

# Conducting a Literature for a Systematic Review

2011 SMP Research Skills Program



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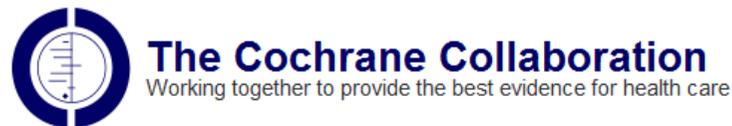
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- › We wish to extend our thanks to the following staff for sharing their time, research and expertise in developing this program.
- › Margot Day, Sub-Dean USydMP Honours Post-graduate Co-ordinator
- › Pauline Kennedy, Medical Program Administration Unit
  
- › Asbestos Disease Research Institute, Bernie Banton Centre



- › Cochrane Collaboration



› You will learn to:

- Identify the differences between the different types of review article.
- Know where literature searching falls in the systematic review process.
- Identify the databases and resources required for a comprehensive review search.
- Construct a search strategy for use in a systematic review.
- Search databases using a combination of subject headings and keyword search techniques.
- Document and report a search strategy.

Systematic  
review  
Narrative  
Overview  
Meta-analysis  
Literature  
Literature

What's the difference?

Image generated from Wordle <http://www.wordle.net/>

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## Literature Review

- › Also known as “Narrative’ or ‘Overviews’
- › Section of a dissertation/thesis or a paper in their own right.
- › gives an overview and provides comprehensive knowledge of what has been written on a particular topic.
- › Conveys...what has been said, who the key writer’s are, what are the prevailing theories and hypotheses, what questions are being asked.
- › summarises and synthesises the arguments and ideas of others and highlights the strengths and weaknesses of these ideas and arguments
- › provides a solid background for a research paper’s investigation.

## Systematic Reviews

“attempts to collate all empirical evidence that fits pre-specified eligibility criteria to answer a specific research question”

### **Key characteristics:**

- Clearly stated set of objectives with pre-defined eligibility criteria for studies;
- An explicit reproducible methodology;
- A systematic search that attempts to identify all studies that meet the eligibility criteria;
- An assessment of the validity of the findings of the studies, through the assessment of risk of bias;
- A systematic presentation and synthesis of the characteristics and findings of the included studies;
- May or may not include a meta-analysis.



# What is a systematic review?

## Radiotherapy for malignant pleural mesothelioma (Review)

Chapman E, García Diéguez M



This is a reprint of a Cochrane review, prepared and maintained by The Cochrane Collaboration and published in *The Cochrane Library* 2010, Issue 4

<http://www.thecochranelibrary.com>



Radiotherapy for malignant pleural mesothelioma (Review)  
Copyright © 2010 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

## Differences between Narrative and Systematic Reviews

- › Traditional reviews are prone to bias. The author may:
  - Only choose studies which they are familiar with or support their view-point [selection bias]
  - Only select studies which have been published [publication bias]
  - Only select studies written in English [language bias]
  - Only search a limited number of databases or use a limited choice of search terms [indexing bias]

# What is a systematic review?

## What is a meta-analysis?

- › A research review method that combines the evidence of multiple primary studies by employing statistical methods, enhancing the objectivity and validity of findings.
  - › The research design and hypotheses of primary studies need to be very similar if not identical.
  - › Not always appropriate for use in a systematic review.
-

## Systematic Literature Review

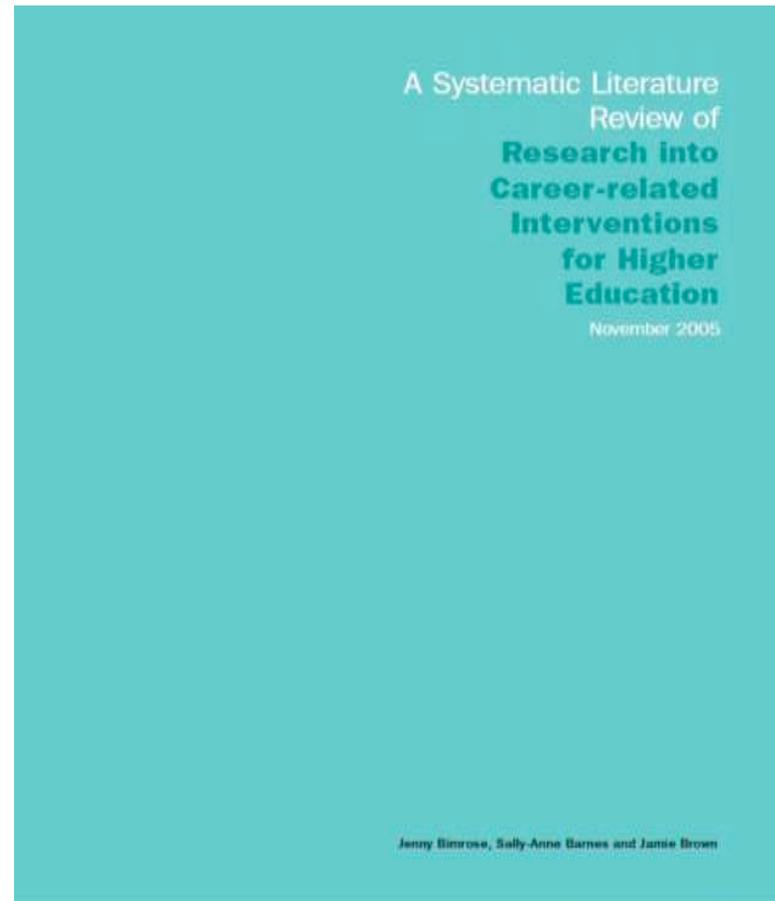
- › A systematic literature review is a review where all procedures are documented – the research trail of databases and search terms used is made explicit.
  - ...involves adopting a "replicable, scientific and transparent" approach and providing an "audit trail of reviewers' decisions, procedures and conclusions.
  - Based upon work by down by researches in the medical and health sciences but expanded to other disciplines such as education and management.

