –	С	d

- Sensitivity: a/(a+c)
- Specificity: d/(b+d)
- Positive Predictive Value: a/(a+b)
- Negative Predictive Value: d/(c+d)



PHQ-9 Screening Score

4



# PHQ-9 diagnostic accuracy



#### RESEARCH

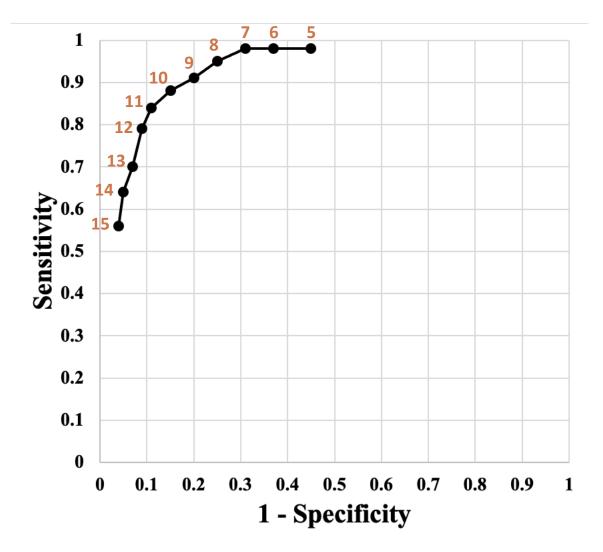




Accuracy of Patient Health Questionnaire-9 (PHQ-9) for screening to detect major depression: individual participant data meta-analysis

Brooke Levis,<sup>1</sup> Andrea Benedetti,<sup>2</sup> Brett D Thombs,<sup>1</sup> on behalf of the DEPRESsion Screening Data (DEPRESSD) Collaboration





A cutoff of ≥ 10 maximized combined sensitivity and specificity

6

- Sensitivity = 0.88
- Specificity = 0.85



# What does this mean?

### Findings

 PHQ-9 has the greatest combination of sensitivity and specificity at a cutoff of ≥ 10

### So what?

- What cutoff should clinicians use in clinical practice?
- Should clinicians <u>always</u> use a cutoff of ≥ 10?
- Are sensitivity and specificity equally important?
- What does "88% sensitivity" even mean?



Knowledge Translation web tool: <u>http://depressionscreening100.com/phq/</u>

### Objective and purpose:

- To create a user-friendly knowledge translation tool based on sensitivity and specificity estimates from the IPDMA
- The tool allows clinicians to estimate, for a given depression prevalence and PHQ-9 screening cutoff score:
  - How many patients would screen positive versus negative
  - How many in each group would be correctly versus incorrectly identified

#### Knowledge user involvement:

• We consulted with family physicians during development



# Knowledge Translation web tool: http://depressionscreening100.com/phq/

#### Please type in the prevalence in %

Enter an integer between 1 and 100

#### Prevalence

#### 10

The default cutoff threshold is the standard cutoff score of 10 or greater. Use the slider below to select a lower or higher cutoff threshold

Cutoff		
	•	 
	Calculate	

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Based on the prevalence you entered, **22** of **100** patients (22%) in your practice would screen positive for possible depression.

Of the 22 patients who screen positive:

**9** (39%) would meet diagnostic criteria for major depression (true positives)

**13** (61%) would not meet diagnostic criteria for major depression (false positives)

Based on the prevalence you entered, **78** of **100** patients (78%) in your practice would screen negative for possible depression.

Of the **78** patients who screen negative:

**77** (99%) would be correctly ruled out (true negatives)

1 (1%) would be missed major depression cases (false negatives)



- This web-tool improves clinician understanding of results from our meta-analysis by
  - Translating results into numbers that are more readily understood
  - Providing guidance



# Acknowledgements

- Co-authors: Yuliia Dehtiarova, Ying Sun, Yin Wu, Andrea Benedetti, and Brett D. Thombs
- PhD Supervisors: Brett Thombs and Andrea Benedetti
- DEPRESSD Steering Committee: Brett Thombs, Andrea Benedetti, Jill Boruff, Pim Cuijpers, Simon Gilbody, John Ioannidis, Lorie Kloda, Sarah Markham, Dean McMillan, Scott Patten, Ian Shrier, and Roy Ziegelstein
- DEPRESSD Project Team
- DEPRESSD-PHQ Data Contributors
- DEPRESSD Funding: CIHR
- PhD Funding: CIHR and FRQS





### Progress toward a reporting guideline for overviews of reviews of healthcare interventions: Preferred Reporting Items for Overviews of Reviews (PRIOR)

Michelle Gates<sup>1</sup>, Allison Gates<sup>1</sup>, Michelle Pollock<sup>2</sup>, Ricardo M Fernandes<sup>3,4</sup>, Dawid Pieper<sup>5</sup>, Andrea C Tricco<sup>6,7,8</sup>, Lisa Hartling<sup>1</sup>

<sup>1</sup>Alberta Research Centre for Health Evidence, Department of Pediatrics, University of Alberta, Canada
<sup>2</sup>Institute of Health Economics, Canada
<sup>3</sup>Clinical Pharmacology Unit, Instituto de Medicina Molecular, Faculty of Medicine, University of Lisbon, Portugal
<sup>4</sup>Department of Pediatrics, Santa Maria Hospital, Portugal
<sup>5</sup>Institut für Forschung in der Operativen Medizin, Department für Humanmedizin, Universität Witten/Herdecke, Germany
<sup>6</sup>Li Ka Shing Knowledge Institute of St. Michael's Hospital, Canada
<sup>7</sup>Dalla Lana School of Public Health & Institute of Health Policy, Management, and Evaluation, University of Toronto, Canada
<sup>8</sup>Queen's Collaboration for Health Care Quality Joanna Briggs Institute Centre of Excellence, Queen's University, Canada

North American Systematic Review Methods Virtual Research Day - October 2020



#### Abstract accepted for the 2020 Cochrane Colloquium.

Gates M, Gates A, Pollock M, Fernandes RM, Pieper D, Tricco AC, Hartling L. Progress toward a reporting guideline for overviews of reviews of healthcare interventions: Preferred Reporting Items for Overviews of Reviews (PRIOR). In: Advances in Evidence Synthesis: special issue. Cochrane Database Syst Rev. 2020;9 Suppl 1:400.

The authors have no conflicts of interest to declare.

### **Background and rationale**

#### WHAT ARE OVERVIEWS OF REVIEWS?

Overviews of reviews use **systematic methodology** to search for and synthesize data from **multiple systematic reviews (SRs)** on a similar topic with the purpose of mapping, synthesizing, and/or exploring discrepancies in the evidence.

