Part One: Background, objectives, outcome measures and types of studies

COI statement

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

Campbell & Cochrane Economics Methods Group
http://methods.cochrane.org/economics/
Learning objectives

- Incorporate economic perspectives into ‘Background’
- Formulate an ‘Objective’ for a critical review of health economic studies
- Identify measures of resource use, costs and cost-effectiveness to be included in ‘Types of outcome measures’
- Identify types of health economic studies to be included in ‘Types of studies’
Preliminary points

- Advisory support from a health economist useful
  - Check with CRG - health economist advisor?
  - Contact Economics Methods Group
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- Focus on how to prepare protocol for a critical review of health economics studies
Chapter 15: Incorporating economics evidence


Key points

- Economics is the study of the optimal allocation of limited resources for the production of benefit to society and is therefore relevant to any healthcare decision.
- Optimal decisions also require best evidence of effectiveness.
- This chapter describes methods for incorporating economics perspectives and evidence into Cochrane reviews, with a focus on critical review of health economics studies.
- Incorporating economics perspectives and evidence into Cochrane reviews can enhance their usefulness and applicability for healthcare decision-making and new economic analyses.

15.1 The role and relevance of economics evidence in Cochrane reviews
15.2 Planning the economics component of a Cochrane review
15.3 Locating studies
15.4 Selecting studies and collecting data
15.5 Addressing risk of bias
15.6 Analysing and presenting results
15.7 Addressing reporting biases
15.8 Interpreting results
15.9 Conclusions

Box15.8.a: Highlighting a need for further economic evidence

15.11 References
Background

Well-formulated review questions occur in the context of an already-formed body of knowledge. The background should address this context, help set the rationale for the review, and explain why the questions being asked are important. It should be concise (generally around one page when printed) and be understandable to the users of the intervention under investigation. All sources of information should be cited.

Description of the condition

The review should begin with a brief description of the condition being addressed and its significance. It may include information about the biology, diagnosis, prognosis and public health importance (including prevalence or incidence).

Description of the intervention

A description of the experimental intervention(s) should place it in the context of any standard, or alternative interventions. The role of the comparator intervention(s) in standard practice should be made clear. For drugs, basic information on clinical pharmacology should be presented where available. This information might include dose range, metabolism, selective effects, half-life, duration and any known interactions with other drugs. For more complex interventions, a description of the main components should be provided.

How the intervention might work

This section might describe the theoretical reasoning why the interventions under review may have an impact on potential recipients, for example, by relating a drug intervention to the biology of the condition. Authors may refer to a body of empirical evidence such as similar interventions having an impact or identical interventions having an impact on other populations. Authors may also refer to a body of literature that justifies the possibility of effectiveness.
Faecal incontinence... can be a debilitating problem with medical, social and economic implications... In the United States more than $400 million is spent each year on a range of both urinary and faecal incontinence products... During 1991 the direct costs of pads, appliances and other prescription items throughout hospitals and long term care settings in the UK for incontinence in general was estimated at £68 million... With the rise in numbers of elderly people in the world, this condition will be an increasing challenge to both healthcare services and home carers.
Background: Describe potential impacts of intervention(s) on resource use (costs)

- ‘Resource inputs’ (‘input costs’)
  - e.g. staff time and skills, equipment, devices, drugs, hospital care, patient out-of-pocket expenses...

- ‘Resource consequences’ (‘downstream costs’)
  - e.g. health care and other resources used to manage sequelae and complications of treatment, time off work...
It is important to consider whether use of Bone Morphogenetic Protein is worthwhile...given the incremental costs (resource use) and benefits (effects) which may be associated with the intervention.
Objectives

This should begin with a precise statement of the primary objective of the review, ideally in a single sentence. Where possible the style should be of the form "To assess the effects of [intervention or comparison] for [health problem] for/in [types of people, disease or problem and setting if specified]". This might be followed by a series of specific objectives relating to different participant groups, different comparisons of interventions or different outcome measures. It is not necessary to state specific hypotheses.
Objectives: Clinical effects

To assess the [clinical] effectiveness of Bone Morphogenetic Protein for fracture healing in skeletally mature adults, compared to current standard treatments
Objectives: Economics

To critically appraise and summarise current evidence on the [resource use/ costs/ and cost-effectiveness] associated with [intervention or comparison] for [health problem] for/in [types of people, disease or problem and setting if specified]
Objectives: Economics