## Systematic Literature Reviews

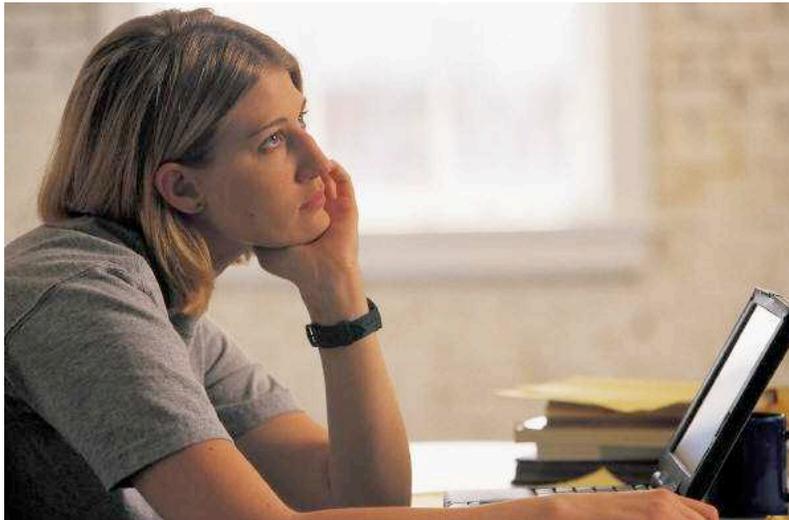
- › Systematic literature reviews include the following
  - The type of literature searched, and excluded (e.g. articles in particular types of journals, but not conference papers, dissertations, textbooks) giving the reason.
  - The method of classification – for example, years of publication, theme.
  - Databases searched, search and thesaurus terms used.
  - Any other type of searching used, for example hand searching.
  - Limiting criteria, for example dates, geographical coverage, type of publication etc
  - Tracking method for citations.
  - What the criteria were for inclusion and exclusion in the final review.
  - How the papers were analysed.