### **UNIQUE CHALLENGES** when the unit of analysis is the SR

+

#### **INCREASED COMPLEXITY**

Need to deal with two layers of information:

Systematic reviews

their included primary studies

#### METHODOLOGICAL UNCERTAINTY

Recommendations are abundant but fragmented and inconsistent

EVIDENCE- and CONSENSUS-BASED REPORTING GUIDANCE UNAVAILABLE

#### > PRIOR





**OBJECTIVE** To develop an **evidence- and agreement-based reporting guideline** for overview of reviews of healthcare interventions using explicit, systematic, transparent methods based on guidance of the EQUATOR Network

#### **METHOD**

<ul> <li>PROJECT LAUNCH</li> <li>Identify and invite experts</li> </ul>	<ul> <li>LIT REVIEW</li> <li>Methodological review of reporting</li> <li>Scoping review of methods for conduct</li> </ul>	<b>DELPHI</b> <ul> <li>Recruit participants</li> <li>Two online rounds</li> <li>One in-person round (virtual)</li> </ul>	<ul> <li><b>DEVELOP GUIDANCE</b></li> <li>– Write initial draft</li> <li>– Pilot test</li> <li>– Disseminate</li> </ul>	
$\boldsymbol{\bigotimes}$	$\bowtie$	$\bowtie$		
Expert advisory board (EAB)	Prospective item list	Agreement on preliminary item list	Final PRIOR guideline	





### ESTABLISHED A CORE TEAM responsible for day-to-day operations, and

an international and interdisciplinary expert advisory board

#### ROLE OF THE EXPERT ADVISORY BOARD

- Provide expertise related to overview methodology and guideline development
- Nominate participants for the Delphi exercise, participate, provide feedback
- Help to plan and facilitate the in-person meeting
- Assist in producing the guideline, dissemination, knowledge translation

**REGISTERED INTENT** to develop PRIOR with the EQUATOR Network (2017)

**PLANNED** the project goals, steps, preliminary timelines, and published an apriori protocol (2019)

#### **OBTAINED ETHICS APPROVAL** to undertake the project





### **Step 2: Literature reviews**

#### **METHODOLOGICAL REVIEW - REPORTING**

- <30% describe a protocol, synthesis methods, quality of primary studies with the reviews, certainty of evidence
- <10% describe how primary studies were considered, how they dealt with discordant reviews

#### **SCOPING REVIEW - METHODS GUIDANCE**

- 77 guidance documents available
- Several areas of conflicting or lacking guidance
  - Whether, how, and when to include primary studies
  - How best to identify and manage primary study overlap
  - Rating the certainty of the evidence
- Limited evidence to support methodological decisions





EVIDENCE-BASED preliminary list of candidate items





Icon courtesy of Freepik via flaticon.com

### **Preliminary item list**

### **STAND-ALONE GUIDELINE (not a PRISMA extension)**

- Many items similar to PRISMA, but unit of analysis differs (systematic reviews)
- Allowed us to focus on particular challenges related to overviews
- Intended to facilitate future guideline extensions (e.g., diagnostic overviews)

### **ITEMS UNIQUE TO OVERVIEWS (examples)**

#### **RATIONALE and SCOPE**

Describe why an overview of reviews format is the most appropriate methodology for answering the research question



#### **ELIGIBILITY CRITERIA**

Specify the pre-established definition of a systematic review used as a criterion for inclusion in the overview of reviews.



#### DATA EXTRACTION

State any methods used to deal with overlapping data from primary studies within the included systematic reviews during data extraction. State the method used to illustrate and/or quantify the degree of overlap across included systematic reviews.





Icons courtesy of fjdesigns and Freepik via flaticon.com

### **Step 3: Modified Delphi**

"The Delphi is a group facilitation technique that aims to obtain consensus from a group of experts"

- Iterative process where participants provide feedback in multiple rounds
- After each round, the findings are analyzed and summarized for participants
- Participants review group responses and re-consider their original decision in subsequent rounds, until a high level of agreement is reached (≥70%)

#### **PURPOSIVE SAMPLE**

100 international participants with diverse expertise in conducting, reviewing, disseminating, and using overviews

#### ONLINE DELPHI 1 & 2

Given preliminary items and available evidence, participants vote to include/exclude (5-point scale)

#### **SELECTED PARTICIPANTS**

Subset of 10 expert panelists invited to an in-person meeting using the nominal group technique to reach agreement on final items





Hasson et al. J Adv Nurs 2000;32(4):1008-15.

## **Step 3: Modified Delphi - progress**

#### **ROUND 1:** 53 participants (53% response)

- ✓ Agreement (≥70%) for the **inclusion of 48** of 52 items
- More than 500 unique qualitative comments

#### ROUND 2: 44 participants (83% return)

- × No agreement on 9 remaining items (many were close)
- ✓ More than 250 unique qualitative comments

#### **IN-PERSON:** 13 selected participants

- 2 x 2-hour virtual meetings to deliberate and re-vote
- Discussion of next steps (pilot-testing, dissemination)





Reworded 1 item Added 5 items







Icon courtesy of Freepik via flaticon.com

### **Step 4: Guidance statement development**

#### **REFINEMENT OF ITEM LIST**

- Edit items based on participant comments, with the aim of producing clear, concise, and unambiguous wording for each item
- May involve condensing and/or re-organizing the checklist to a manageable length to enhance usability (e.g., 20 items + sub-items)

#### **PILOT-TESTING**

- A group of potential users will test the checklist
- Finalization of the checklist based on user feedback and input of the expert panel (in-person attendees)

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Icon courtesy of Freepik via flaticon.com

### **Step 4: Guidance statement development**

#### WRITING

A writing group consisting of the core team, EAB, and expert panelists from inperson meeting will draft the initial manuscript, explanation and elaboration document

#### DISSEMINATION







Icon courtesy of icongeek36, Freepik via flaticon.com



# **THANK YOU** Look forward to PRIOR in 2021!

**Acknowledgments:** We thank all participants of the Delphi rounds, and Drs. Amanda Newton and Shannon Scott for input into the protocol. LH is supported by a Tier 1 Canada Research Chair in knowledge synthesis and translation. ACT is supported by a Tier 2 Canada Research Chair in knowledge synthesis.



mgates1@ualberta.ca





# Accelerating integration of emerging evidence into health care delivery: rapid reviews for learning health systems

Presenter: Marcy Hager, Director of Evidence-based Practice Program, Oregon Health & Science University

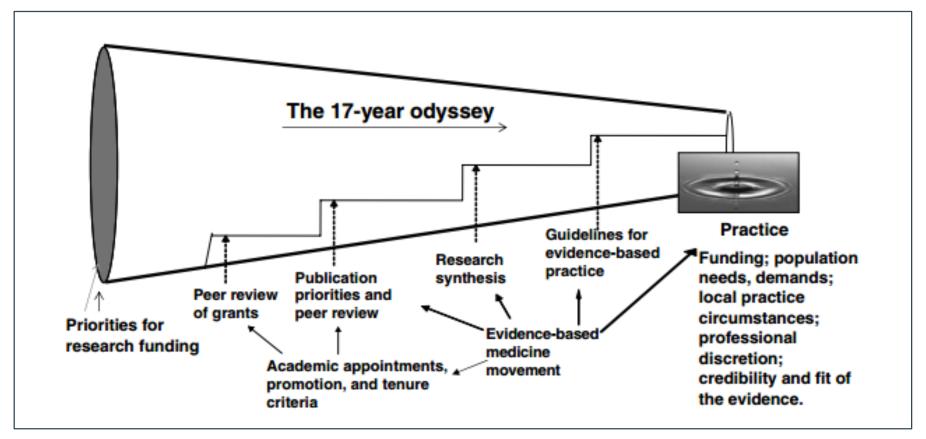


Presenter: Marcy Hager

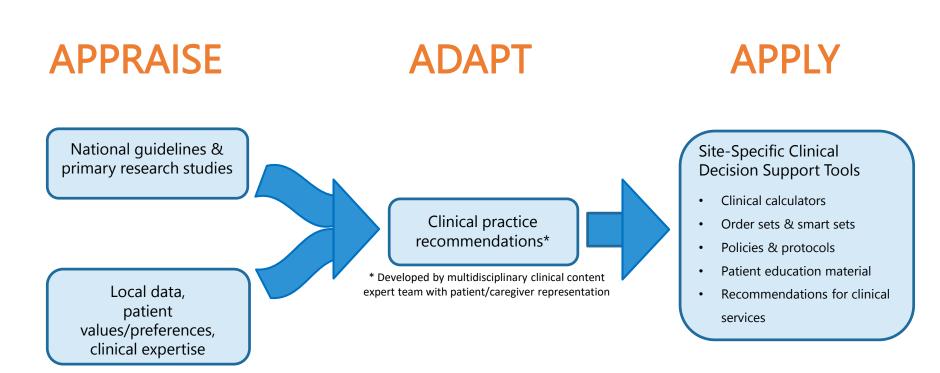
I have no personal financial relationships with commercial interests relevant to this presentation to disclose.

