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[Intervention Review]

Single versus multiple visits for endodontic treatment of permanent teeth

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ABSTRACT

Background

Root canal treatment (RoCT), or endodontic treatment, is a common procedure in dentistry. The main indications for RoCT are irreversible pulpitis and necrosis of the dental pulp caused by carious processes, tooth cracks or chips, or dental trauma. Successful RoCT is characterised by an absence of symptoms (i.e. pain) and clinical signs (i.e. swelling and sinus tract) in teeth without radiographic evidence of periodontal involvement (i.e. normal periodontal ligament). The success of RoCT depends on a number of variables related to the preoperative condition of the tooth, as well as the endodontic procedures. This review updates the previous version published in 2007.

Objectives

To determine whether completion of root canal treatment (RoCT) in a single visit or over two or more visits, with or without medication, makes any difference in term of effectiveness or complications.

Search methods

We searched the following electronic databases: Cochrane Oral Health's Trials Register (to 14 June 2016), Cochrane Central Register of Controlled Trials (CENTRAL) (the Cochrane Library, 2016, Issue 5), MEDLINE Ovid (1946 to 14 June 2016), and Embase Ovid (1980 to 14 June 2016). We searched ClinicalTrials.gov and the [World Health Organization International Clinical Trials Registry Platform](http://WorldHealthOrganization.org/clinical-trials-registry) for ongoing trials to 14 June 2016. We did not place any restrictions on the language or date of publication when searching the electronic databases.

Selection criteria

We included randomised controlled trials (RCTs) and quasi-RCTs of people needing RoCT. We excluded surgical endodontic treatment. The outcomes of interest were tooth extraction for endodontic problems; radiological failure after at least one year, i.e. periapical radiolucency; postoperative pain; swelling or flare-up; painkiller use; sinus track or fistula formation; and complications (composite outcome including any adverse event).

Data collection and analysis

We collected data using a specially designed extraction form. We contacted trial authors for further details where these were unclear. We assessed the risk of bias in the studies using the Cochrane tool and we assessed the quality of the body of evidence using GRADE criteria. When valid and relevant data were collected, we undertook a meta-analysis of the data using the random-effects model. For dichotomous outcomes, we calculated risk ratios (RRs) and 95% confidence intervals (CIs). For continuous data, we calculated mean differences (MDs) and 95% CIs. We examined potential sources of heterogeneity. We conducted subgroup analyses for necrotic and vital teeth.

Main results

We included 25 RCTs in the review, with a total of 3780 participants, of whom we analysed 3751. We judged three studies to be at low risk of bias, 14 at high risk, and eight as unclear.

Only one study reported data on tooth extraction due to endodontic problems. This study found no difference between treatment in one visit or treatment over multiple visits (1/117 single-visit participants lost a tooth versus 2/103 multiple-visit participants; odds ratio (OR) 0.44, 95% confidence interval (CI) 0.04 to 4.78; very low-quality evidence).

We found no evidence of a difference between single-visit and multiple-visit treatment in terms of radiological failure (risk ratio (RR) 0.91, 95% CI 0.68 to 1.21; 1493 participants, 11 studies, $I^2 = 18\%$; low-quality evidence); immediate postoperative pain (dichotomous outcome) (RR 0.99, 95% CI 0.84 to 1.17; 1560 participants, 9 studies, $I^2 = 33\%$; moderate-quality evidence); swelling or flare-up incidence (RR 1.36, 95% CI 0.66 to 2.81; 281 participants, 4 studies, $I^2 = 0\%$; low-quality evidence); sinus tract or fistula formation (RR 0.98, 95% CI 0.15 to 6.48; 345 participants, 2 studies, $I^2 = 0\%$; low-quality evidence); or complications (RR 0.92, 95% CI 0.77 to 1.11; 1686 participants, 10 studies, $I^2 = 18\%$; moderate-quality evidence).

The studies suggested people undergoing RoCT in a single visit may be more likely to experience pain in the first week than those whose RoCT was over multiple visits (RR 1.50, 95% CI 0.99 to 2.28; 1383 participants, 8 studies, $I^2 = 54\%$), though the quality of the evidence for this finding is low.

Moderate-quality evidence showed people undergoing RoCT in a single visit were more likely to use painkillers than those receiving treatment over multiple visits (RR 2.35, 95% CI 1.60 to 3.45; 648 participants, 4 studies, $I^2 = 0\%$).

Authors' conclusions

There is no evidence to suggest that one treatment regimen (single-visit or multiple-visit root canal treatment) is better than the other. Neither can prevent all short- and long-term complications. On the basis of the available evidence, it seems likely that the benefit of a single-visit treatment, in terms of time and convenience, for both patient and dentist, has the cost of a higher frequency of late postoperative pain (and as a consequence, painkiller use).

PLAIN LANGUAGE SUMMARY

Should root canal treatment be performed in one dental visit or over several visits?

Review question

Is there any difference in effectiveness when undertaking root canal treatment in one visit compared to over several visits, and what are the effects on pain and complications, regardless of whether medication is used?

Background

This is an update of a review first published in 2007.

Root canal treatment, or endodontic treatment, is a common procedure in dentistry. The main reasons that root canal treatment are needed are persistent inflammation of the dental pulp (pulpitis) and death of the dental pulp (dead or non-vital tooth) caused by tooth decay, cracks or chips, or other accidental damage to teeth.

Root canal treatment is considered successful when there are no symptoms, for example pain, and when x-rays show no signs of damage to bone and other supporting tissues of the tooth. The success of root canal treatment depends on the preoperative condition of the tooth, as well as the endodontic procedures used.