# What is a systematic review?

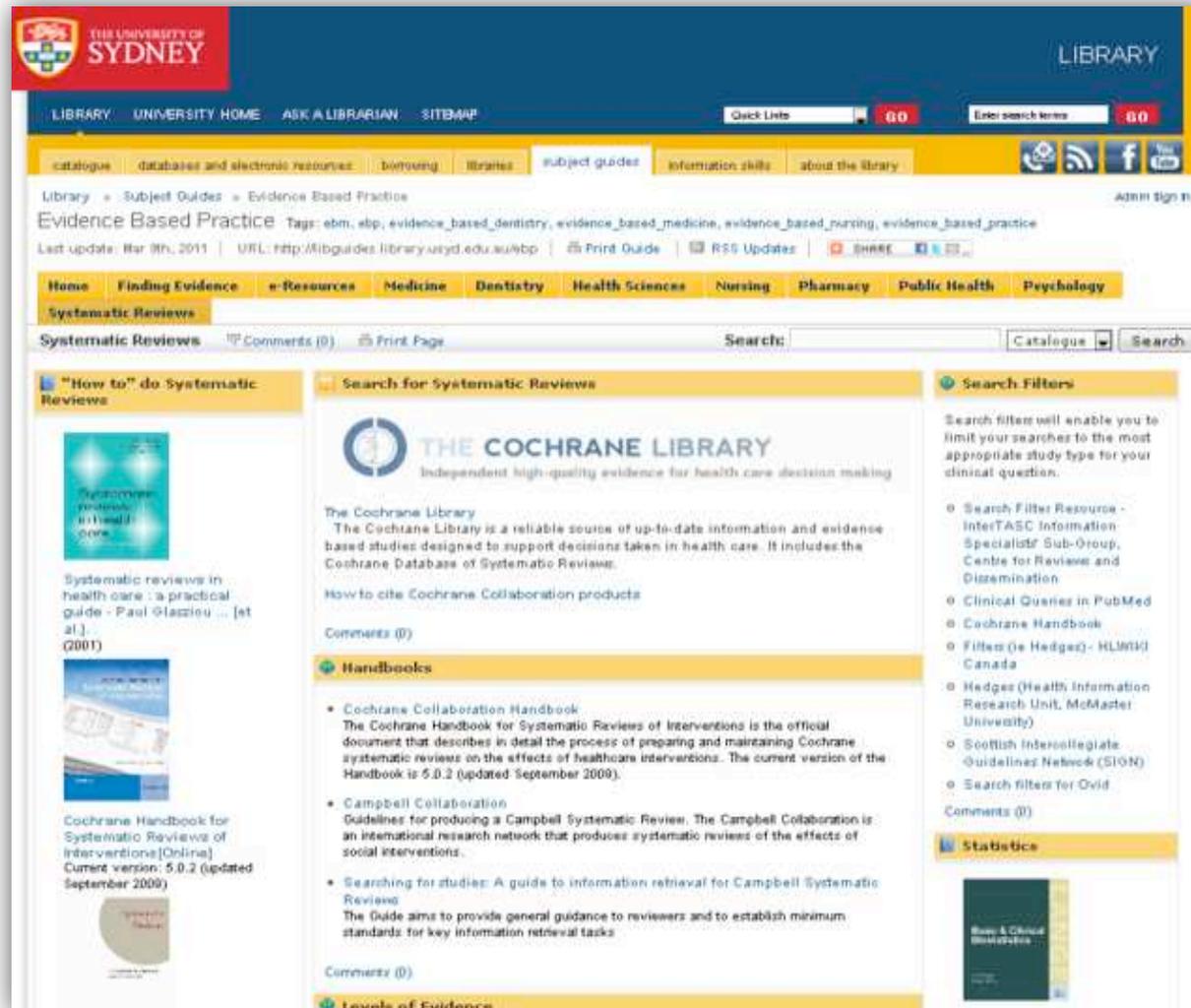
## A Systematic Literature Review



You've decided to do a systematic review – where do you begin?



# Where can I find info on systematic reviews?



The screenshot displays the University of Sydney Library website. At the top, there is a navigation bar with links for 'LIBRARY', 'UNIVERSITY HOME', 'ASK A LIBRARIAN', and 'SITEMAP'. A search bar is located on the right side of the navigation bar. Below the navigation bar, there is a secondary navigation bar with links for 'catalogue', 'databases and electronic resources', 'borrowing', 'libraries', 'subject guides', 'information skills', and 'about the library'. The main content area is titled 'Evidence Based Practice' and includes a search bar, a 'Comments (0)' link, and a 'Print Page' link. The page is divided into three main sections: 'How to do Systematic Reviews', 'Search for Systematic Reviews', and 'Search Filters'. The 'How to do Systematic Reviews' section features a book cover for 'Systematic reviews in health care: a practical guide' by Paul Glasziou et al. (2001) and a link to the 'Cochrane Handbook for Systematic Reviews of Interventions [Online]'. The 'Search for Systematic Reviews' section features the Cochrane Library logo and a description of the library as a reliable source of up-to-date information and evidence-based studies. The 'Search Filters' section provides a list of search filters to refine searches, including 'Search Filter Resource - InterTASC Information Specialists' Sub-Group', 'Clinical Queries in PubMed', 'Cochrane Handbook', 'Filters (ie Hedges) - KUMC/Canada', 'Hedges (Health Information Research Unit, McMaster University)', 'Scottish Intercollegiate Guidelines Network (SIGN)', and 'Search filters for Ovid'. The page also includes a 'Statistics' section with a book cover for 'Basic & Clinical Biostatistics'.

# Where can I find resources on systematic reviews?

Investigate Cochrane Collaboration resources

› Australasian Cochrane Centre

- Individual Cochrane Review Groups:

- Renal Group (Children's Hospital at Westmead):

<http://www.cochrane-renal.org/>

- Public Health (Melbourne): <http://ph.cochrane.org/welcome>

› Campbell Collaboration (Education, Crime & Justice, Social Welfare)

- <http://www.campbellcollaboration.org/>

## Process of undertaking a Systematic Review

1. Question formulation & protocol development.
2. Finding studies: searching the literature.
3. Study selection and appraisal.
4. Summary and synthesis of relevant studies.
5. Determining the applicability of results.

Advisable to develop a protocol outlining the question and proposed methodology before proceeding with the review

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## How much work is a systematic review?

A 1999 analysis of 37 meta-analyses showed that the average hours for a review was 1139 (median 1110) or about 30 person weeks of full-time work.

Breakdown:

- › 588 hours protocol development, searching & retrieval.
- › 144 hours statistical analysis.
- › 206 hours writing.
- › 201 hours administration.

# How long does it take to do a review?

Depends on the resources available:

Cochrane Reviews:

- › Title to Protocol stage: approximately 3 months
- › Protocol to Review: 6-12 months
- › Updates at least every 2 years.