Learning Health Systems Why is there a need for a localized rapid review process?

# The 17-Year evidence-topractice gap

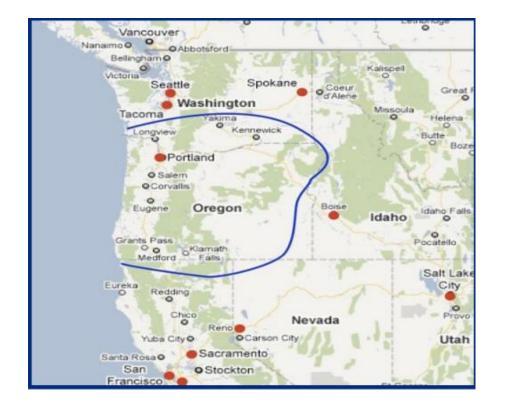


**Source:** Green LW, Ottoson J, Garcia C, Robert H (2009). Diffusion theory and knowledge dissemination. *Annual Review of Public Health* 30, 151.



OHSU Health Evidence-based Practice Program How do we get there?

## **OHSU Health System**



Oregon Health Science University (OHSU), Hillsboro Medical Center and Adventist Health formed OHSU Health System with the goal of transforming members into a value-based care organization through partnership and innovation.

# Office of Clinical Integration and Evidence-based Practice

- Established with the goal of integrating best research evidence into clinical practice.
- Supports the development and implementation of evidence-based guidelines
- Goal: "One Standard of Care" across OHSU Health

# **EBP Deliverables**

### OHSU Health System Evidencebased Clinical Guidelines

**Evidence Briefs** 

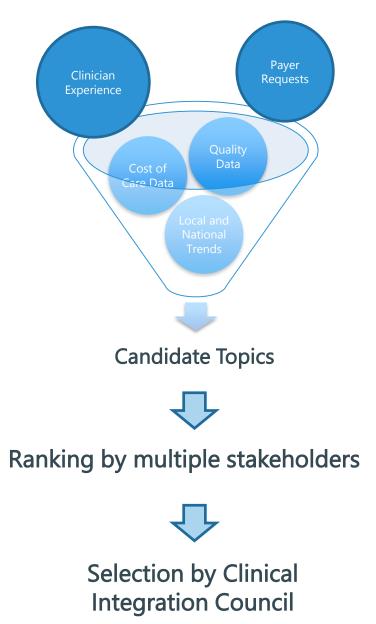
**EBP** Interdisciplinary Course

# **Clinical Guidelines**

# **Clinical Integration Council**

- Interdisciplinary body was formed to oversee care standardization throughout the health system by:
  - Prioritizing OHSU Health System Guidelines
  - Identifying clinical champions
  - Removing barriers
  - Providing resources
- Clinical Advisory Council includes following members:
  - Chief Medical Officers (OHSU, Hillsboro and Adventist)
  - Chief Nursing Officers (OHSU, Hillsboro and Adventist)
  - Physician Champions (OHSU, Hillsboro and Adventist)
  - Quality Officer
  - Senior Associate Dean

### **Guideline Selection Process**

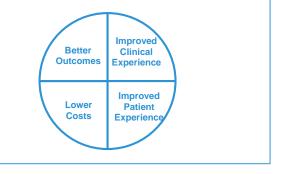


Weighted Overall Project Score:	Project Numbe	÷r:	1	
0.00				
Criteria	Score	Weight	Weighted Score*	Issues/Concerns
1.0 Sponsorship		0.10	0.00	
2.0 Quality, Pt Safety and Experience Improvement Benefit		0.30	0.00	
<list be="" impacted="" outcomes="" to=""></list>				
3.0 Financial Benefits		0.25	0.00	
<list areas="" be="" impacted="" to=""></list>				
4.0 At Risk Populations		0.10	0.00	
5.0 Impact/Effort		0.15	0.00	
6.0 Best Practices		0.05	0.00	
7.0 Provider Wellness/Satisfaction		0.05	0.00	
TOTAL (sum of weighted score column)		1	0.00	

Note: Any criterion scores of zero must be addressed before project is approved

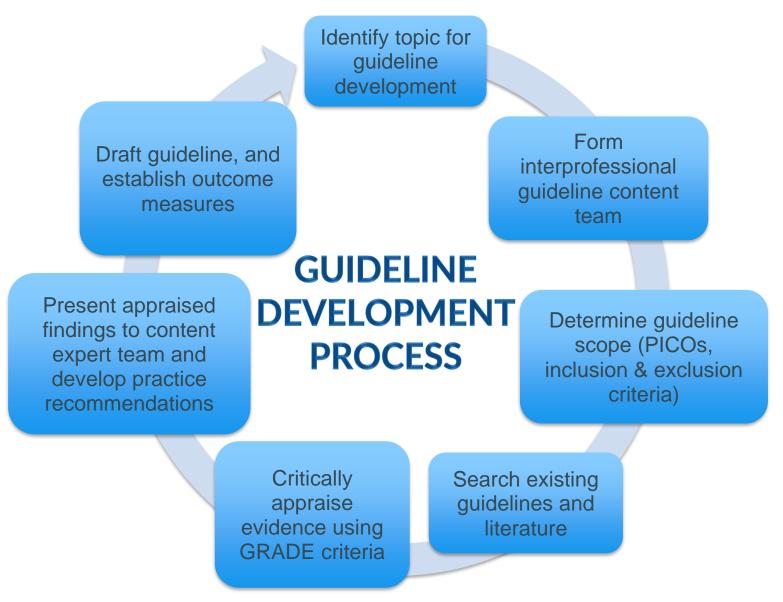
\* Weighted score = project's score for each criterion times the weight.