Study characteristics

We searched the literature up to 14 June 2016. We found 25 relevant studies with a total of 3780 participants. The studies compared root canal treatment performed at a single appointment with root canal treatment performed over two or more appointments on vital permanent teeth, non-vital permanent teeth, or both.

Key results

No apparent difference exists between single- and multiple-visit root canal treatment on x-ray examination, an indicator which does not affect the patient directly, but is known to be important as a measure of effective treatment. Only one study measured the likelihood of tooth extraction due to endodontic problems and did not find evidence of a difference between single- and multiple-visit treatment. Most short- and long-term complications (pain, swelling, fistula, and tooth extraction) were similar in terms of frequency, although people undergoing a single visit were more likely to experience pain in the first week and to take painkillers.

Quality of the evidence

We assessed the available evidence as moderate to low quality because a number of the studies were at high risk of bias, there was inconsistency between study results, and results were imprecise.

SUMMARY OF FINDINGS FOR THE MAIN COMPARISON *[Explanation]*

Single-visit compared to multiple-visit treatment for endodontic treatment of permanent teeth						
Patient or population: people receiving endodontic treatment of permanent teeth Setting: university dental clinics, dental hospitals, and private dental practices Intervention: single-visit treatment Comparison: multiple-visit treatment						
Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	Number of participants (studies)	Quality of the evidence (GRADE)	Comments
	Risk with multiple-visit treatment	Risk with single-visit treatment				
Tooth extraction due to endodontic problems	19 per 1000	9 per 1000 (1 to 88)	RR 0.44 (0.04 to 4.78)	220 (1 RCT)	⊕○○○ VERY LOW ^{1,2,3}	
Radiological failure	146 per 1000	132 per 1000 (99 to 176)	RR 0.91 (0.68 to 1.21)	1493 (11 RCTs)	⊕⊕○○ LOW ^{4,5}	
Pain (dichotomous) - pain in the immediate postoperative period (until 72 hours postobturation)	379 per 1000	375 per 1000 (318 to 443)	RR 0.99 (0.84 to 1.17)	1560 (9 RCTs)	⊕⊕⊕○ MODERATE ⁴	
Pain (dichotomous) - pain at 1 week	109 per 1000	164 per 1000 (108 to 249)	RR 1.50 (0.99 to 2.28)	1383 (8 RCTs)	⊕⊕○○ LOW ^{4,6}	
Swelling or flare-up	81 per 1000	110 per 1000 (53 to 226)	RR 1.36 (0.66 to 2.81)	281 (4 RCTs)	⊕⊕○○ LOW ^{4,5}	
Painkiller use	97 per 1000	228 per 1000 (155 to 335)	RR 2.35 (1.60 to 3.45)	648 (4 RCTs)	⊕⊕⊕○ MODERATE ⁴	

Sinus track or fistula formation	12 per 1000	12 per 1000 (2 to 78)	RR 0.98 (0.15 to 6.48)	345 (2 RCTs)	⊕⊕○○ LOW ^{4,7}
Any complication	286 per 1000	263 per 1000 (220 to 317)	RR 0.92 (0.77 to 1.11)	1686 (10 RCTs)	⊕⊕⊕○ MODERATE ⁴

***The risk in the intervention group** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; **RCT:** randomised controlled trial; **RR:** risk ratio.

GRADE Working Group grades of evidence

High quality: we are very confident that the true effect lies close to that of the estimate of the effect.

Moderate quality: we are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low quality: our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect.

Very low quality: we have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

¹ The study was judged at high risk of bias.

² The results are based on a single study performed in a university centre.

³ The results are based on an single study, with relatively few participants and events.

⁴ The results are based on a number of studies judged at high risk of bias.

⁵ The results showed significant imprecision.

⁶ The studies showed significant heterogeneity.

⁷ The results are based on two studies only, with relatively few participants and events.

BACKGROUND

Description of the condition

Irreversible pulpitis of the dental pulp and its subsequent necrosis (death of dental pulp) are caused by carious processes, tooth cracks or chips, or dental trauma and represent the main indications for root canal treatment (RoCT). The primary symptom of irreversible pulpitis is severe pain, and RoCT is necessary to relieve symptoms and to avoid tooth extraction due to complications following necrosis of the dental pulp.

Description of the intervention

RoCT is a common procedure in dentistry that is performed to remove organic tissue, infected debris, and pathogenic bacteria from the root canal system by means of mechanical instrumentation associated with copious irrigation with disinfectant agents. Two approaches have been proposed to solve this problem. In the first approach, residual bacteria are eliminated or prevented from repopulating the root canal system by introducing an interappointment dressing into the root canal, generally falling into the following categories: phenolic derivatives (eugenol, camphorated para-monochlorophenol, camphorated phenol, metacresyl acetate, beechwood creosote), aldehydes (formocresol), halides (iodine-potassium iodide), calcium hydroxide, antibiotics, or other combinations. The most popular intracanal medication currently in use is calcium hydroxide. Some studies have shown that calcium hydroxide fails to produce sterile root canals and even allows regrowth in some cases (Kvist 2004; Orstavik 1991; Reit 1988). However, even a negative culture before obturation gives no guarantee of healing in all cases (Trope 1999; Weiger 2000). The second approach aims to eliminate remaining bacteria or render them harmless by entombing them by complete and three-dimensional obturation, finishing the treatment in one visit to deprive the micro-organisms of nutrition and the space required to survive and multiply (Soltanoff 1978; Weiger 2000). The antimicrobial activity of the sealer or the zinc ions of gutta-percha can kill the residual bacteria (Moorer 1982; Siqueira 2000).

