BMJ Open Outcomes to evaluate care for adults with acute dental pain and infection: a systematic narrative review

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ABSTRACT

Objective To identify outcomes reported in peer-reviewed literature for evaluating the care of adults with acute dental pain or infection. **Design**

Systematic narrative review.

Setting/participants Primary research studies published in peer-reviewed literature and reporting care for adults with acute dental pain or infection across healthcare settings. Reports not in English language were excluded. Study selection Seven databases (CINAHL Plus, Dentistry and Oral Sciences Source, EMBASE, MEDLINE, PsycINFO, Scopus, Web of Science) were searched from inception to December 2020. Risk of bias assessment used the Critical Appraisal Skills Programme checklist for randomised controlled trials and Quality Assessment Tool for Studies of Diverse Design for other study types.

Outcomes Narrative synthesis included all outcomes of care for adults with acute dental pain or infection. Excluded were outcomes about pain management to facilitate treatment, prophylaxis of postsurgical pain/ infection or traumatic injuries.

Results Searches identified 19 438 records, and 27 studies (dating from 1993 to 2020) were selected for inclusion. Across dental, pharmacy, hospital emergency and rural clinic settings, the studies were undertaken in high-income (n=20) and low/middle-income (n=7) countries. Two clinical outcome categories were identified: signs and symptoms of pain/infection and complications following treatment (including adverse drug reactions and reattendance for the same problem). Patient-reported outcomes included satisfaction with the care. Data collection methods included patient diaries, interviews and in-person reviews.

Discussion A heterogeneous range of study types and qualities were included: one study, published in 1947, was excluded only due to lacking outcome details. Studies from dentistry reported just clinical outcomes; across wider healthcare more outcomes were included. **Conclusions** A combination of clinical and patientreported outcomes are recommended to evaluate care for adults with acute dental pain or infection. Further research

adults with acute dental pain or infection. Further research is recommended to develop core outcomes aligned with the international consensus on oral health outcomes. **PROSPERO registration number** CRD42020210183.

INTRODUCTION

Acute dental pain has a significant impact on quality of life.^{1 2} Timely intervention for the

Strengths and limitations of this study

- The first systematic review to examine outcome measures of care for adults with acute dental pain or infection across healthcare settings.
- The outcomes will be important for evaluating new dental antibiotic and opioid stewardship interventions, as these drugs are frequently overprescribed for adults with acute dental pain and infection, exacerbated by the COVID-19 pandemic.
- Studies about paediatric patients, studies about the postoperative management of pain, studies about local anaesthesia to facilitate dental treatment, studies about traumatic injuries and papers not in English language were excluded due to key differences in clinical management.
- Two independent reviewers extracted data and two different reviewers assessed the quality using either the Critical Appraisal Skills Programme (for the randomised controlled trials) or the Quality Assessment Tool for Studies with Diverse Designs.
- Reporting based on the Preferred Reporting Items for Systematic Reviews and Meta-Analyses 2020 guidelines, with searches of seven major electronic databases from their inception to December 2020.

relief of dental pain and infection is essential to prevent worsening of ill health and reduce the risk of potentially life-threatening complications, such as sepsis, airway occlusion or analgesic overdose.^{3 4} Failure of initial treatment to relieve dental pain and infection can result in patient reattending for further treatment, including to emergency medical care.⁵ Thus, ensuring high-quality care for people with acute dental problems is critical for both patient safety and service efficiency. Outcomes to evaluate the care provided for people with acute dental pain and/or infection are important.

Evidence-based clinical guidelines can improve the provision of quality healthcare and patient outcomes.⁶ Guidelines for treating acute dental pain and infection are generally based on the principle that operative dental procedures (such as removal of a tooth or its pulp) are indicated to address the cause and prevent symptoms recurring.⁷ Drugs such as analgesics and antibiotics have a limited role in dentistry and should usually only be used in addition to dental procedures.^{8 9} Suboptimal treatment of dental pain and infection with drug prescriptions instead of dental procedures is common, including by general medical practitioners and in emergency departments.^{10–12} The contribution of dentistry to global efforts to tackle antibiotic resistance¹³ and opioid substance misuse disorder has been highlighted, with a call for the profession to improve its approach to stewardship of these drugs.^{7 14 15}

While a plethora of drug trials for the treatment of dental pain or infection have been published, there is little research on patient outcomes following urgent dental care for acute dental pain or infection.⁵ A rise in the number of trials to evaluate dental antibiotic stewardship and opioid stewardship interventions is anticipated, with a focus on optimising care and judicious use of medicines for adults (where more than 90% of dental prescribing occurs).¹⁶ To evaluate the effectiveness of these sorts of interventions and to enable improvements in the quality of urgent dental care, this study aimed to identify outcomes from the peer-reviewed literature for evaluating care for adults with acute dental pain and/or infection.

Objectives

The research question was 'What measures in the published literature have been employed to evaluate the outcome of care for adults with acute dental pain and/or infection?'

METHODS

Patient and public involvement

A coproduction team designed and delivered this systematic narrative review. Experts by experience (patients) of urgent dental care and/or complications of dental antibiotics (CC and CP) and academic dental professionals (LT, SH and WT) were involved in all stages of this study, from refining the research question and search terms which had been drafted by WT through to disseminating the results. Through discussion between the members of the coproduction team, involvement with each step of the review was allocated according to the skills they wished to develop and the time they had available to contribute at the relevant stages. Individual contributions are indicated in the following sections.

Eligibility criteria

Primary research studies published in peer-reviewed journals were included if they reported outcomes of care for adults (aged over 18 years) treated for acute dental pain and/or infection with advice, prescriptions, or interventions (such as dental extraction). There was no restriction on the year of dissemination.

Studies which included care for children or for people with other oral or dental conditions (such as cervicofacial infections treated as hospital inpatients or postsurgical pain control) were excluded. Studies of urgent dental care for traumatic injuries were excluded as this is a markedly different population and the subject of a separate study.¹⁷ Reports which did not include the outcomes of care provided (or details of how those outcomes were measured) were also excluded, such as studies about the efficacy of local anaesthesia to facilitate the provision of dental procedures at point of care. Primary research studies not published in peer reviewed journals (such as conference abstracts, case studies and other grey literature) were excluded as the research was seeking tried and tested outcomes for use in clinical trials. Studies not in the English language were excluded due to lack of translation facilities. Full details of the inclusion/exclusion criteria are detailed in online supplemental table 1.