Guideline Topic	Sponsorship	Quality/Pt Experience	Financial Benefit	At-Risk Population	High Impact/ Low Effort	Best Practices	Provider Wellness
Adult Cystic Fibrosis Pain and Anxiety	Ø		Ø				
Safe Opioid Prescribing for Chronic, Non-End-of-Life Pain	Ø	<b>I</b>	Ø				
Supplemental Feeding in Healthy, Term Neonates			Ø			Ø	Ø
Colorectal Cancer Screening		Ø	Ø				$\bigcirc$
Heart Failure			Ø	Ø		Ø	Ø
Acute Low Back Pain				Ø			
Pancreatitis	Ø						Ø
Pediatric Urinary Tract Infection	Ø		Ø			$\bigcirc$	
Sickle Cell Disease	Ø		Ø				$\bigcirc$
Induction of Labor	Ø	Ø		Ø		$\bigcirc$	Ø

# Step 1: Guideline Development



# **Step 2: Guideline Implementation**

Collect Baseline Data	Communicate and Educate	Develop Decision Support Tool	Develop the Process Metrics	Identify Programmatic Recommendations
<ul> <li>What can we collect already</li> <li>Data work and workflow impact</li> <li>Workflow integration</li> </ul>	<ul> <li>Patient Education Materials</li> <li>Staff Materials</li> <li>Patient Communication Materials</li> <li>Expert Talking Points</li> <li>Tool Kits for Site Implementation</li> </ul>	<ul> <li>Links to Guidelines</li> <li>Document Templates</li> <li>Flow Sheets</li> <li>EHR Changes</li> <li>Best Practice Alerts</li> <li>Order Sets</li> </ul>	<ul> <li>Answer how well we are using the tools</li> <li>Resource</li> <li>Communication</li> <li>Project Coordination</li> <li>Site Coordination</li> <li>Site Coordination</li> <li>Develop the reporting tool</li> <li>Publish the reporting tool</li> <li>Identify resources</li> </ul>	<ul> <li>Develop a Business Case</li> <li>Develop an ROI evaluation</li> <li>Develop Budgeting Reports</li> </ul>

# Impact from Guidelines

Outcome Measures Post-Implementation						
Cystic Fibrosis	<ul> <li>Opioid-using encounters during hospitalization decreased 9% at 1- year post-implementation.</li> <li>Encounters in which IV opioids were prescribed decreased 8%.</li> <li>Morphine equivalent dose/day (MEDD) was reduced by 1.4.</li> <li>Length of stay reduced 1/5 days.</li> </ul>					
Opioid Prescribing	<ul> <li>Reduced <b>4660</b> opioid prescriptions (11% to 9%) during 17 months of follow-up (P &lt; .0001).</li> <li>Reduced average MEDD per prescription from 21.1 to 16.8 (P = .009).</li> </ul>					
Supplemental Feeding	• Pre-intervention, median documentation was 0; this rose to:					
	> 78.6% after provider and staff education					
	> 84.9% after the integration of charting tools					
	100% after RNs began calculating & documenting with tool					
Heart Failure	<ul> <li>All cause readmissions decreased from 17.5% to 11%</li> <li>Related readmissions decreased from 12% to 7%</li> </ul>					

# Lessons Learned

- Executive sponsorship and support is essential
- Include patients in guideline development and implementation
- Engage EHR team early on to ensure recommendations are realistic
- Ensure appropriate stakeholders are at the table
- Focus implementation efforts
- Time-consuming
- Competing demands
- Behavior change is more than just Epic tools

# **EBP Program Partnerships**

- Medical Librarian
- Pacific Northwest Evidence-based Practice Center
- Data Analyst/Report Writing
- Health System Effectiveness
- Quality/Performance Improvement
- Alignment with Health System Initiatives such as
  - Integrated Delivery System
  - Population Health
  - Value-based Care



# Thank You

Contact Information: hagerm@ohsu.edu

Inspired by research. Inspiré par la recherche. Driven by compassion. Guidé par la compassion.

# Rapid Reviews Methods – A Scoping Review

CANDYCE HAMEL, MSC, PHD(C) KNOWLEDGE SYNTHESIS GROUP, OHRI

NORTH AMERICA SYSTEMATIC REVIEW METHODS VIRTUAL RESEARCH DAY OCTOBER 30, 2020



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**Declaration of interest: None.** 

#### **Rapid review methods**

Check for updates



Journal of Clinical Epidemiology 126 (2020) 131-140

#### REVIEW

Journal of Clinical

Epidemiology

#### Few evaluative studies exist examining rapid review methodology across stages of conduct: a systematic scoping review

Candyce Hamel<sup>a,b,\*</sup>, Alan Michaud<sup>a</sup>, Micere Thuku<sup>a</sup>, Lisa Affengruber<sup>c</sup>, Becky Skidmore<sup>a</sup>, Barbara Nussbaumer-Streit<sup>c</sup>, Adrienne Stevens<sup>a</sup>, Chantelle Garritty<sup>a,b</sup>

<sup>a</sup>Ottawa Hospital Research Institute, Knowledge Synthesis Group, Ottawa, Ontario K1H 8L6, Canada <sup>b</sup>School of Medicine, University of Split, Split, Croatia 21000 <sup>c</sup>Cochrane Austria, Department for Evidence-based Medicine and Evaluation, Danube University Krems, Krems, Austria Accepted 23 June 2020; Published online 26 June 2020

#### Abstract

Objectives: The objective is to identify studies that have assessed methodological shortcuts for undertaking rapid reviews (RRs) and mapping these to review conduct stages and Methodological Expectations of Cochrane Intervention Reviews (MECIR) guidance.

Study Design and Setting: We conducted a systematic scoping review. We searched multiple databases (e.g., MEDLINE, Embase), which were supplemented by grey literature searching. Methods were defined *a priori* in a published protocol.

**Results:** Out of 1,873 records, 90 publications were divided into four RR categories: formal evaluation (n = 14), development, which included four subcategories (n = 65), comparison (n = 2), and applying reporting guidelines/critical appraisal tools (n = 3), and a systematic review surrogate category (n = 6). Four formal evaluation studies were composite evaluations, including more than one shortcut simultaneously. The remaining 10 studies evaluated viable (e.g., including English-only publications) and unviable (e.g., single-reviewer screening) shortcuts, covering five key dimensions and five 'other' (e.g., involving stakeholders) considerations while conducting a review. Because of complexities around shortcuts evaluated, only a cursory mapping to MECIR criteria was possible.

Conclusion: Some methods shortcuts may be valid in the context of RRs, but limitations in the studies may limit their applicability. The results will serve to inform discussions within Cochrane regarding possible future implementation of RRs. © 2020 Elsevier Inc. All rights reserved.

Keywords: Rapid reviews; Methodology; Shortcuts; Formal evaluations; Abbreviated methods; Scoping review

### **Objective**

- Protocol:
  - Open Science Framework (https://osf.io/dekx6/)
- Objective: to conduct a scoping review of the literature
  - <u>assessing one or more method(s) applicable for undertaking rapid</u> <u>reviews (e.g., single reviewer screening vs. double reviewer</u> screening) or
  - <u>comparing the results of rapid reviews to those of systematic</u> <u>reviews (e.g., do conclusions change?) across all stages of conduct.</u>
- Abstract in special Supplement to the Cochrane Database of Systematic Reviews<sup>1</sup>

#### **Methods**

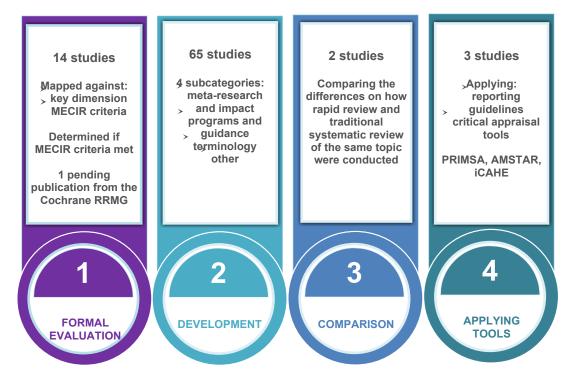
- Eligibility criteria
  - Published in English, since 1997
- Search for studies
  - MEDLINE<sup>®</sup> ALL, Embase Classic + Embase, PsycINFO, ERIC, Cochrane Library, CINAHL, Web of Science, Epistemonikos
  - Additional searching: grey literature (e.g., organizations that produce RRs), bibliographies of included studies, contacting experts in the field, bibliography of Robson 2018<sup>2</sup>
- Study selection
  - Piloting at title and abstract and full text screening
  - Liberal accelerated at title and abstract
  - Dual, independent at full text
- Data charting
  - Piloting performed on 5 records
  - One reviewer extracted, a second reviewer verified all data