Endodontic techniques can claim many improvements through the use of rubber dam, magnifying devices, electronic apex locators, and engine-driven rotary nickel titanium files, which have improved the success rate of endodontic treatment and shortened the time needed for treatment (Bystrom 1981; Orstavik 1998; Reit 1988). The basic biological rationale for achieving final success of RoCT consists primarily of eliminating micro-organisms from the entire root canal system and preventing their re-entry. Different therapeutic procedures can be employed, depending upon the biological condition of the tooth being treated, its pathological state, clinician expertise, instrument availability, and patient preference. Successful RoCT is characterised by the absence of symptoms and

clinical signs of infection in a tooth without radiographic evidence of periodontal involvement (Friedman 2002). The success of RoCT depends on variables related to the preoperative condition of the tooth, as well as the endodontic procedures.

How the intervention might work

RoCT can be followed by numerous short- and long-term complications (Battrum 1996). The former include immediate postoperative inflammation of periradicular tissues associated with pain, either spontaneous or provoked. The correlation of postoperative pain with different variables, including the number of visits needed to complete RoCT, operative procedures, pulp vitality and dental anatomy, has been the objective of numerous studies (Albashaireh 1998; DiRenzo 2002; Gambarini 1991; Soltanoff 1978). The main long-term complications include the persistence of inflammation of fistula or sinus track, pain, or both, and an absence of radiographic healing. Several studies have investigated the frequency of radiographic healing in teeth with preoperative periapical pathology and have compared single- and multiple-visit approaches, employing interappointment medication (Katebzadeh 2000; Peters 2002; Soltanoff 1978; Trope 1999; Weiger 2000). The results of such investigations have led to conflicting conclusions. Some studies have suggested that the use of different medications between visits can contribute to the elimination of all bacteria (Fava 1995). In contrast, others have emphasised the need to seal the endodontic space as quickly as possible, i.e. in a single visit, as temporary cements are unreliable in maintaining a good coronal seal during the time between visits. Postoperative complications have been reported with both methods, varying from 5% in Abbott 2000 to more than 20% in Friedman 1995.

Why it is important to do this review

Cochrane Oral Health undertook an extensive prioritisation exercise in 2014 to identify a core portfolio of titles that were the most clinically important ones to maintain in the Cochrane Library (Worthington 2015). The operative and prosthodontic dentistry expert panel identified this review as a priority title (Cochrane Oral Health priority review portfolio).

RoCT is an extremely common procedure, performed on thousands of people every day. If a single-visit approach is shown to be as effective and safe as multiple-visit treatment, it could result in a significant saving in time and money for patients and dentists.

OBJECTIVES

To determine whether completion of root canal treatment (RoCT) in a single visit or over two or more visits, with or without medi-

cation, makes any difference in term of effectiveness or complications.

METHODS

Criteria for considering studies for this review

Types of studies

We included randomised controlled trials (RCTs) and quasi-RCTs (i.e. those using an alternative assignment based on, for example, birth date). We also considered split-mouth studies. We excluded studies that did not measure at least one of our outcomes.

Types of participants

Participants aged 10 years or above who required root canal treatment. All participants had teeth with a completely formed apex and without internal resorption.

Types of interventions

Root canal treatment in a single visit or multiple visits, i.e. two or more appointments.
Any systemic medical treatment (antibiotics, non-steroidal anti-inflammatories or analgesics) was to be the same in both groups.

Types of outcome measures

Primary outcomes

- Tooth extraction due to endodontic problems (binary, yes/no).
- Radiological failure after at least one year, i.e. the presence of any periapical radiolucency (binary, yes/no). Additional [Table 1](#) summarises how we adapted the most common scales of radiological healing to a binary outcome.

Secondary outcomes

- Postoperative pain (binary, yes/no; continuous).
- Swelling or flare-up (binary, yes/no).
- Painkiller use (binary, yes/no).
- Sinus track or fistula formation (binary, yes/no).
- Any complication defined as a composite outcome including any adverse event (pain, painkiller use, swelling or flare-up) occurring within one month from the treatment (binary, yes/no).

Search methods for identification of studies

To identify studies for this review, we developed detailed search strategies for each database searched. These were based on the search strategy developed for MEDLINE Ovid ([Appendix 3](#)), but revised appropriately for each database.

Electronic searches

We searched the following electronic databases.

- Cochrane Oral Health Group's Trials Register (to 14 June 2016) (see [Appendix 1](#)).
- Cochrane Central Register of Controlled Trials (CENTRAL; 2015, Issue 5) in the Cochrane Library (searched 14 June 2016) (see [Appendix 2](#)).
- MEDLINE Ovid (1946 to 14 June 2016) (see [Appendix 3](#)).
- Embase Ovid (1980 to 14 June 2016) (see [Appendix 4](#)).

We did not place any restrictions on the language or date of publication when searching the electronic databases.

Searching other resources

We searched the following trial registries for ongoing studies (see [Appendix 5](#)).

- US National Institutes of Health Ongoing Trials Register ClinicalTrials.gov (clinicaltrials.gov; searched 14 June 2016).
- World Health Organization International Clinical Trials Registry Platform (apps.who.int/trialsearch; searched 14 June 2016).

We did not perform a separate search for adverse effects of interventions; we considered adverse effects as described in included studies only.