Population groups identified for subgroup analysis during the synthesis phase were dental vs other healthcare settings, and high-income versus low-income and middle-income countries (LMICs).

Information sources

On 29 November 2020, seven databases were searched from their earliest dates: CINAHL Plus, Dentistry and Oral Sciences Source, Ovid EMBASE, Ovid Medline, PyscINFO, Scopus and Web of Science.

Search strategy

The search strategy used to identify relevant papers from the database searches was developed in consultation with an information specialist at the University of Manchester. It consisted of 'population' AND 'intervention' terms. Population terms were: (Acute* OR Urgent OR Unschedul* OR Emergenc*) AND (Dental* OR Odontogenic OR Dentoalveolar) AND (Pain OR Toothache OR Pulpitis OR Infection OR Swell* OR Abscess OR Pericoronitis OR Osteitis OR Socket OR Periodontitis OR Implantitis OR Ulcer* OR Stomatitis). Intervention terms were: Patient Care OR Dental Care OR Procedure OR Treat* OR Endodont* OR Exodont* OR Extract* OR Extirpat* OR Incis* OR Drain* OR Debrid* OR Irrigat* OR Prescri* OR Antibiotic* OR Antimicrob* OR Antiseptic OR Analgesi* OR Advice OR Refer*. Full details of the search terms and limits employed with each database are detailed in online supplemental table 2).

Limits included: 'human' as animal and laboratory studies were not eligible for the review, and 'English language' as justified in the 'eligibility criteria' section. There were no limits on the date of included studies.

Selection process

Titles and abstracts from the database searches (undertaken by WT) were transferred into Endnote V.X9 where duplicates were removed (by WT) and the title/abstracts were screened (independently by WT and SH) for potential inclusion. Full texts of all shortlisted studies were

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assessed for eligibility (independently by WT and LT). Where necessary, corresponding authors were contacted to confirm whether the included population met our inclusion criteria. Disagreements at each stage of the process were resolved through discussion between the screeners.

Data collection process

The characteristics (study type, objective and population) and outcomes, data source (patient-reported, clinician observed or administrative system) and data collection instrument were collected from each report by two reviewers (LT and SH) working independently. Disagreements at each stage of the process were resolved through discussion between the reviewers.

Data items: outcomes and other variables

All outcomes relating to the outcomes of care provided to adults with acute dental pain or infection were sought, together with details about the sources of data and timescales between urgent dental treatment received by the participants and completion of data collection. In addition, specific details about the types of studies (eg, randomised controlled trial (RCT) or questionnaire study) and population were sought, including age range of patients, type of healthcare setting (such as dental clinic or pharmacy), country in which the study took place, and whether a high-income or LMIC (based on World Bank definitions.¹⁸ Details about study type, patient age, healthcare setting and country for each included study are provided in table 1, details about which countries were LMICs are highlighted (in bold) in table 2. There was no restriction on time frames for the outcomes and where missing data was identified this was recorded in the results tables. Where necessary, corresponding authors were contacted to provide details relating to the data items sought (such as the age of participants).

Quality assessment

The shortlisted studies were assessed using the Critical Appraisal Skills Programme (CASP) Checklist for RCTs.¹⁹ For studies which used a design not valid for an RCT (as assessed via the CASP RCT checklist), the Quality Assessment Tool for Studies with Diverse Design (QATSDD) was used.²⁰ Quality assessment of all studies was undertaken by WT, with 30% of studies (selected at random from across the CASP and QATSDD sets) independently assessed by CP. Discrepancies in relation to each element of the assessment framework were resolved through discussion between the assessors and, where differences were just one point, the scores were averaged.

Synthesis methods

All studies which had been selected for inclusion and which had passed the quality assessment were eligible for inclusion in synthesis. Outcome data collected were initially categorised by WT based on a framework advocated for antimicrobial stewardship interventions²¹ as the outcomes identified in this study were

The tabular structure displays a summary of outcomes for each study, using the structure identified. Table 2 presents clinical outcomes ('signs/symptoms of dental pain or infection' and 'complications or harm') and patientreported outcomes ('satisfaction with the outcome of care' and 'other') for each study with details of how the outcome was measured (such as numeric pain scale). Sources of data employed in each study and the timescales between treatment provided to participants and completion of data collection are presented in table 3.

RESULTS

Study selection

Of the 19438 records identified from database searches, 27 studies were selected for inclusion (see figure 1). One study, published in 1947, was excluded as it was impossible to tell how the outcomes had been measured.²³ Another study²⁴ which may look like it should be included was excluded as it reported secondary analysis of data collected in other studies.^{25 26}

Study characteristics

The included studies dated between 1993 and 2020 and encompassed a heterogeneous range of designs, from RCTs to questionnaire surveys. Most studies (n=23) took place in dental settings, one was in a hospital emergency department, another in a rural community healthcare clinic and a third was in community pharmacy; the setting for one study was unclear. The earliest 14 studies all took place in high income countries (during the period 1993–2012). Of the 13 studies which took place between 2013 and 2020, seven were based in LMICs (Brazil, Egypt, India, Tanzania and Turkey). Further characteristics of the included studies, including their objectives, are presented in table 1.

Quality assessment

Following application of the inclusion/exclusion criteria, 11 studies were quality assessed using the CASP framework for RCTs (see online supplemental table 3) and 16 using the QATSDD tool (see online supplemental table 4). Many of the studies assessed using the QATSDD criteria scored poorly, for example, due to failure to justify the sample size or provision of a rationale for the analytic method used, and few studies covered the QATSDD criterion about patients being involvement in the study design.