## How many people does it take to do a review?

A systematic review is best done by a team of people rather than one person, consisting of:

- Someone to write the review
  - Content or subject experts;
  - A methodologist, or someone experienced in the methods of a conducting a review;
  - A statistician;
  - Someone experienced in literature searching;
- › Two people to do the:
- Study selection and appraisal.
  - Data collection and analysis

## Question Formulation

- Need to develop a relevant and answerable question – this can take some time.
  - Has a review been published on the topic – do a scoping search!
-

## Process of undertaking a systematic review

Let's think about your review question

How broadly or narrowly focused should it be?

A narrow review question:

Easy for reviewer but less useful for the reader.

A broad review question :

More difficult for the reviewer but more useful for the reader.

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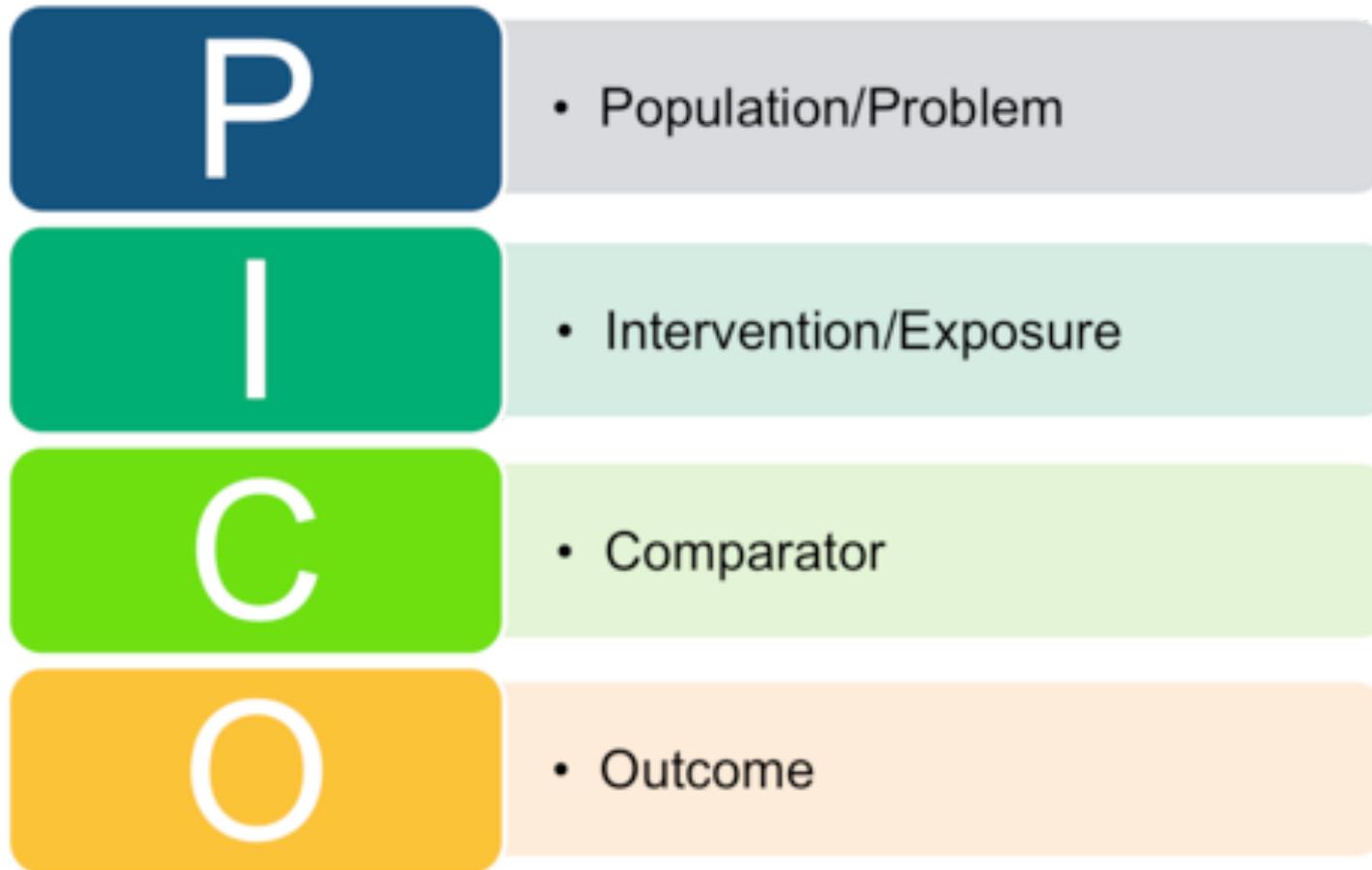
› Broad Review Questions:

- Surgical interventions for patients with malignant mesothelioma
- Incentive-based interventions to increase physical activity

Narrow Review Questions

- Extra-corporeal shock wave lithotripsy versus ureteroscopic management for ureteri calculi in middle-aged men
- Antibiotic prophylaxis to reduce respiratory tract infections and mortality in adults receiving intensive care