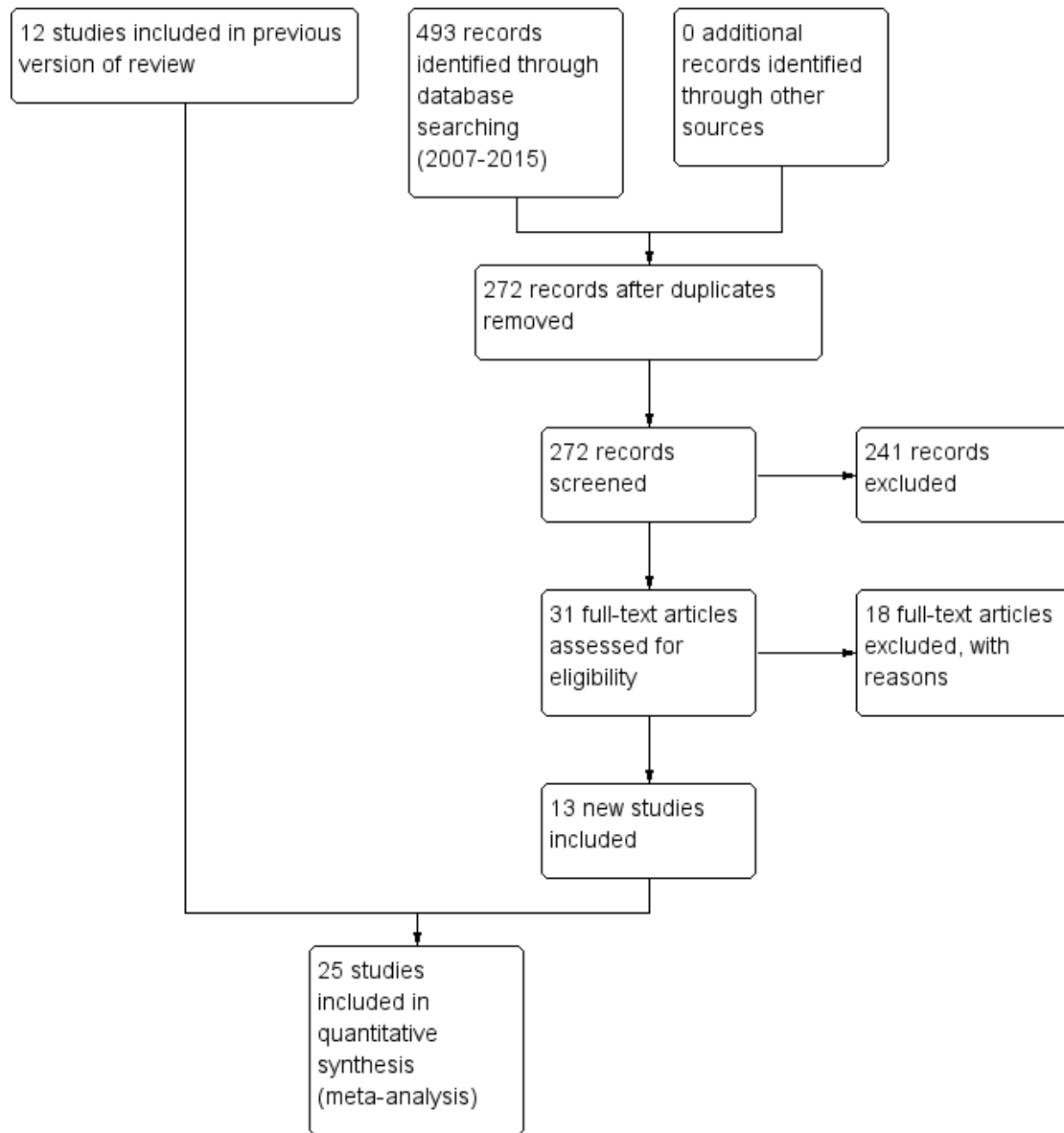
We checked all references in the identified papers and we contacted the authors to identify any additional published or unpublished data.

Data collection and analysis

Selection of studies

Two review authors (MM and LF) independently examined the title and abstract (when available) of each article identified by the search strategy. Where studies appeared to meet the inclusion criteria for this review or where there were insufficient data in the title and abstract to make a clear decision, we obtained the full report. The full report was then assessed by at least two of the review authors (MM, LF, GL, MG), to determine whether studies met the inclusion criteria. We resolved any disagreements by discussion. We recorded studies rejected at this or subsequent stages in the [Characteristics of excluded studies](#) table, with the reason(s) for exclusion. See [Figure 1](#) for a flow chart that summarises the results of the search.

Figure 1. Study flow diagram: review update



Data extraction and management

At least two review authors (MM, LF) independently extracted data using a specially designed data extraction form and entered

them into a spreadsheet. At least two review authors (GL, MM) authors independently assessed the risk of bias in the included studies. We discussed any discrepancies to reach agreement. When necessary, we contacted study authors for clarification or missing

information. For each trial, we recorded the following data.

- Year of publication, country of origin, number of centres, source of study funding, recruitment period.
- Details of the participants including demographic characteristics and criteria for inclusion and exclusion, type of tooth treated (vital or non-vital) and reasons of the treatment, diagnostic criteria for pulpal or periapical disease, numbers randomised to each treatment group.
- Details about the number of visits performed to treat the root canal (single or multiple), number of operators involved in the treatment, use of rubber dam and magnification loupes, type of canal shaping, type of irrigation, method used to establish the working length of the root canal, type of obturation of the canal. In the multiple visits approach, we also recorded the type of medication used in the interappointment period.
 - Details of pain management.
 - Details of the outcomes reported, including method of assessment, and time(s) assessed.
 - Description of operators.

Assessment of risk of bias in included studies

Two review authors (GL, MM) independently assessed the risk of bias of the included trials and any disagreement was resolved through discussion and consensus. We used the recommended

approach for assessing risk of bias in studies included in Cochrane Reviews ([Higgins 2011](#)). We addressed six domains:

- random sequence generation (selection bias);
- allocation concealment (selection bias);
- blinding of outcome assessment (detection bias);
- incomplete outcome data (attrition bias);
- selective reporting (reporting bias);
- other bias.