Results of individual studies

The outcomes recorded in each individual study are presented in table 2, including details about how they were measured. Two categories of clinical outcomes and one of patient-report outcomes were identified. Clinical

Table 1 Cha

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Table 1 Characte	eristics of included studie	S	
Study	Study type	Objective	Population* (patient age, setting, country)
Fazakerley <i>et al</i> , 1993 ⁴⁶	Comparative double-blind trial.	To evaluate the efficacy of cephradine, amoxicillin and phenoxymethylpenicillin in the treatment of dentoalveolar infections.	18–65 years. University dental clinic, UK.
Gibson <i>et al</i> , 1993 ⁴⁷	Prospective survey.	To investigate the success of treatment in resolving the chief complaint of pain and to determine the compliance with further dental care for the original dental problem.	18 years or older. University dental clinic, Canada.
Fouad <i>et al</i> , 1996 ⁴⁸	Double-blind, placebo- controlled clinical trial.	To examine the effect of penicillin on the reduction of symptoms and the course of recovery of the localised acute apical abscess after emergency endodontic treatment.	18 years or older. University dental clinic, USA.
Penniston and Hargreaves, 1996 ⁴⁹	Prospective, randomised, double-blind, placebo- controlled clinical trial.	To compare the analgesic efficacy of ketorolac tromethamine following intraoral periapical infiltration injection or intramuscular injection of the drug.	18–65 years. University dental clinic, USA.
Adriaenssen, 1998 ⁵⁰	Open, randomised, multicentre comparative study.	Comparison of the efficacy, safety and tolerability of azithromycin and co-amoxiclav in the treatment of acute periapical abscesses.	18–75 years. Dental practices, Belgium
Doroschak <i>et al</i> , 1999 ⁵¹	Randomised, double- blind, placebo-controlled study.	To determine if a combination of an Non-Steroidal Anti-Inflammatory Drug (NSAID) and an opioid provide greater pain relief than either drug alone.	18–65 years. University dental clinic, US
Gallatin <i>et al</i> , 2000 ⁵²	Prospective, double-blind, randomised study.	To evaluate pain reduction in untreated irreversible pulpitis using an intraosseous injection of Depo-Medrol.	18 years or older. University dental clinic, USA.
Houck <i>et al</i> , 2000 ²⁵	Prospective, randomised blinded study.	To evaluate postoperative pain and swelling after performing a trephination procedure in symptomatic necrotic teeth with radiolucencies.	Adults*. University dental clinic, USA.
Nagle <i>et al</i> , 2000 ⁵³	Prospective, randomised, double-blind study.	To determine the effect of penicillin on pain in untreated teeth diagnosed with irreversible pulpitis.	Adults* University dental clinic, USA.
Henry <i>et al</i> , 2001 ²⁶	Prospective, randomised, double-blind, placebo- controlled study.	To determine the effect of penicillin on postoperative pain and swelling in symptomatic necrotic teeth.	18 years or older. University dental clinic, USA.
Hersh <i>et al</i> , 2003 ⁵⁴	Randomised, double- blind, placebo-controlled clinical trial.	Efficacy and safety of a benzocaine intraoral patch in patients presenting with spontaneous toothache pain	18–65 years. University dental clinic, USA.
Runyon <i>et al</i> , 2004 ⁵⁵	Prospective, randomised, double-blind, placebo- controlled trial.	To determine if penicillin is necessary or beneficial in the treatment of undifferentiated dental pain without overt infection.	18 years or older. Emergency department, USA
Campanelli <i>et al</i> , 2008 ⁵⁶	Clinical study.	To record the objective and subjective systemic signs of emergency patients presenting with pulp necrosis and localised acute apical abscess.	18 years or older. University dental clinic, USA.
Cohen <i>et al</i> , 2009 ²⁸	Cross-sectional survey.	The pharmacist's role in managing toothache pain from the perspective of the patient.	21 years or older. Community pharmacy, USA.
Wilson <i>et al</i> , 2013 ²⁷	Retrospective questionnaire survey.	To record the levels of patient satisfaction with oral urgent treatment and to highlight areas for improvement in both training and service provision.	
Sethi <i>et al</i> , 2014 ⁵⁷	Randomised clinical trial.	To compare and evaluate the effect of an oral dose of 100 mg tapentadol, 400 mg etodolac or 10 mg ketorolac as a pretreatment analgesic for the prevention and control of postoperative endodontic pain in patients with irreversible pulpitis.	18–60 years. Dental college clinic, India.
Pavithra <i>et al</i> , 2015 ⁵⁸	Randomised double blind trial.	To compare and evaluate analgesic effectiveness of Ibuprofen and Aceclofenac in management of acute irreversible pulpitis.	20–50 years. Dental college clinic, India.
Bultema <i>et al</i> , 2016 ⁵⁹	Prospective, double-blind randomised trial.	To compare liposomal bupivacaine versus bupivacaine for pain control in untreated, symptomatic irreversible pulpitis.	18 years or older. University dental clinic, USA.

Continued

Study	Study type	Objective	Population* (patient age, setting, country)
Sebastian <i>et al</i> , 2016 ⁶⁰	Prospective, randomised study.	To compare debridement vs no debridement on postoperative pain in emergency patients with symptomatic pulpal necrosis, and apical radiolucency.	18 years or older. University dental clinic, USA.
Santini <i>et al</i> , 2017 ⁶¹	Double-blind, controlled parallel design.	To compare the overall analgesic effectiveness of two combinations of opioid and non-opioid analgesics for acute periradicular abscess.	Over 18 years. Dental hospital, Brazil
Taggar <i>et al</i> , 2017 ⁶²	Randomised, double- masked, controlled, parallel-group trial.	To compare the analgesic effect of a single dose of ibuprofen sodium dihydrate with that of a comparable dose of ibuprofen acid in endodontic pain patients presenting with moderate to severe pain.	18–60 years. (Setting unclear), USA.
Aaron and Steier, 2018 ⁶³	Single-centre prospective clinical Study.	To determine if dentists are successful in reducing pain caused by acute apical abscess in a National Health Service emergency setting and if different treatment strategies result in different levels of pain reduction.	20–68 years. Primary care dental clinic, UK.
Beus <i>et al</i> , 2018 ²⁹	Prospective, randomised, single-blind study.	To compare the postoperative course of incision and drain with drain placement versus mock incision and drainage procedure with mock drain placement after endodontic debridement in swollen emergency patients.	18 years or older. University dental clinic, USA.
Eren <i>et al</i> , 2018 ⁶⁴	Single-blinded, single- centre, randomised controlled trial.	To evaluate three emergency procedures for their ability to alleviate clinical symptoms associated with symptomatic teeth having signs of (at least) partial irreversible pulpitis.	18–60 years. University dental clinic, Turkey.
Wolf <i>et al</i> , 2019 ⁶⁵	Prospective randomised study.	To compare the outcomes of two emergency treatment procedures to alleviate pain from localised symptomatic apical periodontitis.	18 years or older. University dental clinic, Sweden.
Al-Rawhani <i>et al</i> , 2020 ⁶⁶	Randomised placebo- controlled double-blind trial.	To evaluate the effect of preoperative administration of a single, oral dose of 50 mg diclofenac on postoperative pain in patients with symptomatic irreversible pulpitis.	18 years or older. University dental clinic, Egypt.
da Silva <i>et al</i> , 2020 ⁶⁷	Double-blind, randomised clinical trial.	To compare the acetaminophen administration efficacy or its combination with codeine for pain control in acute apical abscesses cases.	18 years or older. University dental clinic, Brazil.