# Your systematic review question in PICO



# Identifying the Key Concepts using PICO

## Surgical interventions for patients with malignant mesothelioma

	Question	Search Terms
<b>P</b>	Malignant mesothelioma patients	Malignant mesothelioma
<b>I</b>	Surgical interventions	Pneumonectomy, Extrapleural pneumonectomy, pleurectomy, decortication,
<b>C</b>	Surgical interventions	
<b>O</b>	Quality of life Survival Improvement in symptoms	
<b>Question Type: Intervention / Therapy</b>		
<b>Best Study Design: Meta-Analysis / Systematic Review &gt; Randomized Controlled Trial</b>		

## Resources to Help you with Brainstorming your question

- › Literature you have already read
- › Subject experts
- › Drug information
  - eTG: Treatment guidelines for Australia (what drugs are used?)
  - AusDI / MIMS: Class, Generic and Brand Names
- › Synonyms
  - Dictionaries
  - Encyclopaedias
  - eMedicine (synonym tool)
  - MediLexicon for acronyms
  - Plurals, truncation, phrasing
- › Spelling: US / English
- › Prognostic Indicators
  - Think of every possible outcome (big picture to the minutiae)

# What Type of Clinical Question Are You Asking?

## Therapy / Intervention

- To determine the most appropriate treatment for a particular condition (drug treatment, surgical procedure, other)

## Diagnosis

- Questions the validity and reliability of a particular diagnostic test or screening method

## Prognosis

- To find likely outcomes for patients where their illness is identified early

## Aetiology / Harm

- Is there a relationship between exposure to a potentially harmful agent and developing a particular condition

# What study design fits my question?

**Table 2** Appropriate study designs for addressing different clinical questions

Question type	Generic examples	Best design†	Other possible designs
Intervention	<ul style="list-style-type: none"> <li>• Drug vs control</li> <li>• Drug vs drug</li> <li>• Procedure vs control</li> <li>• Decision support system vs usual practice</li> </ul>	Randomized controlled trial	Cohort study Case-control study
Diagnostic accuracy	<ul style="list-style-type: none"> <li>• Test A vs reference test</li> <li>• Test A vs Test B vs reference</li> </ul>	Cross-sectional study	–
Aetiology	<ul style="list-style-type: none"> <li>• Exposure present vs absent</li> </ul>	Cohort study	Case-control study
Prognosis	<ul style="list-style-type: none"> <li>• Exposure present vs absent</li> </ul>	Cohort study	–

†In all cases, a systematic review of all available studies of the best type is preferable to single studies.

Cross, Nicholas B., Craig, Jonathan C., Webster, Angela C. Asking the right question and finding the right answers. *Nephrology* 2010;15(1):8-11.

- › A scoping search is designed to give an overview of the published literature on your research question.
- › Can be done quite quickly using a simple search strategy.
- › Use it to find out if any systematic reviews have already been published, and to get an idea of how much literature is available.
  - Use Database limits to find systematic reviews or RCT's etc.

Once you have your question you should prepare a protocol

A protocol is:

A pre-defined plan or methodology that outlines the approach to a systematic review.

## What does a protocol look like?

Title*
<b>Protocol information:</b>
Authors*
Contact person*
Dates
What's new
History
<b>The protocol:</b>
Background*
Objectives*
Methods:
Criteria for selecting studies for this review:
Types of studies*
Types of participants*
Types of interventions*
Types of outcome measures*
Search methods for identification of studies*
Data collection and analysis*
Acknowledgements
References:
Other references:
Additional references
Other published versions of this review
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Contributions of authors
Declarations of interest*
Sources of support:
Internal sources
External sources
Published notes

Green S, Higgins JPT (editors). Chapter 2: Preparing a Cochrane review. In: Higgins JPT, Green S (editors). *Cochrane Handbook for Systematic Reviews of Interventions* Version 5.0.1 (updated September 2008). The Cochrane Collaboration, 2008. Available from [www.cochrane-handbook.org](http://www.cochrane-handbook.org).



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## Mycophenolic acid versus azathioprine as primary immunosuppression for kidney transplant recipients (Protocol)

Wagner M, Balk EM, Webster AC, Raman G, Trikalinos TA, Schmid CH, Uhlig K



**THE COCHRANE  
COLLABORATION**<sup>®</sup>

This is a reprint of a Cochrane protocol, prepared and maintained by The Cochrane Collaboration and published in *The Cochrane Library* 2009, Issue 2

<http://www.thecochranelibrary.com>

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Publishers Since 1807

Mycophenolic acid versus azathioprine as primary immunosuppression for kidney transplant recipients (Protocol)  
Copyright © 2009 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

# What makes a great systematic review literature search?

It's systematic:

- › Focused question
  - › Comprehensive search of all resources
  - › Free of bias
-

## › Be **Sensitive**

- use multiple sources and try to find all the available literature

## › Be **efficient**

- start where you expect highest yield, keep your search within reason

## › Minimise **bias**

- don't limit your search by language, year, publication status



## Structure of a search strategy

- › The strategy should be based on your eligibility criteria
- › Use the 2 or 3 most important concepts in your question
- › Use those most likely to be found in the title and the abstract of an article.
- › Use PICO

- › Aim for high sensitivity
- › Use synonyms – express your concepts as in many ways as you can.
- › Use the controlled vocabulary of each database you search.
- › Use textwords in addition to your subject headings.
- › Test and revise your strategy for each database you search – this may lead to more results.