Each domain in the tool includes one or more entries in the 'Characteristics of included studies' table. Within each entry, we described what was reported to have happened in the study, in sufficient detail to support a judgement about the risk of bias. We then assigned a judgement relating to the risk of bias for that entry, either 'low', 'high', or 'unclear' risk of bias. After taking into account the additional information provided by the authors of the trials, we summarised the risk of bias in the studies as:

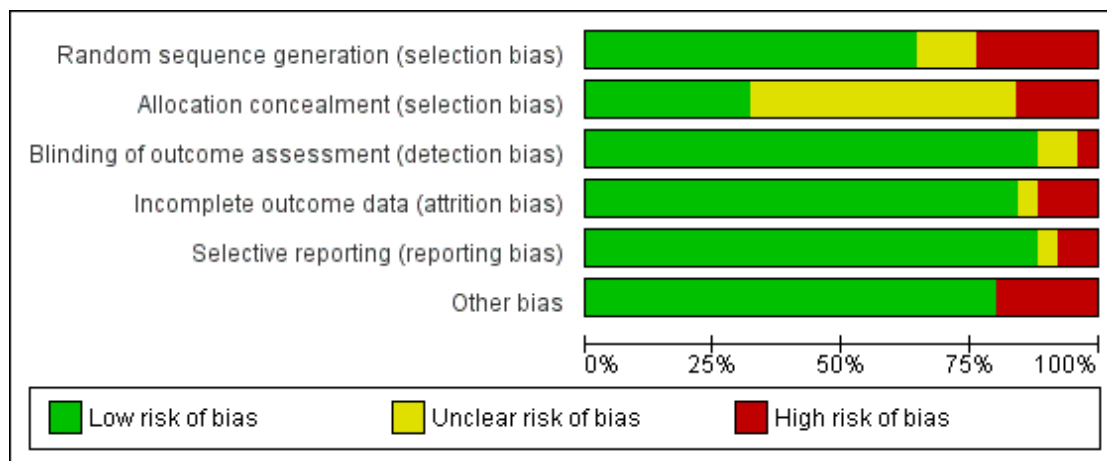
- low risk of bias = low risk of bias for all key domains;
- unclear risk of bias = unclear risk of bias for one or more key domains;
- high risk of bias = high risk of bias for one or more key domains.

We completed a 'Risk of bias' table for each included study (see [Characteristics of included studies](#)), and presented results graphically by study and by domain across all studies ([Figure 2](#); [Figure 3](#)).

Figure 2. 'Risk of bias' summary: review authors' judgements about each risk of bias item for each included study

	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of outcome assessment (detection bias)	Incomplete outcome data (attrition bias)	Selective reporting (reporting bias)	Other bias
Akbar 2013	+	?	+	+	+	+
Albashaireh 1998	-	-	+	+	+	+
Al-Negrish 2006	-	-	+	+	+	+
DiRenzo 2002	+	?	+	+	+	+
Dorasani 2013	-	?	?	-	+	-
Gesi 2006	+	?	+	+	+	+
Ghoddusi 2006	+	?	+	+	+	+
Ince 2009	?	?	+	+	+	+
Molander 2007	+	+	+	+	+	+
Mulhern 1982	?	?	+	+	+	+
Oginni 2004	?	?	+	?	+	-
Paredes-Vieyra 2012	+	?	+	+	+	-
Patil 2016	+	+	-	+	-	+
Penenis 2008	+	+	+	-	+	+
Peters 2002	-	-	+	+	+	+
Rao 2014	-	?	+	+	?	+
Risso 2008	+	?	+	+	-	+
Singh 2012	+	+	+	+	+	+
Soltanoff 1978	+	?	?	+	+	+
Trope 1999	+	+	+	+	+	-
Wang 2010	+	+	+	+	+	+
Weiger 2000	+	?	+	+	+	+
Wong 2015	+	+	+	-	+	+
Xiao 2010	+	+	+	+	+	-
Yoldas 2004	-	-	+	+	+	+

Figure 3. 'Risk of bias' graph: review authors' judgements about each risk of bias item presented as percentages across all included studies



Measures of treatment effect

Our primary measures of intervention effect were tooth extraction due to endodontic problems and radiological failure after one year. We dichotomised data on radiological healing when this was measured on ordinal scales (see Table 1 for details). Our other outcomes were incidence of postoperative pain, presence of swelling or flare-up, painkiller use, sinus track or fistula formation, and any complication. We analysed dichotomous data by calculating risk ratios (RRs) and 95% confidence intervals (CIs). When pain was recorded as a continuous outcome, we analysed data by calculating mean differences (MDs) and 95% CIs.

Unit of analysis issues

The statistical unit was the individual participant. We considered studies using the tooth as the statistical unit, making the assumption that participants were randomised a number of times equal to the number of teeth. When this was not explicitly stated, we considered the study potentially affected by a bias of allocation.

Dealing with missing data

When raw data were not available, we obtained them by consulting tables and graphs, or by contacting the trial authors.

Assessment of heterogeneity

We assessed heterogeneity by inspection of the point estimates and CIs on the forest plots. We assessed the variation in treatment effects by means of Cochran's test for heterogeneity and quantified by the I^2 statistic. We considered heterogeneity statistically significant if $P < 0.1$. A rough guide to the interpretation of the I^2 statistic given in the *Cochrane Handbook for Systematic Reviews of Interventions* is: 0% to 40% might not be important, 30% to 60% may represent moderate heterogeneity, 50% to 90% may represent substantial heterogeneity, 75% to 100% may represent very substantial ('considerable') heterogeneity (Higgins 2011).