*Where not specified in the paper, authors were contacted to confirm participants were all aged >18 years and care was for only people with acute dental pain or infection.

outcomes included: 'signs and symptoms of dental pain/ infection', and 'complications or other harm' resulting from treatment or disease progression. Patient-reported outcomes included patient satisfaction with the outcome of care.

As also shown in table 2, various approaches were used for measuring the clinical outcomes, including unidimensional pain scales (such as a Visual Analogue Scale (VAS) or category pain scale), amount of rescue medication taken, and the presence of absence of various signs and symptoms such as swelling, trismus or fever. Complications were assessed by recording whether unplanned visits had been required or whether the patient had experienced symptoms of drug allergy or other adverse effects (such as gastrointestinal symptoms and headaches).

Details about data sources for the outcomes and duration of data collection in each study are presented in table 3. Most of the outcomes were reported by patients (n=20) through diaries, questionnaires or interviews. A minority of studies (n=7) employed clinical observations from in person monitoring or review during or after their treatment appointment. None of the studies used a combination of patient-reported and clinician observed data. No studies employed data from healthcare administrative systems. Data collection in most studies took place over less than a week (n=17). In six studies, the duration of data collection was 1 week, and two of the remaining four studies data collection completed 1 year after the participant received urgent dental treatment.

Results of syntheses

Pain was the most commonly reported sign/symptom (see table 2), including unstimulated/spontaneous pain (n=24), pain stimulated by percussion, chewing or thermal stimulus (n=7) or the need for additional pain relief through use of rescue medication (n=14). Complications or other harm related to the treatment provided included adverse outcomes (such as drug allergy or nausea) and progression of the acute dental condition requiring unplanned visits for additional treatment. Patient satisfaction was only recorded in studies in non-dental healthcare settings^{27 28} and only one dental study included patient-reported outcomes.²⁹

Comparing results between high-income countries and LMICs found just one difference in the outcomes reported: none of the studies undertaken in LMICs reported on swelling as a sign of infection, compared with 35% (n=7/20) of studies undertaken in high-income

Оре	n ac	cess	;																6
	rted	Other															Cost of care		Continued
	Patient-reported outcomes	Satisfaction														Category scale	Category scale		
	is or harm	Juplanned visits		Yes/No	Yes/No														
	Complications or harm	Adverse drug reaction			Allergy GI Tract	Injection pain	Headache	Gl tract Headache Euphoria Sedation										GI Tract Dizziness Headache Heartburn	
		Other signs/ symptoms	Temperature, Lymphadenopathy		Fever, Trismus or Swallowing difficulty		Gingival redness, Bone loss							Temperature, Purulence, Trismus	Malaise				
		Swelling	Numeric scale		Category scale		Yes/No			Numeric scale		Numeric scale		Yes/No					
		Rescue pain relief taken		Yes/No	Amount				Amount	Amount & type	Amount	Amount and type							
d study		Pain Reduction											Verbal pain relief scale				Category scale		
oyed in each include	Signs/symptoms of dental pain or infection	Pain intensity – stimulated					Category scale		Category scale	Numeric scale	Numeric scale	Numeric scale							
Outcome measures employed in each included study	Signs/symptoms of d	Pain intensity— unstimulated	VAS	Yes/no	VAS	VAS, HP-VAS and Category Scale	Category scale	VAS, HP-VAS and Category Scale	Category scale	Numeric scale	Numeric scale	Numeric scale		VAS	VAS			VAS	
Table 2 Out			Fazakerley <i>et</i> <i>al</i> , 1993 ⁴⁶	Gibson <i>et al</i> , 1993 ⁴⁷	Fouad <i>et al</i> , 1996 ⁴⁸	Penniston and Hargreaves, 1996 ⁴⁹	Adriaenssen, 1998 ⁵⁰	Doroschak <i>et</i> <i>al</i> , 1999 ⁵¹	Gallatin <i>et al</i> , 2000 ⁵²	Houck <i>et al</i> , 2000 ²⁵	Nagle <i>et al</i> , 2000 ⁵³	Henry <i>et al</i> , 2001 ²⁶	Hersh <i>et al</i> , 2003 ⁵⁴	Runyon <i>et al,</i> 2004 ⁵⁵ *	Campanelli <i>et</i> al, 2008 ⁵⁶	Cohen <i>et al</i> , 2009 ²⁸ *	Wilson <i>et al</i> , 2013 ²⁷ *	Sethi <i>et al</i> , 2014 ⁵⁷	