To critically appraise and summarise current evidence on the (incremental) resource use, costs and cost-effectiveness of Bone Morphogenetic Protein for fracture healing in skeletally mature adults, compared to current standard treatments.
Objectives: Economics

To critically appraise and summarise current evidence on the (incremental) resource use, costs and cost-effectiveness of Bone Morphogenetic Protein for fracture healing in skeletally mature adults, compared to current standard treatments.
Use of ‘Clinical event pathway descriptions’

Person with stroke

Admission to specialist acute stroke unit

Brain imaging (if required)

Thrombolysis & mechanical clot retrieval

Decompressive hemicranietomy

Managing acute intracerebral haemorrhage

Antiplatelet & anticoagulant treatment

Maintaining or restoring homeostasis

Assessing swallow function & oral nutrition

Statin treatment

Assessing & managing carotid stenosis following acute non-disabling stroke

Early mobilization

Stroke rehabilitation

Clinical event

Clinical event management

Resources used to manage Clinical events
Use of ‘Clinical event pathway descriptions’

Person who had stroke → Information, support & training → Screening & assessment → Setting goals → Rehabilitation plan → Therapy → Early support discharge → Transfer of care from hospital to community → Return to work → Long term support → Clinical event

Special rehabilitation services → Multidisciplinary rehabilitation team → Intensity of stroke rehabilitation in terms of functioning, movement, communication, self care etc.

System to ensure transfer of care → Movement difficulties after stroke → Physiotherapy and different body trainings

Subsequent clinical events → Resources used to Manage subsequent Clinical events
Criteria for considering studies for this review

Types of studies

Eligible study designs should be stated here, along with any thresholds for inclusion based on the conduct of the studies or their risk of bias. For example, 'All randomized controlled comparisons' or 'All randomized controlled trials with blind assessment of outcome'. Exclusion of particular types of randomized studies (for example, cross-over trials) should be justified.

See also

- Eligibility criteria for types of study designs are discussed in Chapter 5 (Section 5.5)

Types of participants

The diseases or conditions of interest should be described here, including any restrictions such as diagnoses, age groups and settings. Subgroup analyses should not be listed here (see 'Subgroup analysis and investigation of heterogeneity' under 'Methods').

See also

- Eligibility criteria for types of participants are discussed in Chapter 5 (Section 5.2)

Types of interventions

Experimental and comparator interventions should be defined here, under separate subheadings if appropriate. It should be made clear which comparisons are of interest. Restrictions on dose, frequency, intensity or duration should be stated. Subgroup analyses should not be listed here (see 'Subgroup analysis and investigation of heterogeneity' under 'Methods').

See also

- Eligibility criteria for types of interventions are discussed in Chapter 5 (Section 5.3)

Types of outcome measures

Note that outcome measures do not always form part of the criteria for including studies in a review. If they do...
Types of outcome measures: economic outcomes

- Resource use
- Costs
- Cost-effectiveness
- Magnitude
- Time horizon
- Analytic perspective
Magnitude and analytic perspective

Bone Morphogenetic Protein for fracture healing

- Costs of acute treatment and care
- Costs of revisional procedures
- Costs of secondary interventions
- Cost of antibiotics
- Cost of outpatient visits
- Cost of travel to outpatient visits
- Cost of physiotherapy
- Cost of child care
- Lost wages
- Lost productivity (work output)
- Wages paid to temporary staff to cover absence
Types of outcome measures: economic outcomes

- **Resource use**
  - Specific items of resource use
    - Length of hospital stay (days)
    - Duration of operation (minutes)
    - Outpatient visits (number)
    - Pharmaceuticals (treatment duration and dosage)
    - Time to return to work (days)
Types of outcome measures: economic outcomes

- Resource use
  - Exceptions
    - Other direct resource use associated with complications of treatment
Types of outcome measures: economic outcomes

- **Costs**
  - Specific cost items?
    - Cost of hospital stay
    - Cost of operation
    - Cost of outpatient visits
  - Specific cost categories
    - Direct medical costs
    - Non-medical costs
Types of outcome measures: economic outcomes

- **Costs**
  - **Level**
    - Cost of ‘X’ per patient (specific cost items)
      - Average (mean) cost of ‘X’ per patient
    - Total direct medical costs per patient
      - Average (mean) total direct medical costs per patient
    - Total non-medical costs per patient
      - Average (mean) total non-medical costs per patient
Types of outcome measures: economic outcomes

- **Cost-effectiveness**
  - Incremental cost-effectiveness ratios (ICERs)
  - Incremental cost per quality-adjusted life year (QALY)
  - Incremental cost per disability-adjusted life year (DALY)
  - Incremental cost-benefit ratios
  - Net benefits
Types of outcome measures

Note that outcome measures do not always form part of the criteria for including studies in a review. If they do not, then this should be made clear. Outcome measures of interest should be listed in this section whether or not they form part of the eligibility criteria.