Assessment of reporting biases

Only a proportion of research projects conducted are ultimately published in an indexed journal and become easily identifiable for inclusion in systematic reviews. Reporting biases arise when the reporting of research findings is influenced by the nature and direction of the findings of the research. We attempted to minimise potential reporting biases including publication bias, time lag bias, multiple (duplicate) publication bias, and language bias in this review. If there had been more than 10 studies evaluating one outcome, we planned to construct a funnel plot. If there had been asymmetry in the funnel plot indicating possible publication bias, we planned to undertake statistical analysis using the methods introduced by Egger 1997 (continuous outcome) and Rucher

2008 (dichotomous outcome). We attempted to avoid time lag bias, multiple (duplicate) publication bias, and language bias by conducting a detailed sensitive search, including searching for ongoing studies. We did not impose any restrictions on language, and we found translators for potentially relevant trials published in other languages.

Data synthesis

For each intervention, we sought and summarised data on the number of participants in intervention and control groups who experienced the event (outcome) and the total number of participants. We only conducted a meta-analysis if there were studies of similar comparisons reporting the same outcome measures. We combined RRs for dichotomous data, and MDs for continuous data, using a random-effects model in Review Manager software (RevMan 2014).

Subgroup analysis and investigation of heterogeneity

We had planned subgroup analyses to investigate the relevance of pretreatment conditions (vital teeth versus necrotic teeth), pretreatment symptoms (symptomatic versus asymptomatic teeth), pretreatment radiographic periapical appearance (apical radiolucency versus no apical radiolucency), endodontic technique, and antimicrobials employed (antimicrobial A versus antimicrobial B). Only data on pretreatment conditions (vital teeth versus necrotic teeth) were available for a subgroup analysis.

Sensitivity analysis

We performed a sensitivity analysis, excluding studies at high risk of bias.

Presentation of main results

Using GRADEpro GDT software (GRADEpro GDT 2014), we produced Summary of findings table 1 for all outcomes. We assessed the quality of the evidence as high, moderate, low, or very low, with reference to study limitations, inconsistency, imprecision, indirectness, and risk of publication bias (Atkins 2004). We explained decisions to downgrade the quality of evidence using footnotes in the 'Summary of findings' table.

RESULTS

Description of studies

Results of the search

We found 25 studies suitable for inclusion in this review (see Figure 1).

Included studies

Characteristics of trial design and settings

For a summary of the characteristics of each of the included studies, see Characteristics of included studies.

Of the 25 included studies, five were conducted in the USA (DiRenzo 2002; Mulhern 1982; Penenis 2008; Soltanoff 1978; Trope 1999), four in India (Dorasani 2013; Patil 2016; Rao 2014; Singh 2012), two in Jordan (Albashaireh 1998; Al-Negrish 2006), three in China (Wang 2010; Wong 2015; Xiao 2010), two in Turkey (Ince 2009; Yoldas 2004), and single studies were conducted in Italy (Gesi 2006), Iran (Ghoddusi 2006), Sweden (Molander 2007), Nigeria (Oginni 2004), Mexico (Paredes-Vieyra 2012), the Netherlands (Peters 2002), Brazil (Risso 2008), Germany (Weiger 2000), and Pakistan (Akbar 2013).

Twenty-two of the studies were performed in university clinics or hospitals. Gesi 2006 was undertaken in private practice. Two studies did not provide details about the settings of the study (Mulhern 1982; Rao 2014).

All the studies used parallel group designs and most had two treatment arms. Two studies had three arms, which compared a single visit, multiple visits without intracanal medication, and multiple visits with intracanal medication (calcium hydroxide) (Ghoddusi 2006; Trope 1999). In order to include such data in the meta-analysis, we combined the two multi-visit arms. In the same meta-analysis, we pooled data from studies that used or did not use a dressing.

Four studies reported that they had received research grants for the conduct of the study (Ghoddusi 2006; Mulhern 1982; Penenis 2008; Risso 2008).

Characteristics of participants

A total of 3780 participants were enrolled in the 25 studies included in this review, and a total of 3571 participants were analysed.

All studies considered one tooth per participant, with the exception of Dorasani 2013, Oginni 2004, Paredes-Vieyra 2012, Trope 1999, Wong 2015, and Xiao 2010. Dorasani 2013 considered 64 teeth in 43 participants (5 participants contributed 2 teeth and 1 participant contributed 3 teeth). In Oginni 2004, participants requiring root canal treatment (RoCT) on more than one tooth, underwent consecutive treatment of each tooth with an interval of at least four weeks to allow proper evaluation: 283 teeth were randomised in 255 participants. In Paredes-Vieyra 2012, the study authors reported that 287 participants and 300 teeth were enrolled; they stated that 21 participants contributed more than one tooth, but according to these figures, no more than 13 participants could

have contributed more than one tooth. Trope 1999 considered 102 teeth in 81 participants (61 participants had a single tooth, 18 had 2 teeth and 2 participants had 3 teeth). Finally in Xiao 2010, 86 participants were enrolled with a total of 138 teeth. For these five studies, the analysis was conducted at the level of the tooth. Fourteen studies included participants with necrotic teeth only (Akbar 2013; Al-Negrish 2006; Dorasani 2013; Ghoddsu 2006; Molander 2007; Mulhern 1982; Paredes-Vieyra 2012; Penenis 2008; Peters 2002; Rao 2014; Risso 2008; Trope 1999; Weiger 2000; Xiao 2010). Yoldas 2004 is the only study that included retreatment, while Gesi 2006 and Wang 2010 included only participants with vital teeth. Seven studies included both necrotic teeth and vital teeth (Albashaireh 1998; DiRenzo 2002; Ince 2009; Oginni 2004; Patil 2016; Singh 2012, Wong 2015), but two of them did not provide details on the numbers in the two categories (DiRenzo 2002; Oginni 2004). One study did not provide details on the pretreatment status (Soltanoff 1978). One study included maxillary central incisors only (Patil 2016). None of the data reported by the included studies allowed us to analyse the effects of the two approaches stratified on the basis of preoperative conditions of patients, specifically presence of symptoms (pain) or signs (infection).