Table 2 Cor	Continued									
	Signs/symptoms of dental pain or infection	ntal pain or infection					Complications or harm	s or harm	Patient-reported outcomes	ed
	Pain intensity – unstimulated	Pain intensity – stimulated	Pain Reduction	Rescue pain relief taken	Swelling	Other signs/ symptoms	Adverse drug reaction	Unplanned visits	Satisfaction	Other
Pavithra <i>et al</i> , 2015 ⁵⁸	VAS									
Bultema <i>et al</i> , 2016 ⁵⁹	VAS			Delayed prescription			Numbness	Yes/No		
Sebastian <i>et al</i> , 2016 ⁶⁰	, HP-VAS			Delayed prescription						
Santini <i>et al</i> 2017 ⁶¹	VAS			Yes/No			GI Tract Dizziness Drowsiness Headache			
Taggar <i>et al</i> , 2017 ⁶²	VAS	Bite force to elicit pain	Time to 50% pain relief							
Aaron and Steier, 2018 ⁶³	Modified pain quality assessment scale									
Beus <i>et al</i> , 2018 ²⁹	HP-VAS			Amount and type	Patient perception: 'swelling becoming smaller'	Experience of bad taste or pus drainage				Patient perception: 'feeling better'
Eren <i>et al</i> , 2018 ⁶⁴	VAS	Yes/No on chewing and thermal stimulus		Amount						
Wolf <i>et al</i> , 2019 ⁶⁵	Numeric scale			Yes/No Opioid/ Non-opioid		Antibiotics prescribed		Yes/No		
Al-Rawhani <i>et</i> al, 2020 ⁶⁶	HP-VAS			Yes/No						
da Silva et al, 2020 ⁶⁷	VAS			Yes/No			Gl Tract Dizziness Drowsiness Headache			
*Study undertal Gl, gastrointesti	*Study undertaken in non-dental setting. Gl, gastrointestinal; HP-VAS, Heft Parker Visual Analogue Scale.	sual Analogue Scale.								

6

Eren et al, 2018⁶⁴

Wolf et al, 201965

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Table 3 Data sources and of			Oliziaian ahaamuud	
	Patient report	Questionnaires or interviews	Clinician observed	In novem menitering
16	Patient diary	Questionnaires or interviews	In-person review	In-person monitoring
Fazakerley <i>et al</i> , 1993 ⁴⁶			5 days	
Gibson <i>et al</i> , 1993 ⁴⁷		2 days		
Fouad <i>et al</i> , 1996 ⁴⁸	3 days			
Penniston and Hargreaves, 1996 ⁴⁹	6 hours			
Adriaenssen, 1998 ⁵⁰			10 days	
Doroschak et al, 1999 ⁵¹	1 day			
Gallatin <i>et al</i> , 2000 ⁵²	1 week			
Houck <i>et al</i> , 2000 ²⁵	1 week			
Nagle et al, 2000 ⁵³	1 week			
Henry <i>et al</i> , 2001 ²⁶	1 week			
Hersh <i>et al</i> , 2003 ⁵⁴				90 min
Runyon <i>et al</i> , 2004 ⁵⁵			1 week	
Campanelli <i>et al</i> , 2008 ⁵⁶			2 weeks	
Cohen <i>et al</i> , 2009 ²⁸		1 year		
Wilson <i>et al</i> , 2013 ²⁷		1 year*		
Sethi et al, 2014 ⁵⁷	1 day			
Pavithra et al, 2015 ⁵⁸		45 min		
Bultema <i>et al</i> , 2016 ⁵⁹	3 days			
Sebastian et al, 2016 ⁶⁰	5 days			
Santini <i>et al</i> 2017 ⁶¹	3 days			
Taggar <i>et al</i> , 2017 ⁶²				1 hour
Aaron <i>et al</i> , 2018 ⁶³		1 day		
Beus <i>et al</i> , 2018 ²⁹	4 days			

Al-Rawhani et al, 2020⁶⁶ 2 days da Silva et al, 2020⁶⁷ 3 days

Studies highlighted in bold are those undertaken in LMICs.

*Where not specified in the paper, authors were contacted to confirm the timescales.

5 days

1 week

LMICs, low-income and middle-income countries.

countries. There was also one difference found in data sources for the outcomes: none of the LMIC-based studies recorded clinician observed outcomes compared with 30% (n=6/20) of studies in high-income countries. No differences were found in data collection periods.

DISCUSSION

A diverse range of measures were identified to assess the outcomes of care for adults presenting with acute dental pain and/or infection across a range of healthcare settings in high income and LMICs. Most were clinical outcomes, such as signs and symptoms of pain and infection and complications or other harms following treatment (such as drug allergy). Patient-reported outcomes relating to satisfaction were only used in studies from

non-dental settings. The range of outcomes and data collection periods were similar between high income countries and LMICs. Just one key difference was noted in their assessment: none of the LMIC studies reported clinician-observed data. This is the first study to focus comprehensively on outcomes relating to acute dental conditions and a lack of consensus in outcomes reported across the studies was found.

Due to the heterogeneous range of studies identified for inclusion, a systematic narrative review was selected to enable synthesis of the results. This type of review is, however, more subjective, and open to potential bias than conventional systematic reviews. Core outcome sets (COS) can improve consistency in reporting and maximise the value derivable from studies.³⁰ Further research

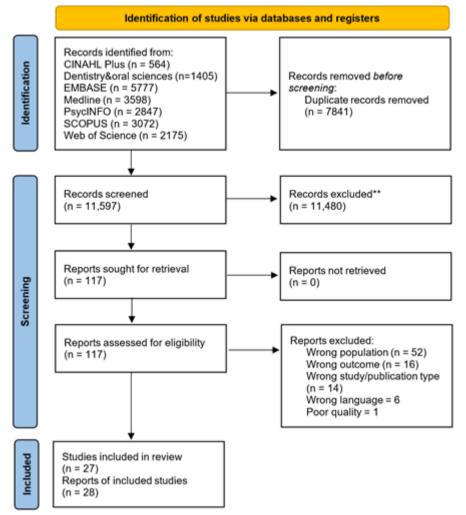


Figure 1 PRISMA flow chart detailing selection of the included studies. PRISMA, Preferred Reporting Items for Systematic Reviews and Meta-Analyses.

is indicated to develop a COS relating to the care of people presenting with acute dental pain or infection across healthcare settings internationally. Given the high rates of inappropriate antibiotic prescribing for people with acute dental conditions¹⁶³¹ and the increasing recognition of the important contribution dentistry can make to global efforts to tackle antibiotic resistance,⁷ this COS will be particularly important.