# Creating a search strategy

Initial Search Strategy for all databases	
Patient Population	Intervention
malignant mesothelioma [as subject heading]	Pneumonectomy [as subject heading]
malignant mesothelioma [as textword]	Pneumonectomy [as textword]
	Extrapleural pneumonectomy [as subject heading]
	Extrapleural pneumonectomy [as textword]
	EPP [as textword]
	pleurectomy [as subject heading]
	pleurectomy [as keyword]
	decortication [as subject heading]
	decortication [as textword]

## Using a Controlled Vocabulary

- › Use the subject terms (known as the controlled vocabulary) for each database you search.
- › Will find relevant articles even if different terms are used for each concept.
- › Use the database interface to find the best subject heading – they will guide you through the process.
- › For databases such as Embase and Medline, don't automatically explode – take care in selecting your subject headings.
- › Subject headings may be different for each database.
- › Indexers are human, therefore you need to use textwords.

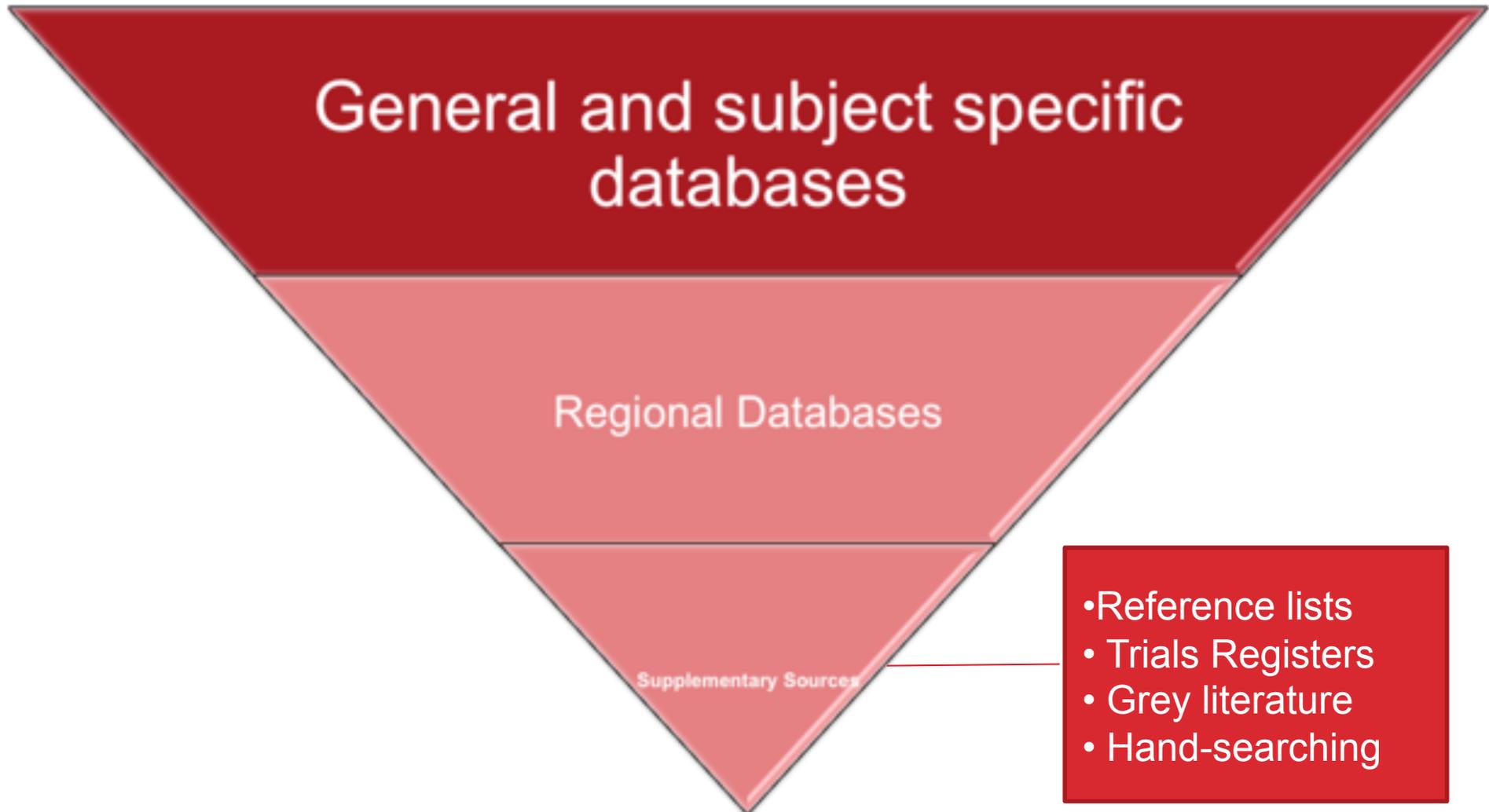
## Using Textwords

- › These are words appearing in the title and abstract of a database record.
- › In Medline add the field suffix .tw to your word:
  - For example: pleurectomy.tw
- › Allows you to use synonyms, related terms, alternative spelling, abbreviations, acronyms:
  - For example: lung cancer, lung neoplasms
- › Use truncation and wildcards - \* \$ ?
  - For example: transplant\* will retrieve transplants, transplantation
- › Proximity indicators – ADJ, NEAR, NEXT
  - For example: malignant ADJ3 meothelioma\*.tw will retrieve malignant mesothelioma, malignant pleural mesothelioma etc.
- › Syntax may be different for each database you search. Use the help for each database.

## What study design filter should I use?

- › A search strategy designed to limit your results to specific study types.
  - For example, randomised controlled trials
- › Research by different organisations has been done to identify the most sensitive and efficient search terms
- › Select your filter:
  - Database and interface to be searched
  - Study design required for your review.
- › You can use:
  - Built in database filters
  - Filters developed by Cochrane or Health Information Research Unit at McMaster University etc.
- › Need to document what you have used

Be Comprehensive – but efficient



Where do you look for studies?

**Start where you expect to retrieve the most evidence**

- › General Medical databases
  - Medline
  - Embase
  - CENTRAL
- › Subject Specific databases:
  - Cinahl - Nursing
  - ERIC - Education
  - PsycInfo - Psychology

Ask your Faculty Liaison Librarian for advice

<http://www.library.usyd.edu.au/contacts/subjectcontacts.html>

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One of the major medical databases containing:

- › Over 18 million **citations** from more than 5,500 international journals
- › Indexes content published in over 39 languages