Characteristics of interventions

Of the 25 studies included in this review, the majority compared RoCT performed in a single visit with RoCT performed in two visits (Akbar 2013; Al-Negrish 2006; Albashaireh 1998; DiRenzo 2002; Dorasani 2013; Gesi 2006; Ghoddsu 2006; Ince 2009; Molander 2007; Paredes-Vieyra 2012; Patil 2016; Penenis 2008; Peters 2002; Rao 2014; Singh 2012; Trope 1999; Weiger 2000; Wang 2010; Yoldas 2004). In one study, the multiple-visit RoCT lasted three visits (Mulhern 1982). One study (Wong 2015) reported that multiple visit treatment had been performed in two or three appointments depending on the complexity of the RoCT. In two studies, the number of visits was not specified (Oginni 2004; Soltanoff 1978).

In the multiple-visit approach, nine studies did not use any intracanal medications in the interappointment period (Albashaireh 1998; DiRenzo 2002; Ghoddsu 2006; Ince 2009; Mulhern 1982; Patil 2016; Rao 2014; Singh 2012; Trope 1999). In 12 studies, the root canals were medicated with a calcium hydroxide paste (Akbar 2013; Al-Negrish 2006; Dorasani 2013; Gesi 2006; Molander 2007; Paredes-Vieyra 2012; Peters 2002; Risso 2008; Wang 2010; Weiger 2000; Wong 2015; Xiao 2010), while two studies used a medication with calcium hydroxide and chlorhexidine (Penenis 2008; Yoldas 2004). Two studies did not specify the type of interappointment medication (Oginni 2004; Soltanoff 1978).

Eleven studies did not report use of rubber dam to isolate the tooth during RoCT (Albashaireh 1998; Ince 2009; Molander 2007; Oginni 2004; Rao 2014; Risso 2008; Singh 2012; Soltanoff 1978; Wang 2010; Xiao 2010; Yoldas 2004), and use of magnification

loupes was reported by three studies only (Penenis 2008; Peters 2002; Wong 2015).

Working length was established using an electronic apex locator and radiographs in seven studies (DiRenzo 2002; Paredes-Vieyra 2012; Patil 2016; Peters 2002; Rao 2014; Risso 2008; Yoldas 2004), by an electronic apex locator only in four studies (Penenis 2008; Singh 2012; Wang 2010; Xiao 2010), and by radiographs only in nine studies (Akbar 2013; Al-Negrish 2006; Dorasani 2013; Gesi 2006; Ghoddsu 2006; Ince 2009; Mulhern 1982; Trope 1999; Weiger 2000). Five studies did not report information about working length evaluation (Albashaireh 1998; Molander 2007; Oginni 2004; Soltanoff 1978; Wong 2015).

Canal shaping was performed using conventional hand instruments in 10 studies (Akbar 2013; Al-Negrish 2006; Dorasani 2013; Gesi 2006; Ghoddsu 2006; Ince 2009; Peters 2002; Risso 2008; Soltanoff 1978; Weiger 2000), a combination of hand files and nickel titanium rotary files in seven studies (DiRenzo 2002; Molander 2007; Paredes-Vieyra 2012; Rao 2014; Singh 2012; Wang 2010; Yoldas 2004), and rotary nickel titanium files only in four studies (Patil 2016; Penenis 2008; Wong 2015; Xiao 2010). Four studies did not report canal shaping instrumentation in a satisfactory way (Albashaireh 1998; Mulhern 1982; Oginni 2004; Trope 1999).

Twenty-four studies reported that canal obturation was performed with gutta-percha using the lateral condensation technique; only one study reported use of the vertical condensation technique (Penenis 2008).

In 21 studies, sodium hypochlorite with a range between 0.5% to 5.25% was used as irrigant (Akbar 2013; Al-Negrish 2006; Albashaireh 1998; DiRenzo 2002; Dorasani 2013; Gesi 2006; Ince 2009; Molander 2007; Mulhern 1982; Paredes-Vieyra 2012; Patil 2016; Penenis 2008; Peters 2002; Rao 2014; Singh 2012; Trope 1999; Wang 2010; Weiger 2000; Wong 2015; Xiao 2010; Yoldas 2004), while two studies used saline solution as irrigant (Ghoddsu 2006; Soltanoff 1978). In Risso 2008, a combination of sodium hypochlorite, citric acid, and sodium thiosulfate was used as irrigant. In Oginni 2004, the type of irrigant used was not specified.

Nine studies reported that RoCT was performed by a single operator (Al-Negrish 2006; Albashaireh 1998; Dorasani 2013; Ghoddsu 2006; Paredes-Vieyra 2012; Patil 2016; Peters 2002; Risso 2008; Trope 1999), while 10 studies reported two or more operators were involved in RoCT (DiRenzo 2002; Gesi 2006; Ince 2009; Molander 2007; Mulhern 1982; Penenis 2008; Wang 2010; Weiger 2000; Wong 2015; Yoldas 2004). Six studies specified that treatment was conducted by trained or experienced operators (Ince 2009; Molander 2007; Penenis 2008; Peters 2002; Wang 2010; Weiger 2000), while three studies reported that RoCT was performed by postgraduate students (DiRenzo 2002; Mulhern 1982; Risso 2008). Six studies did not provide details about the operators (Akbar 2013; Oginni 2004; Rao 2014; Singh 2012; Soltanoff 1978; Xiao 2010).

