BMJ Open Barriers and enablers to monitoring and deprescribing opioid analgesics for chronic non-cancer pain: protocol for a qualitative evidence synthesis using the Theoretical Domains Framework

Amanda J Cross ^(b), ^{1,2} Rachelle Buchbinder, ^{1,2} Allison Bourne, ^{1,2} Christopher Maher, ^{3,4} Stephanie Mathieson, ^{3,4} Chung-Wei C Lin, ^{3,4} Denise A O'Connor^{1,2}

ABSTRACT

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For numbered affiliations see end of article.

Correspondence to

Dr Amanda J Cross; amanda.cross@monash.edu **Introduction** The over-prescription and overuse of opioid analgesics for chronic non-cancer pain (CNCP) is a growing issue. Synthesis of evidence about the barriers and enablers to reducing long-term opioid prescribing and use will enable the development of tailored interventions to address both problems.

Objective To synthesise the barriers and enablers to monitoring the ongoing appropriateness of opioid treatment and deprescribing opioids for CNCP from the clinician, patient and general public point of view, and to map the findings to the Theoretical Domains Framework (TDF).

Methods and analysis We will perform a qualitative evidence synthesis using the TDF. We will include qualitative research that has explored clinician, patient and the general public's perceptions regarding barriers and enablers to monitoring and deprescribing opioids for CNCP. Studies will be identified via searches in MEDLINE, EMBASE, CINAHL, AMED and PsycINFO. Databases will be searched from inception to July 2019, and the studies must be published in English. Article selection and data extraction will be completed independently by two review authors. Methodological guality of included studies will be independently assessed by two review authors using the Critical Appraisal Skills Programme quality assessment tool. We will conduct thematic synthesis and then map identified themes and sub-themes to TDF domains. Confidence in synthesis findings will be evaluated using the Grading of Recommendations Assessment, Development, and Evaluation Confidence in the Evidence from Reviews of Qualitative Research tool. Ethics and dissemination Ethical approval is not required to conduct this review. We will publish the results in a peer-reviewed journal.

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INTRODUCTION

Chronic non-cancer pain (CNCP), defined as pain persisting beyond 3 months unrelated to a malignancy, is highly prevalent worldwide

Strengths and limitations of this study

- This study will inform development of tailored, effective, theory-based interventions for reducing inappropriate use of opioid analgesics in chronic non-cancer pain.
- We will use systematic search strategies to identify relevant qualitative research.
- To minimise bias, two reviewer authors will independently select, extract data and assess the methodological quality of included studies.
- We will use Grading of Recommendations Assessment, Development, and Evaluation Confidence in the Evidence from Reviews of Qualitative Research to report the confidence in synthesis findings.
- Synthesis will be limited to quotes and themes explored in the original reports. We will not seek original data.

with population estimates varying from 20% to 40%.^{1–5} Optimal management of CNCP is essential to minimise the burden on individuals and the wider healthcare system. While evidence supports the use of opioid analgesics for acute and cancer-related pain, there is limited evidence supporting the long-term efficacy and safety of opioids for CNCP.^{6–12} Furthermore, evidence suggests that potential harms such as risk of overdose, opioid abuse and opioid addiction, often outweigh potential benefits.^{13 14}

Opioid use for CNCP is common. In Australia, almost three million adults are using opioids for non-cancer pain each year,¹⁵ and there was a 15-fold increase in opioid dispensing between 1992 and 2012.¹⁶ In 2015, the amount of opioids prescribed in the USA was three times higher than in 1999,¹⁷ and