- › Primarily scholarly content as selected by the US National Institutes of Health Literature Selection Technical Review Committee
- › Uses the Medical Subject Headings (MeSH)
- › 1948 – Present
- › Strong North American focus (~52% of journals are of US origin)

See the Fact Sheets a@

<http://www.nlm.nih.gov/pubs/factsheets/medline.html>

Biomedical database with a major strength in pharmacological / drug :

- › 1966 to present
- › Incorporates Medline
- › 24 million records
- › Indexes 7,500 peer-reviewed journals in 30 languages
- › Includes 2000 journals not indexed in Medline
- › Over 260,000 conference abstracts
- › Extensive citations to drug/pharmaceutical research
- › EMTREE thesaurus
- › Search interfaces and limits specific to drugs and diseases
- › Medline overlap on average 34% but can range from 10-70% depending on the topic.

## Cochrane Central Register of Controlled Trials

- Available via OvidSP and The Cochrane Library.
- Includes Randomised Controlled Trials & Controlled Clinical Trials.
- 625,000 records from:
  - Medline (360,000)
  - Embase (95,000)
  - Other (175,000)
    - Cochrane Review Group specialised registers (unpublished records, non-randomised studies, on-going studies.
    - Hand-searched records from 3000+ journals and conference proceedings

- › LILACS (Latin American and Caribbean Health)
  - <http://bases.bireme.br/cgi-bin/wxislind.exe/iah/online/?IsisScript=iah/iah.xis&base=LILACS&lang=i&form=F>
- › AIM - African Index Medicus
  - <http://indexmedicus.afro.who.int/>
- › IMSEAR – Index Medicus for the South East Asian Region:
  - <http://imsear.hellis.org/>
- › Health Collection via Informit (Australian)
  - Available through University of Sydney Library Databases list

See Chapter 6: Cochrane Handbook for Systematic Reviews of Interventions(2009) for a complete list <http://www.cochrane-handbook.org>

## LILACS

- › LILACS (from the Latin American and Caribbean Health Science Information Center)
- › Uses Medical Subject headings (MeSH)
- › 1982 to present
- › Articles from 19 Latin and Caribbean countries
- › 825 journals (~40 journals overlap Medline & Embase)
- › 544,157 records
  - 437,400 articles
  - 75,087 monographs
  - 25,543 theses
- › 169,513 full-text records



# What Trials Registers are available?

- › United States
  - <http://clinicaltrials.gov>
- › Australia/New Zealand (ANZCTR)
  - <http://www.anzctr.org.au>
- › International
  - <http://www.who.int/ictrp>
- › Individual pharmaceutical company registers

## What's in Clinicaltrials.gov?

- › Up to date database for locating both US govt and privately funded clinical trials.
- › 104,582 clinical trials
- › Contains:
  - Summary of the purpose of the trial
  - Recruiting status
  - Criteria for participation
  - Location of the trial and contact information
  - Research design
  - Phase of the trial
  - Disease/condition & drug/therapy under study
- › Study Results new feature – Only a few studies have results available online