See also

- Types of outcomes are discussed in Chapter 5 (Section 5.4).
- The importance of addressing patient-relevant outcomes is discussed further in Chapter 11 (Section 11.5.2); see also an extended discussion of patient-reported outcomes in Chapter 17.

Primary outcomes

The review’s primary outcomes should normally reflect at least one potential benefit and at least one potential area of harm, and should be as few as possible. It is normally expected that the review should be able to analyse these outcomes if eligible studies are identified, and that the conclusions of the review will be based in large part on the effects of the interventions on these outcomes.

Secondary outcomes

Non-primary outcomes should be listed here. The total number of outcomes addressed should be kept as small as possible.

The following optional (level 4) headings may be helpful, as supplements or replacements for the headings above:

- Main outcomes for ‘Summary of findings’ table
- Timing of outcome assessment
- Adverse outcomes
- Economic data

Search methods for identification of studies

The methods used to identify studies should be described. The following headings may be useful:

- Defining the review question and developing criteria for inclusion and exclusion of studies
- Searching for studies
- Selecting studies and collecting data

Part 2: General methods for Cochrane reviews

- Front page
- Handbook information
- Part 1: Cochrane reviews
  - 1 Introduction
  - 2 Preparing a Cochrane review
  - 3 Maintaining reviews: updates, amendments and feedback
  - 4 Guide to the contents of a Cochrane protocol and review
    - 4.1 Introduction
    - 4.2 Title and review information (or protocol information)
    - 4.3 Abstract
    - 4.4 Plain language summary
    - 4.5 Main text
      - Introductory text
      - Background
      - Objectives
      - Methods
      - Results
      - Discussion
      - Authors’ conclusions
      - Acknowledgements
      - Contributions of authors
      - Declarations of interest
      - Differences between protocol and review
      - Published notes
    - 4.6 Tables
    - 4.7 Studies and references
    - 4.8 Data and analyses
      - Figure 4.8.a: Illustration of the hierarchy of the data
    - 4.9 Figures
    - 4.10 Sources of support to the review
    - 4.11 Feedback
    - 4.12 Appendices
    - 4.13 Chapter information
    - 4.14 References
- Part 2: General methods for Cochrane reviews
- 5 Defining the review question and developing criteria for inclusion and exclusion of studies
- 6 Searching for studies
- 7 Selecting studies and collecting data

Search methods for identification of studies

Part 2: General methods for Cochrane reviews

5 Defining the review question and developing criteria for inclusion and exclusion of studies
6 Searching for studies
7 Selecting studies and collecting data
Types of studies: Economic evaluation studies

| Are both costs (inputs) and consequences (outputs) of the alternatives examined? |
|-------------------------------------------------|-----------------|
| No                                             | Yes             |
| Examine only consequences                      | Examine only costs |
| 1A Partial evaluation                          | 2 Partial evaluation |
| Outcome description                             | Cost description |
| Cost-outcome description                       |                 |
| 3A Partial evaluation                          | 4 Full economic evaluation |
| Efficacy or effectiveness evaluation           | Cost analysis    |
| Cost-effectiveness analysis (CEA)              |                 |
| Cost-utility analysis (CUA)                    |                 |
| Cost-benefit analysis (CBA)                    |                 |

Is there comparison of two or more alternatives?
Bone Morphogenetic Protein
(Example)

Objectives
To critically appraise and summarise current evidence on the (incremental) resource use, costs and cost-effectiveness of Bone Morphogenetic Protein for fracture healing in skeletally mature adults, compared to current standard treatments