#### **Methods**

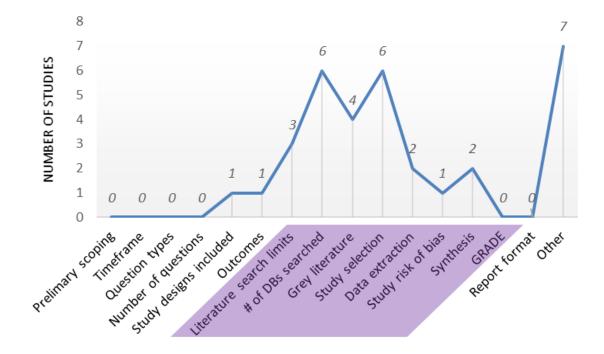
- Data synthesis
  - Two reviewers mapped the studies into 4 categories:
    - 1) Formal evaluation
      - mapped to stages of conduct to identify gaps
      - compared to Methodological Expectations of Cochrane Intervention Reviews (MECIR) guidelines to see whether the method met the MECIR criteria
    - 2) Development: meta-research and impact, programs and guidance, terminology, other
    - 3) Comparison (i.e., comparing RRs to SRs of the same topic)
    - 4) Applying tools (e.g., PRISMA, AMSTAR)
  - All studies narratively described and presented in tables

#### **Results**

- 90 studies, including 6 SR surrogates
  - 68 studies (75.6%) published since 2014
- 14 formal evaluation studies<sup>3-16</sup>
  - 10 evaluated single shortcuts
  - 4 evaluated 'composite' shortcuts
  - 11 studies (78.6%) published since 2017



#### Mapping to key dimensions of the review process



- Includes the composite evaluations studies
- A study could have evaluated more than one shortcut
- 33 total evaluations
- 16 single evaluations

#### **Shortcuts evaluated: 10 single-evaluation studies**

Review stage	Evaluation	Viable?
Literature search limits	• Marshall 2019: Excluding articles older than 5, 7, 10, 15, and 20 years before the search date	×
Number of databases	Marshall 2019: Removing any studies not identified in PubMed	√*
searched / Grey literature	<ul> <li>Nussbaumer-Streit 2019: Abbreviated searches, (i) combining a variety of database searches (ii) with or without gray literature searching</li> </ul>	<b>√/×</b>
	• Pham 2016: (i) including only the bibliographic database that yielded the highest number of records, plus the ancillary sources searched in the original SR/MA, and (ii) limiting the search to bibliographic databases	<b>√/×</b>
Screening	Gartlehner 2020: Single- reviewer screening	√*
	<ul> <li>Gartlehner 2019: Machine-assisted, screening, single-reviewer screening, and machine screening alone</li> </ul>	√*
	Pham 2016: Single-reviewer screening	×
	• Rathbone 2017: Participants, interventions and comparators-based title-only screening	$\checkmark$
Data extraction	Martyn St James 2017: Extracting data from an existing SR	$\checkmark$

\* Not for SRs, but may be viable for RRs or where synthesis is urgently needed.

#### **Shortcuts evaluated: 10 single-evaluation studies**

Review stage	Evaluation	Viable?
Involving stakeholders	Moore 2017: Including knowledge brokers in the review process	$\checkmark$
Inclusion based on study design	<ul> <li>Marshall 2019: Excluding trials with fewer than 50, 100, and 200 participants, and using the largest trial only</li> </ul>	×
Inclusion based on language	Nussbaumer-Streit 2020: Limiting to English-only publications	$\checkmark$
Inclusion based on access to publication	Pham 2016: Including studies that were available electronically	√*
Peer-review search strategy	• Spry 2018: Impact of the peer review of search strategies	✓

\* e-journals became more widely available in the 1990s

#### Discussion

- Little overlap in evaluations
  - Comparisons within a review stage of conduct differed
- Largely based on case studies
  - For example, Pham 2016 evaluated 4 different shortcuts using 3 SRs
- Composite evaluations
  - Recommend reporting the impact of each shortcut separately
- Recommendations may be topic dependent and impact of a shortcut should be considered
  - For example, if a topic was on nursing, then CINAHL should be searched





#### References

- Hamel C, Michaud A, Affengruber L, Skidmore B, Stevens A, Nussbaumer-Streit B, Garritty C. Rapid review methods: a systematic scoping review. In: Advances in Evidence Synthesis: special issue Cochrane Database of Systematic Reviews 2020;(9 Suppl 1):431. https://doi.org/10.1002/14651858.CD202001.
- 2. Robson RC, Pham B, Hwee J, Thomas SM, Rios P, Page MJ, Tricco AC. Few studies exist examining methods for selecting studies, abstracting data, and appraising quality in a systematic review. Journal of Clinical Epidemiology 2019; 106:121-135. Epub 2018 Oct 9.
- 3. CADTH Health Technology Inquiry Service. Transcatheter Aortic Valve Implantation: A Critical Appraisal of a Health Technology Assessment and Comparison with a Rapid Review. 2010.
- 4. Eiring Ø, Brurberg KG, Nytrøen K, Nylenna M. Rapid methods including network meta-analysis to produce evidence in clinical decision support: a decision analysis. Syst Rev. 2018 Oct 20;7(1):168.
- 5. Gartlehner G, Affengruber L, Titscher V, Noel-Storr A, Dooley G, Ballarini N, et al. Single-reviewer abstract screening missed 13 percent of relevant studies: a crowd-based, randomized controlled trial. J Clin Epidemiol. 2020 Jan 20;
- 6. Gartlehner G, Wagner G, Lux L, Affengruber L, Dobrescu A, Kaminski-Hartenthaler A, et al. Assessing the accuracy of machine-assisted abstract screening with DistillerAI: a user study. Syst Rev. 2019 Nov 15;8(1):277.
- 7. Kaltenthaler E, Tappenden P, Booth A, Akehurst R. Comparing methods for full versus single technology appraisal: A case study of docetaxel and paclitaxel for early breast cancer. Health Policy. 2008;87(3):389–400.
- 8. Marshall I, Marshall R, Wallace B, Brassey J, Thomas J. Rapid reviews may produce different results to systematic reviews: a metaepidemiological study. J Clin Epidemiol. 2019;109:30–41.
- 9. Martyn-St James M, Cooper K, Kaltenthaler E. Methods for a Rapid Systematic Review and Metaanalysis in Evaluating Selective Serotonin Reuptake Inhibitors for Premature Ejaculation. Evid Policy. 2017;13(3).
- 10. Moore G, Redman S, D'Este C, Makkar S, Turner T. Does knowledge brokering improve the quality of rapid review proposals? A before and after study. Syst Rev. 2017 28;6(1):23.
- 11. Nussbaumer-Streit B, Klerings I, Dobrescu AI, Persad E, Stevens A, Garritty C, et al. Excluding non-English publications from evidence-syntheses did not change conclusions: a meta-epidemiological study. J Clin Epidemiol. 2020 Feb;118:42–54.
- 12. Nussbaumer-Streit B, Klerings I, Wagner G, Heise TL, Dobrescu AI, Armijo-Olivo S, et al. Abbreviated literature searches were viable alternatives to comprehensive searches: a meta-epidemiological study. J Clin Epidemiol. 2018 Jun;102:1–11.
- 13. Pham MT, Waddell L, Rajić A, Sargeant JM, Papadopoulos A, McEwen SA. Implications of applying methodological shortcuts to expedite systematic reviews: three case studies using systematic reviews from agri-food public health. Res Synth Methods. 2016 Dec;7(4):433–46.
- 14. Rathbone J, Albarqouni L, Bakhit M, Beller E, Byambasuren O, Hoffmann T, et al. Expediting citation screening using PICo-based title-only screening for identifying studies in scoping searches and rapid reviews. Syst Rev. 2017 Nov 25;6(1):233.
- 15. Spry C, Mierzwinski-Urban M. The impact of the peer review of literature search strategies in support of rapid review reports. Res Synth Methods. 2018;9(4):521–6.
- 16. Taylor-Phillips S, Geppert J, Stinton C, Freeman K, Johnson S, Fraser H, et al. Comparison of a full systematic review versus rapid review approaches to assess a newborn screening test for tyrosinemia type 1. Res Synth Methods. 2017 Dec;8(4):475–84.