- › Conference papers, dissertations, government reports, proceedings, technical reports, research reports, in-house journals.
- › Hard to track down, much un-published.
- › OpenSIGL – <http://opensigl.inist.fr>
- › Grey-Matters: A practical tool for evidence based searching
  - [http://cadth.ca/media/pdf/Grey-Matters\\_A-Practical-Search-Tool-for-Evidence-Based-Medicine.doc](http://cadth.ca/media/pdf/Grey-Matters_A-Practical-Search-Tool-for-Evidence-Based-Medicine.doc)

- › Reference lists of relevant articles
- › References from existing systematic reviews in similar topic areas
- › Consultation with subject experts
- › Hand-searching
  - Not all journals are indexed in databases
  - Articles may be indexed incorrectly.
  - Journals may be indexed selectively
  - It may pay to select a 'high yield' journal in your subject area and search it.
  - Organisations such as Cochrane hand-search in their subject areas.

# How do I document my search strategies?

For each database:

Need to include:

- Name of database
- name of database platform (optional)
- years covered by search
- one or two sentence summary of the search strategy, including information on what filter you used.
- complete search strategy
- language restrictions (if any)
- › Any other sources of studies

TIP: Copy and paste search strategies from the database saved searches – reduces risk of error!

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## Medline via OvidSP

Medline via OvidSP (1948 to most recent) using the optimally sensitive search strategy developed for the Cochrane Collaboration for the identification of RCT's (Lefebvre, 2008):

1. Mesothelioma/
  2. (malignant adj3 mesothelioma\*).tw.
  3. or/1-2
  4. Pneumonectomy/
  5. Pneumonectom\*.tw.
  6. extrapleural pneumonectom\*.tw.
  7. epp.tw.
  8. pleurectom\*.tw.
  9. decortication.tw.
  10. or/4-9
  11. and/3,10
-

## CENTRAL via OvidSP

The Cochrane Central Register of Controlled Trials (CENTRAL) via OvidSP (Issue 1, 4 Jan 2011) will be searched using the following terms:

1. Mesothelioma/
  2. (malignant adj3 mesothelioma\*).tw.
  3. or/1-2
  4. Pneumonectomy/
  5. Pneumonectom\*.tw.
  6. extrapleural pneumonectom\*.tw.
  7. epp.tw.
  8. pleurectom\*.tw.
  9. decortication.tw.
  10. or/4-9
  11. and/3,10
-

# How should I document my search strategies?

## Embase.com

Embase.com (1966 to the most recent) was searched using a filter based on research by the UK Cochrane Centre “What is in The Cochrane Central Register of Controlled Trials (CENTRAL) from EMBASE? (Lefebvre 2008).” From: Cochrane Handbook, chapter 6.3.2.2. ], notation as a search string by Ket (Jan. 2009) <http://www.ubvu.vu.nl/pub/index.cfm?objecttype=page&objectid=387&lang=>

#4 #1 AND #2 AND #3

#3 random\* OR factorial\* OR crossover\* OR cross AND over\* OR placebo\* OR (doubl\* AND blind\*) OR (singl\* AND blind\*) OR assign\* OR allocat\* OR volunteer\* OR 'crossover procedure'/exp OR 'double blind procedure'/exp OR 'randomized controlled trial'/exp OR 'single blind procedure'/exp AND [embase]/lim

#2 'lung resection'/de AND [embase]/lim OR ('pleurectomy'/de AND [embase]/lim) OR pneumonectom\*:ab,ti AND [embase]/lim OR (pleurectom\*:ab,ti AND [embase]/lim) OR epp:ab,ti OR ('decortication' AND [embase]/lim) OR ('decortication':ab,ti AND [embase]/lim) OR 'extrapleural pneumonectomy'

#1 'malignant mesothelioma'/de AND [embase]/lim OR ('mesothelioma'/de AND [embase]/lim) OR (malignant NEXT/3 mesothelioma\* AND [embase]/lim)

---

# How should I document my search strategies?

## LILACS

LILACS (1982 to most recent) was searched using the following search strategy

```
((([MH]"MESOTHELIOMA") or ([TI] (mesothelioma$) or ([AB] (mesothelioma$)))) and  
((( [MH]"PNEUMONECTOMY" or ([TI] (pneumonectom$ or pleurectom$ or epp or decortication or  
extra and pleural and pneumonectom$) or ([AB] (pneumonectom$ or pleurectom$ or epp or  
decortication or extra and pleural and pneumonectom$))))))
```

# How should I document my search strategies?

## Any other searching done?

Insert another section into your Search Methods for Identification of studies,

Detailing:

- Details of any hand-searching done – for example, journals searched.
  - Any other means of identifying studies – For example, a statement of experts contacted.
- › Have a look at a Cochrane review.

- Identify the differences between the different types of review article.
- Know where literature searching falls in the systematic review process.
- Identify the databases and resources required for a comprehensive review search.
- Construct a search strategy for use in a systematic review.
- Search databases using a combination of subject headings and keyword search techniques.
- Document and report a search strategy.



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