Measuring what matters to patients has been recognised as central to improving patient care and service delivery, with patients needing to be involved in decisions about what to measure.³² For this reason, experts by experience of urgent dental care were key members of our coproduction team, including when devising the review's search strategy. Funding to reimburse their time for participating in the length process of a systematic review was welcomed by the experts by experience.

The range of healthcare settings included in this review (dental clinics, pharmacies, hospital emergency departments and community clinics) mean the findings of this study are widely generalisable and can be easily translated to different healthcare settings around the world. Even though limited to English language, studies from a wide range of countries were included, across both highincome countries and LMICs. Six papers were excluded due to language (including 50% in Japanese) which may have introduced additional outcomes and differences in cultural practices.

Restricting this paper to published studies relating to adults from the peer-reviewed literature means that additional measures in the grey literature may have been missed as well as meaning that it fails to conform completely to the new Preferred Reporting Items for Systematic Reviews and Meta-Analyses 2020 guidelines for systematic reviews which were published during the course of our study.³³ The authors decided additional searches of the grey literature would not, however, meet the research questions or their intention to identify outcomes which had been successfully tried and tested. Studies including children were excluded from this review as the outcomes (especially patient-reported outcomes) are materially different.³⁴ Further, the trials for which these outcomes will be used by the authors relate to dental antibiotic stewardship and opioid stewardship for adult patients, which is the patient group where most overprescribing of these drugs occurs.^{35 36}

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The importance of valid, reliable and feasible measures for improving the quality of oral healthcare, including patient-reported outcomes and experience measures has been recognised.³⁷ In 2020, an international consensus of patient-centred outcomes to measure adult oral health (focusing on caries and periodontal disease) was published and highlighted that multiple measures are required to capture the effect of oral health on the individual patient.²² Where possible, we have adopted the terminology from this adult oral health standard set of outcomes when presenting our findings, such as 'complications or other harm resulting from treatment or disease progression' and 'unplanned visits.' However, while our findings cover some of the same territory, there are important differences in the detail especially relating to timescales. For example, there is no mention of 'infection' in the oral health outcomes and 'dental pain' covers only the frequency of pain in the last 6 months and 'complications' within 30 days, whereas our study found that these outcomes were measured in hours and days for people with acute dental conditions. Quality of life indicators such as the ability to eat, sleep, speak or carry out usual work activities at home and in the workplace (productivity) are outcomes from the standard oral health set which could be useful for studies of the outcome of care for people with acute dental pain and/ or infection but which were not employed in any of the studies within our review.²²

Primary medical care and to a lesser extent primary dental care have been recent targets of global efforts to tackle antibiotic resistance through stewardship programmes by reducing unnecessary and inappropriate prescribing.^{38 39} A hybrid umbrella/systematic review of measures to evaluate the effectiveness of antibiotic stewardship programmes, in primary medical and dental care respectively, found similar outcomes to this present review, including drug allergy, re-consultation rates and patient satisfaction.⁴⁰ Notably, the study about antibiotic stewardship measures found dental studies focused only on antibiotic use and the authors concluded that a range of metrics encompassing the wider measures employed in studies of medical care, including patientreported outcomes, should also be utilised in dentistry. Our findings reiterate this idea that a diverse range of outcomes should be used to evaluate care for people with acute dental conditions. Clinical outcomes such as signs and symptoms of pain and infection, and complications (including unplanned dental visits) should be employed in future studies, together with patient-reported measures such as satisfaction with the outcome of care.

Most studies in the review used unidimensional pain scales which are recognised to work well for acute pain: VAS, Heft-Parker scale, numeric rating scale and category pain scale.³⁴ Interestingly, none used the unidimensional pain scales based on images: Faces Pain Scale or Wong-Baker Faces Pain Scale.³⁴ Unsurprisingly none used the McGill Pain Scale or other multidimensional scales which are recognised to be more useful for chronic than acute BMJ Open: first published as 10.1136/bmjopen-2021-057934 on 21 February 2022. Downloaded from http://bmjopen.bmj.com/ on December 19, 2022 by guest. Protected by copyright

pain.³⁴ Future research to compare the utility of pain scales based on images with the other unidimensional pain scales for use in urgent dental care settings would be useful.

Dental antibiotic and opioid prescribing are recent priorities for clinicians and policymakers around the world, with overprescribing identified as a problem driving the development and spread of antibiotic resistance⁷ and substance misuse disorder,⁴¹ respectively. Prescribing rates and choices varying between countries, and solutions to tackle the problem of overprescribing need to be tailored to the local context.^{14 42} A recent pilot trial of a clinical decision prescribing tool and targeted education to improve dental antibiotic and opioid prescribing in Australia demonstrated a 41% reduction in antibiotic usage and 59% reduction in opioids.¹⁶ Clinical trials of antibiotic and opioid stewardship interventions are also planned in the UK⁴³ and USA.⁴⁴ Further research to develop a set of core outcomes for studies relating to the care of adults with acute dental pain and infection would be useful in the evaluation of stewardship interventions, to enable direct comparisons between stewardship interventions internationally.45

Standardising the reporting of metrics will facilitate improvements in the quality of care for people with acute dental pain and/or infection. The outcomes identified in this study (both clinical and patient reported) should form the basis on which to build international consensus on a COS as these measures will be useful in research, clinical and public health settings. Future research should be directed towards development and utilisation of this outcome set across healthcare settings where people with acute dental pain and infection present for treatment.