The Next Generation of Systematic Literature Review Platform

# **PICO Portal**

A new machine-learning powered tool to aid citation screening for evidence synthesis

> **Eitan Agai – PICO Portal Founder** eagai@PICOportal.org

Agai E. A New Machine-Learning Powered Tool to Aid Citation Screening for Evidence Synthesis: PICO Portal. In: Advances in Evidence Synthesis: special issue. Cochrane Database of Systematic Reviews 2020;(9 Suppl 1):172 https://doi.org/10.1002/14651858.CD202001



- How to build "trust" in machine learning?
- Where is machine learning is applied in PICO Portal?
  - 'Include' & 'exclude' prediction
  - Study type classification
  - Deduplication
  - Highlighting keywords
  - Crowd sourcing
- Q&A



### A Glance at PICO Portal

Highlights 💙	« Previous	6/15 Next »	PICO Legend	ds 🔻 Rese	t Yes Maybe	<u>₽</u> <u>!</u> <u>⊆</u> <u></u>
				Selec	ct an exclude reason	• e <u>X</u> clude
Title & Abstract				^		
J2017_611403981 Parents in transition: Experiences of parents	of young peo	ople with a liv	ver transplant transferring to adult services 🗗	_	ostract Review Consensus: <mark>Yes</mark> Hur <mark>d</mark> i	
Authors: Wright J., Elwell L., McDonagh J.E., Kelly D.A., Wra Published on: 2017 Publication: Pediatric Transplantation DOI: 10.1111/petr.12760 🗗	-			Article Non R	e Analysis: RCT	
healthcare management throughout childhood and adole adult services can be challenging for parents and young per transplant recipients (15.2-25.1 yr) participated in semistru- impact of transplantation ,"`` protection vs. independence desire to promote independence in their child while still m navigate . Parents are important facilitators of young peop supported to move from a``managerial"to a``supervisor	scence including sople, yet parents actured interview a ,"and``ending aintaining contro le's developmen y"role during trai nphasize their ro	encouraging dev s' views regarding s . Interviews we relationships and ol and protection t of self-manager nsition to help yc	hage and engage with healthcare services . Parents have a key role in velopment of self-management skills in their <b>children</b> . Transition to g transition remain largely unexplored . Nine parents of <b>pediatric</b> liver re analyzed using IPA . Analysis revealed three key themes: ``emotion d changing roles ."Parents expressed the dichotomous nature of the , and discussed how changing roles and relationships were difficult to ment skills for successful transfer to adult services . Parents should be <b>bung people</b> engage independently with the healthcare team . Finding nd guide the transfer of self-management skills from parent to young	al	Tags (5) Notes (0) PDF (0) Search Cond/Dx: Solid Organ Transplant Study Design: Qualitative studies Target Population: Patient Setting: Primary care	



### A Glance at PICO Portal

Highlights V 6/15 No	ext »	Reset     Yes     Maybe     P     I     C     Q       Select an exclude reason <ul> <li>exclude</li> </ul>
Title & Abstract	~	
	MANCHESTER 1824 The University of Manchester	Full Text Review Final Consensus: I Mage
	The University of Manchester Research	Article Analysis: Non RCT
Parents in transition: Experience         people with a liver transplant transplant transplant         DOI:         10.1111/petr.12760         Document Version         Accepted author manuscript         Link to publication record in Manchester Research Explorer         Citation for published version (APA):         Wright, J., Elwell, L., Mcdonagh, J., Kelly, D. A., & Wray, J. (20 young people with a liver transplant transferring to adult service		<ul> <li>Tags (5) Notes (0) PDF (0)</li> <li>Search</li> <li>Cond/Dx: Solid Organ Transplant</li> <li>Study Design: Qualitative studies</li> <li>Target Population: Patient</li> <li>Setting: Primary care</li> <li>Add new tag</li> </ul>



### Building Machine Learning and "Trust"

Machine learning = classification, prediction & clustering

Trust is built when the classification, prediction & data clustering helps you make good research decisions

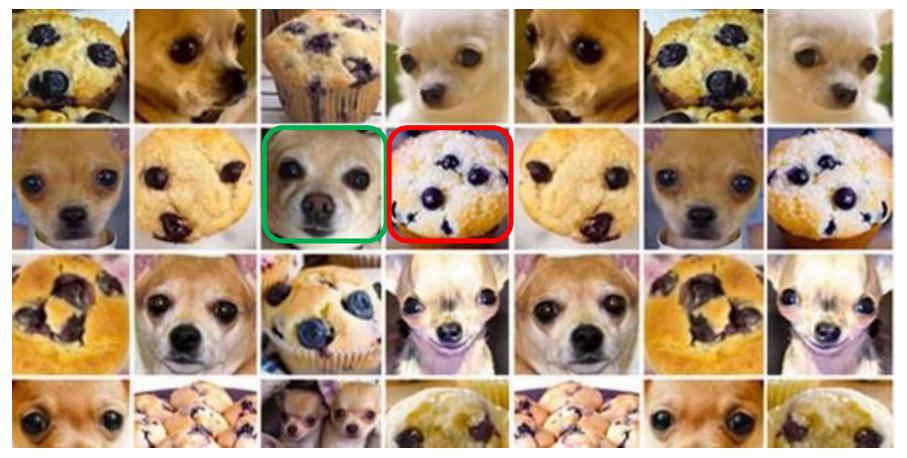
Based on these guiding principals, our approach is:

We **earn user's trust** during the project duration, and we also make sure that the techniques that PICO Portal is using are benchmarked against published research



### Predicting study results in systematic reviews is hard

If Chihuahua is "Include" & Muffin is "exclude", how do we teach a machine to solve that?



https://www.freecodecamp.org/news/chihuahua-or-muffin-my-search-for-the-best-computer-vision-api-cbda4d6b425d/



### Predicting study results in systematic reviews is hard

And in this case of Sheepdog or Mop?