Table 1 WHO guide to good prescribing and opioid prescribing in CNCP guidelines		
WHO Principles of Good Prescribing ²⁸		Guidelines for prescribing opioids for CNCP ^{10 20-22}
Step 1	Define the patient's problem	 Assess the patient using a multidisciplinary approach. Non-pharmacological therapy and non-opioid pharmacological therapy are preferred for CNCP.
Step 2	Specify the therapeutic objective	Before starting opioid therapy, clinicians should establish realistic treatment goals with the patient and set a review date. Written, structured clinician-patient agreements/contracts for opioid use could be considered.
Step 3	Verify the suitability of the medication	A careful assessment of the benefits and risks of prescribing an opioid for each specific patient should be considered.
Step 4	Write a prescription	 Start with a low dose and adjust slowly according to response. Do not introduce an opioid at the same time as another drug.
Step 5	Give information, instructions and warnings	Discuss the adverse effects, possible harms and realistic benefits of long-term opioid therapy with patients.
Step 6	Monitor (and stop?) the treatment	 Regularly review the patient to monitor progress, evaluate benefits and harms, and assess if ongoing treatment is needed. Reviews should be within 4 weeks of starting opioid treatment or of changes in dose, and minimum every 3 months for continued treatment. If opioid treatment is ineffective, or benefits do not outweigh harms, then opioid treatment should be tapered slowly and under supervision.

CNCP, chronic non-cancer pain; WHO, World Health Organization.

the total number of opioid prescriptions peaked at more than 255 million in 2012.¹⁸ While the rates of new patients prescribed opioids in Australia and the USA have started to decline, high-risk prescribing, including initiating with strong opioids and prescriptions for more than 3 days' supply, is still a major problem.^{15 19}

Recent evidence-based guidelines do not recommend opioids as first-line management for CNCP.¹⁰ ²⁰ ²¹ If opioids are prescribed, clinicians should monitor patient progress and evaluate benefits and harms of continuing opioid therapy with patients at least every 3months (table 1).¹⁰ ^{21–23} If opioid treatment is ineffective or if the harms outweigh the benefits, then opioid treatment should be tapered. Unfortunately, many of the harms associated with long-term opioid use, including tolerance and addiction,⁹ can make the process of reducing or deprescribing opioids challenging for both clinicians and patients. There is also a lack of quality evidence on safe and effective interventions for deprescribing opioids in CNCP.²⁴

Understanding barriers and enablers to reducing the long-term prescribing and use of opioids and mapping these factors to theoretical mechanisms of behaviour change, is an important step towards developing tailored, effective approaches for reducing inappropriate use of opioids in CNCP. The Theoretical Domains Framework (TDF), which consists of 14 domains and 84 theoretical constructs drawn from over 30 theories of behaviour change, offers a systematic and theory-based approach for identifying determinants of behaviour change and facilitating design of tailored, theory-based implementation interventions.^{25 26} It allows for key factors that influence guideline recommended behaviour and clinical practice

change to be linked to evidence-based behaviour change techniques.²⁵

To date, there has been only one qualitative evidence synthesis relating to opioid use in CNCP, and it has explored health professionals' experience of prescribing opioids for CNCP.²⁷ It has found that there is often a sense of uncertainty about when to prescribe opioids, and that healthcare professionals struggle to balance their professional duty to 'get rid of pain' with the social suspicion and hostility surrounding prescribing opioids.²⁷ This review did not focus on barriers and enablers to monitoring ongoing appropriateness of opioid treatment, nor barriers and enablers to reducing or deprescribing opioids. A second qualitative evidence synthesis is currently under way which aims to explore patients' experiences taking opioids for CNCP but does not include an explicit aim to map barriers and enablers to a behaviour change framework.²⁸

Aim

To synthesise available qualitative research regarding barriers and enablers to monitoring ongoing appropriateness of opioid treatment and deprescribing opioids for CNCP from the clinician, patient and general public point of view, and to organise the findings using the TDF.

METHODS

Selection criteria

We will include English language full-text articles or reports (ie, excluding conference abstracts) that fulfil the following criteria:

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Types of studies: Studies that used any type of qualitative method(s) to obtain data (eg, focus group, interviews). Studies that included mixed methods would be included where the qualitative data were analysed independently of the quantitative data.

Types of participants: Clinicians (regardless of discipline), patients (previously or currently being treated for CNCP with one or more opioids) and/or their carers, and the general public who have been interviewed about the barriers and enablers to monitoring ongoing appropriateness of opioid treatment and/or deprescribing opioids for CNCP.

Types of settings: All healthcare settings in all countries.

Types of pain: CNCP defined as pain lasting beyond 3 months unrelated to malignancy.