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REFERENCES

- Currie CC, Stone SJ, Durham J. Pain and problems: a prospective cross-sectional study of the impact of dental emergencies. J Oral Rehabil 2015;42:883–9.
- 2 Emmott R, Barber SK, Thompson W. Antibiotics and toothache: a social media review. *Int J Pharm Pract* 2021;29:210–7.
- 3 SDCEP. Management of acute dental problems during COVID-19 pandemic, 2020. Available: https://www.sdcep.org.uk/published-guidance/acute-dental-problems-covid-19/
- 4 Robertson DP, Keys W, Rautemaa-Richardson R, et al. Management of severe acute dental infections. *BMJ* 2015;350:h1300.
- 5 Worsley DJ, Robinson PG, Marshman Z. Access to urgent dental care: a scoping review. *Community Dent Health* 2017;34:19–26.
- 6 Lugtenberg M, Burgers J, Westert G. Effects of evidence-based clinical practice guidelines on quality of care: a systematic review. *BMJ Quality & Safety* 2009;18:385–92.
- 7 Thompson W, Williams D, Pulcini C. *The essential role of the dental team in reducing antibiotic resistance*. Geneva: FDI World Dental Federation, 2020.
- 8 Faculty of General Dental Practitioners (FGDP) U, Surgery FoD. Antimicrobial prescribing in dentistry: good practice guidelines. 3rd ed. London, UK: Royal College of Surgeons of England, 2020.
- 9 Lockhart PB, Tampi MP, Abt E, et al. Evidence-based clinical practice guideline on antibiotic use for the urgent management of pulpal- and periapical-related dental pain and intraoral swelling: a report from the American dental association. J Am Dent Assoc 2019;150:e12:906–21.
- 10 Bassey O, Csikar J, Hallam J, et al. Non-traumatic dental presentations at accident and emergency departments in the UK: a systematic review. Br Dent J 2020;228:171–6.
- 11 Cope AL, Wood F, Francis NA, et al. General practitioners' attitudes towards the management of dental conditions and use of antibiotics in these consultations: a qualitative study. BMJ Open 2015;5:e008551.
- 12 Amen TB, Kim I, Peters G, et al. Emergency department visits for dental problems among adults with private dental insurance: a national observational study. Am J Emerg Med 2021;44:166–70.
- 13 Shah S, Wordley V, Thompson W. How did COVID-19 impact on dental antibiotic prescribing across England? *Br Dent J* 2020;229:601–4.
- 14 Suda KJ, Durkin MJ, Calip GS, et al. Comparison of opioid prescribing by dentists in the United States and England. JAMA Netw Open 2019;2:e194303–e03.
- 15 Teoh L, Hollingworth S, Marino R, et al. Dental opioid prescribing rates after the up-scheduling of codeine in Australia. Sci Rep 2020;10:1–6.
- 16 Teoh L, Stewart K, Marino RJ. Improvement of dental prescribing practices using education and a prescribing tool: a pilot intervention study. Br J Clin Pharmacol 2020.
- 17 Kenny KP, Day PF, Sharif MO, *et al*. What are the important outcomes in traumatic dental injuries? an international approach to the development of a core outcome set. *Dent Traumatol* 2018;34:4–11.

- 18 World_Bank. World bank country and lending groups, 2021. Available: https://datahelpdesk.worldbank.org/knowledgebase/ articles/906519-world-bank-country-and-lending-groups
- 19 Critical Appraisal Skills Programme C. CASP randomised controlled trial standard checklist, 2020. Available: https://casp-uk.b-cdn.net/ wp-content/uploads/2020/10/CASP_RCT_Checklist_PDF_Fillable_ Form.pdf
- 20 Sirriyeh R, Lawton R, Gardner P, et al. Reviewing studies with diverse designs: the development and evaluation of a new tool. J Eval Clin Pract 2012;18:746–52.
- 21 Schweitzer VA, van Heijl I, van Werkhoven CH, et al. The quality of studies evaluating antimicrobial stewardship interventions: a systematic review. *Clin Microbiol Infect* 2019;25:555–61.
- 22 Ni Riordain R, Glick M, Al Mashhadani SSA. Developing a standard set of patient-centred outcomes for adult oral health-an international, cross-disciplinary consensus. *Int Dent J* 2020.
- 23 Culhane MC. Oral penicillin in the treatment of acute mandibular pericoronitis. *Oral Surg Oral Med Oral Pathol* 1947;33:B505–8.
- 24 Nusstein JM, Reader A, Beck M. Effect of drainage upon access on postoperative endodontic pain and swelling in symptomatic necrotic teeth. J Endod 2002;28:584–8.
- 25 Houck V, Reader A, Beck M, et al. Effect of trephination on postoperative pain and swelling in symptomatic necrotic teeth. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2000;90:507–13.
- 26 Henry M, Reader A, Beck M. Effect of penicillin on postoperative endodontic pain and swelling in symptomatic necrotic teeth. *J Endod* 2001;27:117–23.
- 27 Wilson K, Bouchiba M, Vithlani G, *et al.* Patient satisfaction with oral urgent treatment (out) in North West Tanzania. *Br Dent J* 2013;215:131–4.
- 28 Cohen LA, Bonito AJ, Akin DR, *et al*. Role of pharmacists in consulting with the underserved regarding toothache pain. *J Am Pharm Assoc* 2009;49:38–42.
- 29 Beus H, Fowler S, Drum M, et al. What is the outcome of an incision and drainage procedure in endodontic patients? A prospective, randomized, single-blind study. J Endod 2018;44:193–201.
- 30 Kirkham JJ, Gorst S, Altman DG, *et al.* Core outcome set– STAndards for reporting: the COS-STAR statement. *PLoS Med* 2016;13:e1002148.
- 31 Cope AL, Francis NA, Wood F, et al. Antibiotic prescribing in UK general dental practice: a cross-sectional study. Community Dent Oral Epidemiol 2016;44:145–53.
- 32 Calvert M, Kyte D, Price G, et al. Maximising the impact of patient reported outcome assessment for patients and society. BMJ 2019;364:k5267.
- 33 Page MJ, Moher D, Bossuyt PM, et al. PRISMA 2020 explanation and elaboration: updated guidance and exemplars for reporting systematic reviews. BMJ 2021;372:n160.
- 34 Sirintawat N, Sawang K, Chaiyasamut T, et al. Pain measurement in oral and maxillofacial surgery. J Dent Anesth Pain Med 2017;17:253–63.
- 35 Schroeder AR, Dehghan M, Newman TB, et al. Association of opioid prescriptions from dental clinicians for us adolescents and young adults with subsequent opioid use and abuse. JAMA Intern Med 2019;179:145–52.
- 36 Thompson W, Douglas G, Pavitt S. Factors associated with prescribing of systemic antibacterial drugs to adult patients in urgent primary health care, especially dentistry. J Antimicrob Chemother 2019.
- 37 Righolt AJ, Sidorenkov G, Faggion CM, et al. Quality measures for dental care: a systematic review. Community Dent Oral Epidemiol 2019;47:12–23.
- 38 Atkins L, Chadborn T, Bondaronek P, et al. Content and mechanism of action of national antimicrobial stewardship interventions on management of respiratory tract infections in primary and community care. Antibiotics 2020;9:512.
- 39 Löffler C, Böhmer F, Hornung A, et al. Dental care resistance prevention and antibiotic prescribing modification-the clusterrandomised controlled DREAM trial. *Implement Sci* 2014;9:27.
- 40 Teoh L, Sloan AJ, McCullough MJ, et al. Measuring antibiotic stewardship programmes and initiatives: an umbrella review in primary care medicine and a systematic review of dentistry. Antibiotics 2020;9:607.
- 41 Teoh L. Opioid prescribing in dentistry is there a problem? *Aust Prescr* 2020;43:144.
- 42 Thompson W, Teoh L, Hubbard CC, *et al*. Patterns of dental antibiotic prescribing in 2017: Australia, England, United States, and British Columbia (Canada). *Infect Control Hosp Epidemiol* 2021;2:1–8.
- 43 Goulao B, Scott C, Black I, *et al.* Audit and feedback with or without training in-practice targeting antibiotic prescribing (TiPTAP): a