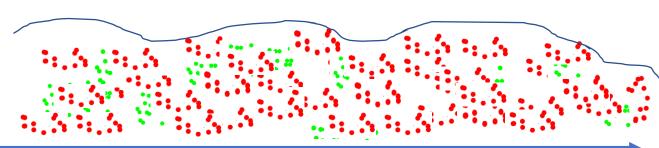
https://www.freecodecamp.org/news/chihuahua-or-muffin-my-search-for-the-best-computer-vision-api-cbda4d6b425d/



### **Traditional Screening**



#### **Traditional Screening**



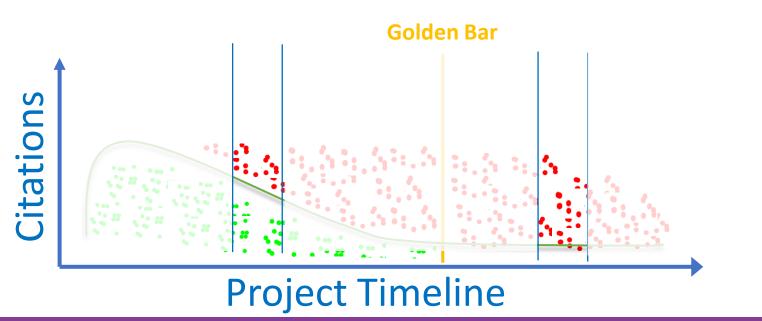
#### **Project Timeline**

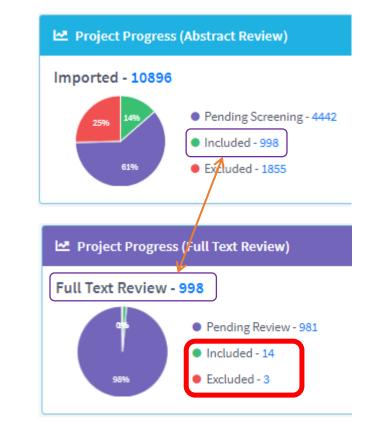
- Citation are screened in random order
- Resources need to be allocated manually
- In many cases the process is sequential and takes longer
- Open source or commercial tools resemble spreadsheets and aren't easy to use



## Screening Using Prediction (Machine Learning)

- Citations are sorted with most likely "include" first
- Users can confirm the model is working for that project
- At the golden bar moment, users can reduce the resources





Screening can immediately help refine the inclusion/ exclusion criteria

PICO

PORTAL

### Study Type Classification: Proceeding with Caution

#### Hall\_2018\_30387126

Caseworker-assigned discharge plans to prevent hospital readmission for acute exacerbations in children with chronic respiratory illness.

Authors: Hall Kerry K, Petsky Helen L, Chang Anne B, O'Grady KerryAnn F Published on: 02 Nov 2018 Publication: Cochrane Database of Systematic Reviews

#### DOI: 10.1002/14651858.CD012315.pub2 🗹

BACKGROUND: Chronic respiratory conditions are major causes of **mortality** and **morbidity**. Children with chronic health conditions have increased **morbidity** associated with their physical, emotional, and general wellbeing. Acute respiratory exacerbations (AREs) are common in children with chronic respiratory disease, often requiring admission to hospital. Reducing the frequency of AREs and recurrent hospitalisations is therefore an important goal in the individual and public health management of chronic respiratory illnesses in children. Discharge planning is used to decide what a person needs for **transition** from one level of care to another and is

Article Analysis: Non RCT Possible Meta-Analysis				
<ul> <li>Tags (0) Notes (0) PDF (0)</li> <li>Search</li> <li>Target Population: Patient</li> <li>Study Design: Program evaluation</li> <li>Target Population: Provider</li> <li>Study Design: Qualitative studies</li> </ul>				
Highlights	110/2889 Next »		PICO Legends ▼	Article Analysis:           Non RCT         Systematic Review
Voices not heard: a systematic revie perspectives of health care transiti Authors: Betz Cecily L, Lobo Marie L, Nehring V Published on: 2013 Sep-Oct Publication: Nur DOI: 10.1016/j.outlook.2013.01.008	on. [Review] ♂ Nendy M, Bui Kim	nd emerging a	dults'	<ul> <li>Tags (0) Notes (0) PDF (0)</li> <li>Search</li> <li>Target Population: Patient</li> <li>Study Design: Program evaluation</li> </ul>
BACKGROUND: A better understanding of the needs (AEA-SHCNs) is essential to provide hea purpose of this systematic review was to evalu	Ith care transition service	s that represent bes	st practices . The	Target Population: Provider Study Design: Qualitative studies



### Sophisticated Deduplication

#### 🗠 Project Progress (Abstract Review)

Imported - 11661

74%



#### PICO Portal deduplication is fast and accurate relative to other similar platforms.

#### Primary

Breakey\_2012\_CN-01008613

#### Usability testing of an online transition program for adolescents with hemophilia

Authors: Breakey VR Warias A Ignas DM Blanchette VS Stinson J Published on: 2012 Publication: Haemophilia Volume (Issue): 18 Journal: Journal DOI: 10.1111/j.1365-2516.2012.02778.x

Aim: To explore the usability of a new Internet-based educational program of Information, self-management strategies and social support for adolescents with hemophilia. Methods: An extensive educational website, "Terens Taking Charge: Managing Hemophilia Online" was developed based on results of an in-depth needs assessment. A purposive sample of adolescents was recruited from two tertiary care centers to as less online program in English and French. The website was tested for usability, using qualitative methods that included semi-structured interviand observation by a trained observer. Testing occurred iteratively, with changes to the prototype made after each cycle. Thematic analysis usin a collaborative and iterative process was used to organize data into categories that reflected the emerging themes. Results: Eighteen participants tested the website in three cycles (age range 12-18 years, mean 15.4 years). All had access to a computer at home and felt comfortable using the internet. Teens responded positively to the content, appearance and theme of the website. Overall, they felt that it was easy to navigate, use and understand. The multimedia components (videos, animations and quizzes) were felt to enrich the experience. Adolescents provided ideas on how the website user-interface could be improved. Minor changes to the website user-interface were made after the first and second cycles of testing in English. Cycle three was done in French and resulted in several additional changes. At the teens' suggestion, additional social media elements were added (discussion board, "ask the expert" section) to increase interactivity. Most participants felt this program would be helpful prior to transition of care and beyond. Conclusions: Usability testing was the crucial first step in ensuring the acceptability and ease of use of this internetbased self-management program. A pilot study is currently underway to determine the feasibility of a randomized controlled trial to assess the online tool.

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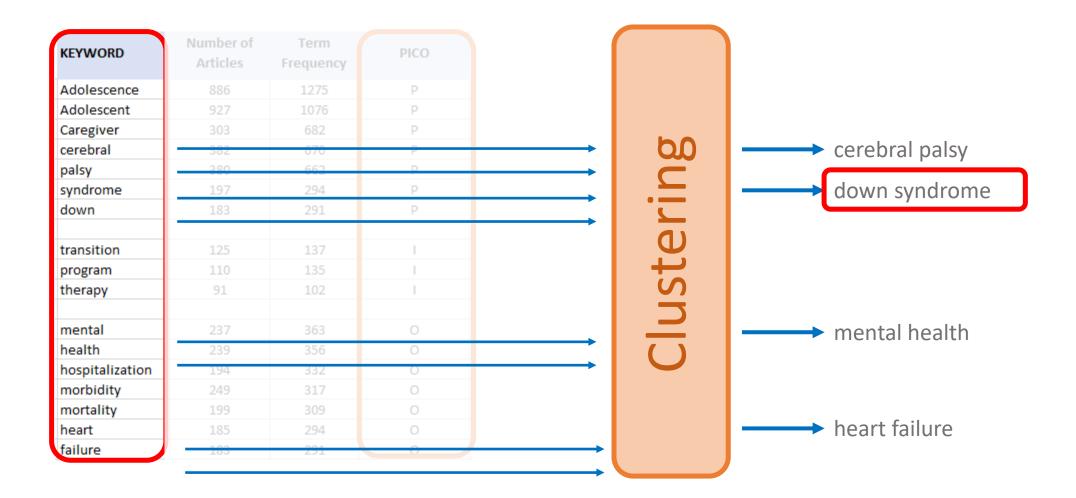
#### Usability testing of an online transition program for adolescents with hemophilia