Types of outcome measures: Studies that report perceptions about barriers and/or enablers to guidelinerecommended safe and appropriate prescribing and use of opioids in CNCP. Specifically, we will focus on guideline recommendations related to step 6 of the World Health Organization (WHO) Principles of Good Prescribing²⁹ (table 1), including monitoring ongoing appropriateness of opioid treatment (ie, effectiveness, benefits and/ or harms) and stopping opioid treatment (ie, tapering, reducing or deprescribing treatment that may be ineffective or harmful). To ensure studies can provide rich data specific to the use of opioids, studies will only be included where their study aim and/or methodology (eg, interview guide) specifically mention aspects of monitoring ongoing appropriateness of opioid treatment or stopping opioid treatment. Where studies include participants with multiple medications, we will include studies where the experiences with opioids are reported separately.

Studies exploring the following topics will not be a focus of this review and such studies will be excluded:

- Initiating opioid treatment
- Opioid use and management of CNCP in the context of substance use disorder, addiction or illicit opioid use.
- Impact or evaluation of policy-level, organisationlevel, practitioner-level or patient-level interventions (eg, legislation change or experiences with a new programme/intervention).

Search methods

We will develop a search strategy with the assistance of an Information Specialist based at Monash University, and with guidance from the qualitative research chapter in the Cochrane Handbook.³⁰

The following databases will be searched to identify relevant articles: MEDLINE, EMBASE, CINAHL, AMED and PsycINFO. We will search databases from inception to July 2019, but will limit the search to articles in English.

The search will be conducted in two parts. Part 1 will combine terms related to: CNCP as recommended by the Cochrane Pain, Palliative and Supportive Care group; opioids; monitoring, deprescribing or reducing; and barriers and enablers. Part 2 will identify studies that have utilised qualitative research methodology. Qualitative filters will be used where available to enhance sensitivity and specificity. To obtain final search results, we will combine parts 1 and 2. The proposed MEDLINE search strategy is outlined in online supplementary appendix 1.

The reference list of included studies will also be searched for relevant articles, and we will track citations of included studies using Web of Science to ensure we gather all relevant literature. Experts in the field will also be approached to identify other potentially relevant studies using the existing professional networks of the authors.

Selection of studies

Two review authors will independently screen all titles and abstracts yielded from the search, and all full-text articles considered relevant. Any disagreements will be resolved via discussion or by consultation with a third review author when necessary. Search and screening results will be summarised in a Preferred Reporting Items of Systematic Reviews and Meta-Analyses flow diagram.³¹

Data collection

Pairs of review authors will independently extract data from each included study using a standardised data collection form. Any disagreements will be resolved by discussion or by consultation with a third review author when necessary. A piloted data extraction form will capture information regarding the:

- ► Study details (authors, year of publication).
- ▶ Research question.
- Participants (number, demographic characteristics, professional background, history of opioid use, method of selection).
- ► Type and duration of CNCP.
- Type and duration of opioid, including stage of deprescribing if applicable.
- ► Setting (type of healthcare, country, urban/rural).
- ▶ Method of data collection (eg, interview, focus group).
- ► Data analysis method (eg, thematic analysis).
- Results (themes, sub-themes, supporting quotes, conclusions).

Box 1 Grading of Recommendations Assessment, Development, and Evaluation Confidence in the Evidence from reviews of Qualitative research approach³⁴ to confidence in findings of qualitative evidence syntheses

- High confidence: highly likely that the review finding is a reasonable representation of the phenomenon of interest.
- Moderate confidence: likely that the review finding is a reasonable representation of the phenomenon of interest.
- Low confidence: possible that the review finding is a reasonable representation of the phenomenon of interest.
- Very low confidence: unclear whether the review findings is a reasonable representation of the phenomenon of interest.

CERQual, Confidence in the Evidence from Reviews of Qualitative Research; GRADE, Grading of Recommendations Assessment, Development, and Evaluation.

Quality assessment

The quality of included studies will be independently assessed by two review authors using the Critical Appraisal Skills Programme (CASP) Checklist.³² Any disagreements will be resolved via discussion or by consultation with a third review author when necessary. The CASP tool uses a checklist of 10 questions, each of which includes multiple signalling questions to help users interpret the items (29 signalling questions in total). Studies will not be excluded based on low quality. A summary table indicating the presence or absence of CASP items will be included, and a narrative summary of the quality of the included studies will be provided.