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study protocol of a cluster randomised trial in dental primary care. *Implementation Science* 2021;16:1–9.

- 44 Durkin M. Using implementation science and informatics to develop and pilot test antibiotic stewardship clinical decision support: NIH, 2021.
- 45 Williamson PR, Altman DG, Bagley H, *et al*. The comet Handbook: version 1.0. *Trials* 2017;18:1–50.
- 46 Fazakerley MW, McGowan P, Hardy P, et al. A comparative study of cephradine, amoxycillin and phenoxymethylpenicillin in the treatment of acute dentoalveolar infection. *Br Dent J* 1993;174:359–63.
- 47 Gibson GB, Blasberg B, Altom R. A prospective survey of hospital ambulatory dental emergencies. Part 2: follow-up to emergency treatment. *Spec Care Dentist* 1993;13:110–2.
- 48 Fouad AF, Rivera EM, Walton RE. Penicillin as a supplement in resolving the localized acute apical abscess. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 1996;81:590–5.
- 49 Penniston SG, Hargreaves KM. Evaluation of periapical injection of ketorolac for management of endodontic pain. *J Endod* 1996;22:55–9.
- 50 Adriaenssen CF. Comparison of the efficacy, safety and tolerability of azithromycin and co-amoxiclav in the treatment of acute periapical abscesses. J Int Med Res 1998;26:257–65.
- 51 Doroschak AM, Bowles WR, Hargreaves KM. Evaluation of the combination of flurbiprofen and tramadol for management of endodontic pain. *J Endod* 1999;25:660–3.
- 52 Gallatin E, Reader A, Nist R, et al. Pain reduction in untreated irreversible pulpitis using an intraosseous injection of Depo-Medrol. J Endod 2000;26:633–8.
- 53 Nagle D, Reader A, Beck M, et al. Effect of systemic penicillin on pain in untreated irreversible pulpitis. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2000;90:636–40.
- 54 Hersh EV, DeRossi SS, Ciarrocca KN, et al. Efficacy and tolerability of an intraoral benzocaine patch in the relief of spontaneous toothache pain. J Clin Dent 2003;14:1–6.
- 55 Runyon MS, Brennan MT, Batts JJ, et al. Efficacy of penicillin for dental pain without overt infection. Acad Emerg Med 2004;11:1268–71.
- 56 Campanelli CA, Walton RE, Williamson AE, et al. Vital signs of the emergency patient with pulpal necrosis and localized acute apical abscess. J Endod 2008;34:264–7.

- 57 Sethi P, Agarwal M, Chourasia HR, *et al.* Effect of single dose pretreatment analgesia with three different analgesics on postoperative endodontic pain: a randomized clinical trial. *J Conserv Dent* 2014;17:517–21.
- 58 Pavithra P, Dhanraj M, Sekhar P. Analgesic effectiveness of Ibuprofen and Aceclofenac in the management of acute pulpitis - a randomized double blind trial. *Int J Pharm Sci Rev Res* 2015;35:70–4.
- 59 Bultema K, Fowler S, Drum M, *et al.* Pain reduction in untreated symptomatic irreversible pulpitis using liposomal bupivacaine (Exparel): a prospective, randomized, double-blind trial. *J Endod* 2016;42:1707–12.
- 60 Sebastian R, Drum M, Reader A, et al. What is the effect of No endodontic debridement on postoperative pain for symptomatic teeth with pulpal necrosis? J Endod 2016;42:378–82.
- 61 Santini MF, Rosa RAda, Ferreira MBC, et al. Comparison of two combinations of opioid and non-opioid analgesics for acute periradicular abscess: a randomized clinical trial. J Appl Oral Sci 2017;25:551–8.
- 62 Taggar T, Wu D, Khan AA. A randomized clinical trial comparing 2 ibuprofen formulations in patients with acute odontogenic pain. *J Endod* 2017;43:674–8.
- 63 Aaron S, Steier L. Efficacy of first aid treatment of acute apical abscess in an NHS emergency clinic. *British Dental Journal* 2018;224:523–7.
- 64 Eren B, Onay EO, Ungor M. Assessment of alternative emergency treatments for symptomatic irreversible pulpitis: a randomized clinical trial. *Int Endod J* 2018;51 Suppl 3:e227–37.
- 65 Wolf E, Dragicevic M, Fuhrmann M. Alleviation of acute dental pain from localised apical periodontitis: a prospective randomised study comparing two emergency treatment procedures. *J Oral Rehabil* 2019;46:120–6.
- 66 Al-Rawhani AH, Gawdat SI, Wanees Amin SA. Effect of diclofenac potassium premedication on Postendodontic pain in mandibular molars with symptomatic irreversible pulpitis: a randomized placebocontrolled double-blind trial. J Endod 2020;46:1023–31.
- 67 da Silva PB, Mendes AT, Cardoso MBF, *et al.* Comparison between isolated and associated with codeine acetaminophen in pain control of acute apical abscess: a randomized clinical trial. *Clin Oral Investig* 2021;25:875–82.