Authors: Breakey VR, Warias A, Ignas DM, Blanchette VS, Stinson J Published on: 2011 Publication: Blood Volume (Issue): 118(21) Journal: Journal DOI: -

ose: This study explored the usability of a new Internet-based educational program of disease-specific information, self-management strategies and social support for adolescents with hemophilia. Methods: A comprehensive eight-module educational website was developed, sults of an in-depth needs assessment. The website was tested for usability, using qualitative methods that included semi-structured, terviews and observation by a trained observer. To determine the usability and intuitiveness of the user interface of the "Teens Managing Hemophilia Online" intervention, testing occurred in three cycles (4 participants per cycle). Participants were asked to standardized parts/features of the program, with changes to the prototype made after each cycle. Thematic analysis using a nd iterative process was used to organize data into categories that reflected the emerging themes. Results: A purposive sample of adolescents (range 12-18 years, mean 15.4 years) was recruited from a Canadian tertiary care center. All of the participants had access to a computer at home and felt comfortable using the Internet. Teens responded positively to the content, appearance and theme of the website (promoting self-management in youth with hemophilia). Subjects thought that it was easy to navigate, use and understand. Overall, they felt the content was appropriate and geared to meet the unique needs of adolescents with hemophilia. The multi-media components (videos, animations and quizzes) were thought to enrich the experience and make the program appealing. Adolescents provided ideas on how the website userinterface could be improved in terms of its usability (navigation, format and layout). Minor changes to the website user-interface were made and tested after the first and second cycles of testing. No further problems were identified in final cycle of testing. At the teens' suggestion, additional social media elements were added (discussion board, "ask the expert" section) to build in elements of support and increase interactivity. Most participants felt this program would be helpful prior to transition of care and beyond. Conclusions: Usability testing was the crucial first step in ensuring the acceptability and ease of use of "Teens Taking Charge: Managing Hemophilia Online". Findings from this study were used to refine the website prototype. A pilot study is underway to determine the feasibility of using a randomized controlled trial to study the Internet-based selfmanagement program.



# Keyword Highlighting



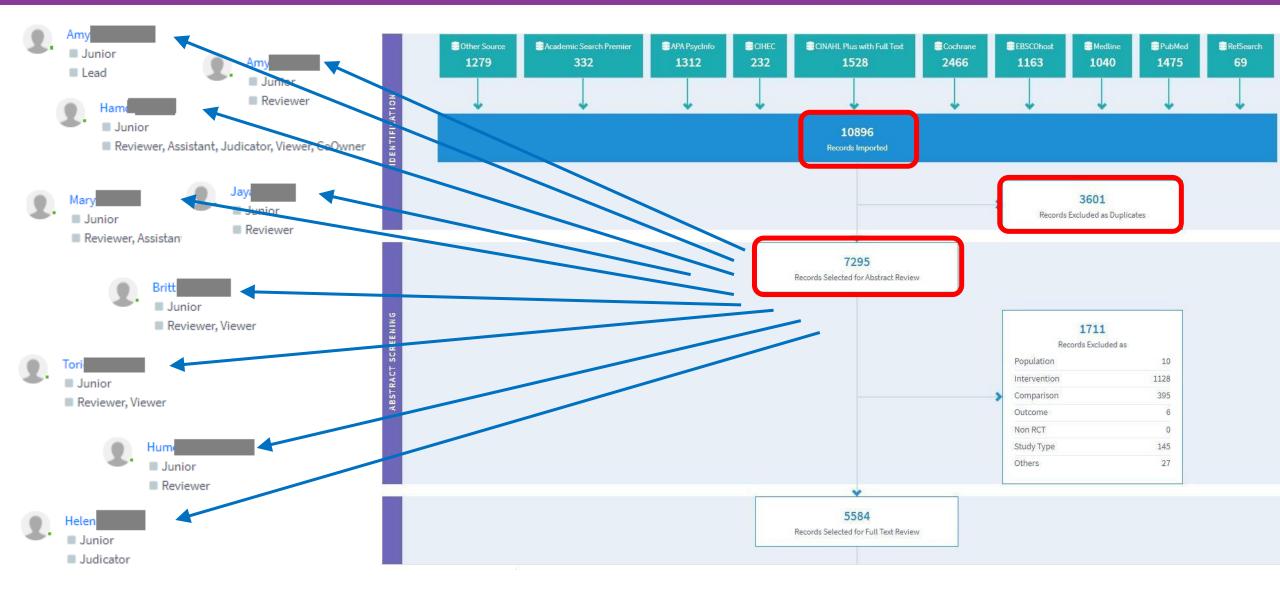


# Keyword Highlighting

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Transition to adulthood for young people with intellectual disab	a <mark>ility</mark> : the experiences of their families 🖻			Abstract Review	
Authors: Leonard H., Foley KR., Pikora T., Bourke J., Wong K., McPherson L., Lennox Published on: 2016 Publication: European Child and Adolescent Psychiatry;25(12):1				Reviewer 1: - Final Consensus: -	
DOI: 10.1007/s00787-016-0853-2 + Add Full Text PDF					
				Article Analysis: Non RCT	
Whilst the transition from school to adult roles can be challenging for any <b>adolescent</b> process may involve coordinated planning, collaboration and decision-making amon with <b>Down syndrome</b> in Western Australia (n = 190) and <b>young people</b> with <b>intellectu</b> comprised two parts: part - collected information about the individual with <b>intellectu</b> collected information about the health and well-being of their family. The majority ( <i>R</i> <b>people</b> were involved in this process . The three most helpful strategies indicated by p transition program and the building of informal community-based supports . A numb to adapt and change to life in adulthood, their difficulty navigating services and prog- about the longer term future. Copyright © 2018, Springer-Verlag Berlin Heidelberg.	ng school staff, families and community agencies . This mixed-methods study ut ual disability (of any cause) in Queensland, Australia (n = 150). The parent-repor tual disability including information on health, functioning and service needs, a (87 %) of parents said that they were involved in decision-making about transiti parents that assisted with transition planning related to the provision of more i ber of themes emerged from the qualitative data which included parents' views grams, issues and challenges around their young person building connectednes	tilised information from two co rt questionnaires administered and about specific transition re ion planning but less than two- information about financial as: and concerns about the capac ss, strain on family wellbeing a	ohorts: young people d in both states elated issues; and part 2 -thirds (59.5 %) of young sistance, the school city of their young adult nd finances and worry	<ul> <li>Tags (0) Notes (0) PDF (0)</li> <li>Search</li> <li>Cond/Dx: ACEs</li> <li>Cond/Dx: Acquired Brain Injury</li> <li>Cond/Dx: ADHD</li> <li>Cond/Dx: Anorectal Malformations (ARMs)</li> <li>Add new tag</li> </ul>	
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Agai E. A New Machine-Learning Powered Tool to Aid Citation Screening for Evidence Synthesis: PICO Portal. In: Advances in Evidence Synthesis: special issue. Cochrane Database of Systematic Reviews 2020;(9 Suppl 1):172 https://doi.org/10.1002/14651858.CD202001