Data analysis and synthesis

We will initially use an inductive approach to coding and conduct thematic synthesis, as outlined by Thomas and Harden.³³ Each line of extracted text will be reviewed and codes will be developed based on the content and meaning of each extract. As new studies are coded, existing codes will be reviewed and revised with new codes added as needed. When all studies have been coded, all text related to each code will be reviewed for consistency of coding across studies. Two review authors will independently code an initial subset of studies and then meet to discuss any discrepancies until consensus is reached. The remaining studies will then be coded by one author and verified by a second author. Codes will be reviewed for similarities and differences and organised into related descriptive themes. A draft summary of the descriptive themes will be prepared by one author and discussed by the review team until consensus is reached.

Next, the emergent themes will be mapped to the domains of the $\text{TDF}^{25\ 26}$ independently by two review authors or if there is no relevant domain placed in an 'other' category. A draft summary of the TDF-aligned themes will be prepared by one author and discussed by the review team until agreement on coding interpretations is reached.

In the final stage, review findings will be drafted to summarise analytical themes and corresponding theoretical domains relating to barriers and enablers to ongoing monitoring of opioid use and deprescribing opioids for CNCP from the clinician, patient and general public point of view. The review findings will be prepared by one author and discussed by the review team until consensus is reached.

Assessment of confidence in review findings

Pairs of independent authors will use the Grading of Recommendations Assessment, Development, and Evaluation (GRADE) Confidence in the Evidence from Reviews of Qualitative Research tool to assess the confidence in the review findings.³⁴ This tool considers four factors that influence the confidence in synthesis of qualitative studies:

- a. Methodological limitations of the studies contributing to a review finding (based on CASP assessments).
- b. Relevance of studies to the review question of studies contributing to a review finding.
- c. Coherence of data contributing to a review finding.
- d. Adequacy of supporting data supporting a review finding.

The overall confidence in each review finding (ie, for each theme generated) will be graded as: high, moderate, low or very low (see box 1).³⁴ A final decision on confidence in review findings will be reached through discussion and consensus among the review team. Review findings, confidence judgement for each finding and an explanation of the judgement will be presented in a Summary of Qualitative Findings table (see table 2).

Reporting

The study will be reported in accordance with the Enhancing transparency in reporting the synthesis of qualitative research statement.³⁵

Patient and public involvement

Patients and the public will not be involved directly in the design and conduct of the review. However, the development of the review questions was informed by the Wiser Health Care Research Collaboration which involves consumer advocates and health professionals.

Ethics and dissemination

Ethical approval is not required to conduct this review. We will publish the results in a peer-reviewed journal.

DISCUSSION

Here, we present the design of a qualitative evidence synthesis using the TDF to explore clinician, patient and general public barriers and enablers to monitoring ongoing appropriateness of opioid treatment and deprescribing opioids for CNCP, and align key findings with evidence-based behaviour change techniques. This is an important step towards developing tailored, effective, theory-based interventions for reducing the inappropriate use of opioids in CNCP.

Study status

Databases were searched in July 2019, screening and study selection occurred in August–September 2019 and data extraction is ongoing. We anticipate completion of this review by June 2020.

Author affiliations

¹Monash Department of Clinical Epidemiology, Cabrini Institute, Malvern, Victoria, Australia

²Department of Epidemiology and Preventive Medicine, School of Public Health and Preventive Medicine, Monash University Faculty of Medicine Nursing and Health Sciences, Melbourne, Victoria, Australia

³Institute for Musculoskeletal Health, Sydney, New South Wales, Australia ⁴Sydney School of Public Health, The University of Sydney Faculty of Medicine and Health, Sydney, New South Wales, Australia

Twitter Amanda J Cross @AmandaJCross

Contributors AJC, DAO'C and RB conceived the idea for the study. AJC and DAO'C wrote the first draft of the manuscript. RB, AB, CM, SM and C-WCL contributed to revising subsequent drafts critically for important intellectual content. All authors approved the final version of the manuscript and agree to be accountable for its content.

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ORCID iD

Amanda J Cross http://orcid.org/0000-0001-6001-9211

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