Types of intervention
BMP versus surgery alone
BMP versus surgery with or without bone graft
BMP and bone substitutes versus surgery and bone substitutes
Types of studies: Economic evaluation studies

| Are both costs (inputs) and consequences (outputs) of the alternatives examined? |
|---------------------------------|-----------------|
|                                 | No              | Yes             |
|                                 | Examine only   | Examine only    |
|                                 | consequences    | costs           |
| 1A Partial evaluation          | 1B Partial      | 2 Partial       |
|                                 | evaluation      | evaluation      |
|                                 | Outcome description | Cost description |
|                                 |                  | Cost-outcome description |
| Is there comparison of two or more alternatives? |      |                  |
|                                 | No              | Yes             |
|                                 | 3A Partial      | 3B               |
|                                 | evaluation      | 4 Full economic  |
|                                 | Efficacy or     | evaluation      |
|                                 | effectiveness   | Cost analysis    |
|                                 | evaluation      |                  |
|                                 |                  | Cost-effectiveness analysis (CEA) |
|                                 |                  | Cost-utility analysis (CUA) |
|                                 |                  | Cost-benefit analysis (CBA) |
Types of studies: Full economic evaluation studies

- Cost-effectiveness analysis: cost per unit of effect (ICER)
- Cost utility analysis: cost per QALY/ cost per DALY (ICER)
- Cost-benefit analysis: cost-benefit ratio/net benefit
Types of studies: health economics studies

Comparative health economics studies

*Resource utilisation studies*
‘Comparative’ resource utilisation studies (e.g. resource use measured within an RCT)

*Partial economic evaluations*
Cost analyses

*Full economic evaluations*
Cost-effectiveness analyses
Cost-utility analyses
Cost-benefit analyses
Types of studies: health economics studies

Types of studies
Randomised controlled trials. Full economic evaluations (cost-effectiveness analyses, cost-utility analyses and cost-benefit analyses), cost analyses and comparative resource utilisation studies.
Types of studies: the issue of scope

- Full economic evaluations, cost analyses and comparative resource utilisation studies can all be conducted alongside an RCT.

- Full economic evaluations can also be conducted as ‘model-based economic evaluations’.

- Cost analyses and comparative resource utilisation studies can also be conducted as ‘stand-alone’ studies.
Types of studies: the issue of scope

Option 1

Include only ‘empirical’ health economics studies conducted alongside single, primary studies of effects which meet eligibility criteria for the review of intervention effects
Types of studies: health economics studies

Types of studies (Option 1)
Randomised controlled trials. Full economic evaluations (cost-effectiveness analyses, cost-utility analyses and cost-benefit analyses), cost analyses and comparative resource utilisation studies conducted alongside a randomised controlled trial.
Types of studies: the issue of scope

Option 2

Include ‘empirical’ health economics studies conducted alongside single, primary studies of effects which meet eligibility criteria for the review of intervention effects

AND

Health economics studies utilising effects data sourced from one or more single, primary studies meeting eligibility criteria for the review of intervention effects
Types of studies: health economics studies

Types of studies (Option 2)
Randomised controlled trials. Full economic evaluations (cost-effectiveness analyses, cost-utility analyses and cost-benefit analyses) conducted alongside a randomised controlled trial or those utilising effects data generated using either a meta-analysis of randomised controlled trials or a single randomised controlled trial. Cost analyses and comparative resource utilisation studies conducted alongside a randomised controlled trial.
Types of studies: the issue of scope

Option 3

Include all health economics studies meeting eligibility criteria, populations and comparisons, whether or not conducted alongside or utilising effects data sourced from studies which meet eligibility criteria for the review of intervention effects.
Types of studies: health economics studies

Types of studies (Option 3)
Randomised controlled trials. Full economic evaluations (cost-effectiveness analyses, cost-utility analyses and cost-benefit analyses), cost analyses and comparative resource utilisation studies – any study design.
The Campbell & Cochrane Economics Methods Group (CCEMG) is an international network of individuals with an interest and expertise in approaches to evidence synthesis that combine economics and systematic review methods. CCEMG strives to promote the inclusion of economic perspectives and evidence in systematic reviews of health care, social welfare, education and criminal justice interventions.

This is achieved through our development and provision of methods guidance, training, peer review and advisory support for economics components of reviews. CCEMG is a Methods Group of Cochrane and a subgroup of The Campbell Collaboration Methods Coordinating Group.

http://methods.cochrane.org/economics/
HOW TO INCLUDE ECONOMICS IN COCHRANE REVIEW PROTOCOLS

Part One: Background, objectives, outcome measures and types of studies

Campbell & Cochrane Economics Methods Group
http://methods.cochrane.org/economics/