Characteristics of outcomes

One study only provided information on tooth extraction due to endodontic problems as outcome measure (Wong 2015).

Radiological failure was investigated in 11 studies (Dorasani 2013; Gesi 2006; Molander 2007; Paredes-Vieyra 2012; Penenis 2008; Peters 2002; Soltanoff 1978; Trope 1999; Weiger 2000; Wong 2015; Xiao 2010). Additional Table 1 shows methods adopted to construct scales for radiological healing or failure. Follow-up varied from one year in Trope 1999 to five years in Weiger 2000. Seventeen studies investigated postoperative pain (Al-Negrish 2006; Albashaireh 1998; DiRenzo 2002; Gesi 2006; Ghoddusi 2006; Ince 2009; Mulhern 1982; Oginni 2004; Patil 2016; Rao 2014; Risso 2008; Singh 2012; Soltanoff 1978; Wang 2010; Wong 2015; Xiao 2010; Yoldas 2004). The methods for evaluating postoperative pain are summarised in the 'Characteristics of included studies' table. Whenever possible, we dichotomised pain data into 'pain' or 'no pain' values. Four studies recorded pain as a continuous variable measured by visual analogue scale (VAS) (DiRenzo 2002; Patil 2016; Singh 2012; Wang 2010). We only considered pain after canal obturation, assessing pain incidence in the immediate postobturation period (until 72 hours), at one week, and at one month. We did not consider pain during the interappointment period in the multiple-visit approach, as we could not compare this with a similar situation in the single-visit approach. Five studies examined the need for participants to take analgesics to relieve pain (Mulhern 1982; Rao 2014; Soltanoff 1978; Wang 2010; Yoldas 2004).

Eight studies investigated the incidence of swelling or flare-up (Akbar 2013; DiRenzo 2002; Ghoddusi 2006; Mulhern 1982; Paredes-Vieyra 2012; Penenis 2008; Rao 2014; Wang 2010). Definitions of flare-up varied (see Table 2). Only DiRenzo 2002, Mulhern 1982, Ghoddusi 2006, and Akbar 2013 clearly defined flare-up as swelling. Therefore, we considered only studies clearly indicating swelling as a specific outcome, not simply as one of the signs related to the tooth infection.

Four studies provided information on fistula or sinus track formation as an outcome measure (Penenis 2008; Paredes-Vieyra 2012; Wong 2015; Xiao 2010).

Excluded studies

Our main reason for excluding each of the 20 studies is recorded in the 'Characteristics of excluded studies' tables.

We excluded 10 studies because they were not RCTs (ElMubarak 2010; Fava 1989; Fava 1994; Friedman 1995; Jabeen 2014; Ng 2004; Prashanth 2011; Roane 1983; Walton 1992; Jabeen 2014). We excluded five studies because they evaluated only the microbiological status of RoCT (Gurgel-Filho 2007; Kvist 2004; Trusewicz 2005; Vera 2012; Xavier 2013), and two because the studies were conducted in animals (Holland 2003; Silveira 2007). We excluded two studies because they included primary teeth (Orhan 2010; Kabaktchieva 2013). We excluded one study because it did not

include any of the outcomes considered in this review (Waltimo 2005).

Risk of bias in included studies

On the basis of criteria used in the critical appraisal of the studies, we assessed three studies as being at overall low risk of bias (Molander 2007; Singh 2012; Wang 2010); eight studies as being at unclear risk of bias (Akbar 2013; DiRenzo 2002; Gesi 2006; Ghoddusi 2006; Ince 2009; Mulhern 1982; Soltanoff 1978; Weiger 2000); and the remaining studies as being at high risk of bias (Albashaireh 1998; Al-Negrish 2006; Dorasani 2013; Oginni 2004; Paredes-Vieyra 2012; Patil 2016; Penenis 2008; Peters 2002; Rao 2014; Risso 2008; Trope 1999; Wong 2015; Xiao 2010; Yoldas 2004). See Figure 2.

Allocation

We assessed the generation of randomisation sequence as being at low risk of bias in 16 trials (64%), unclear risk in three trials (12%), and high risk in six trials (24%). The concealment of allocation was at low risk of bias in eight trials (32%), unclear risk in 13 (52%), and high risk in four (16%). See Figure 3.

Blinding

We assessed blinding of outcome assessment as being at low risk of detection bias for 22 trials (88%), unclear for two trials (8%), and high risk for one trial (4%) (Patil 2016).

Incomplete outcome data

The reported dropout rate ranged from 0% to 35% (Penenis 2008). We assessed 21 (84%) trials as being at low risk with regard to attrition bias, either due to no dropouts or dropouts being unlikely to influence findings. For one study (4%), it was not possible to assess such bias. For three trials, we considered the high dropout rates to put the studies at high risk of attrition bias (12%).

Selective reporting

Two trials (8%) did not report one of the outcomes planned and for this reason we assessed them as being at high risk of reporting bias. For another trial, this bias was impossible to assess (4%). We assessed the remaining 22 studies (88%) as being at low risk of bias.

Other potential sources of bias

We judged five studies (20%) as being at high risk of other bias because they did not explicitly state that participants with more that

one tooth needing treatment were randomised the same number of times.

Effects of interventions

See: [Summary of findings for the main comparison](#) Single visit versus multiple visit treatment for endodontic treatment of permanent teeth

Single visit versus multiple visit root canal treatment

Primary outcomes

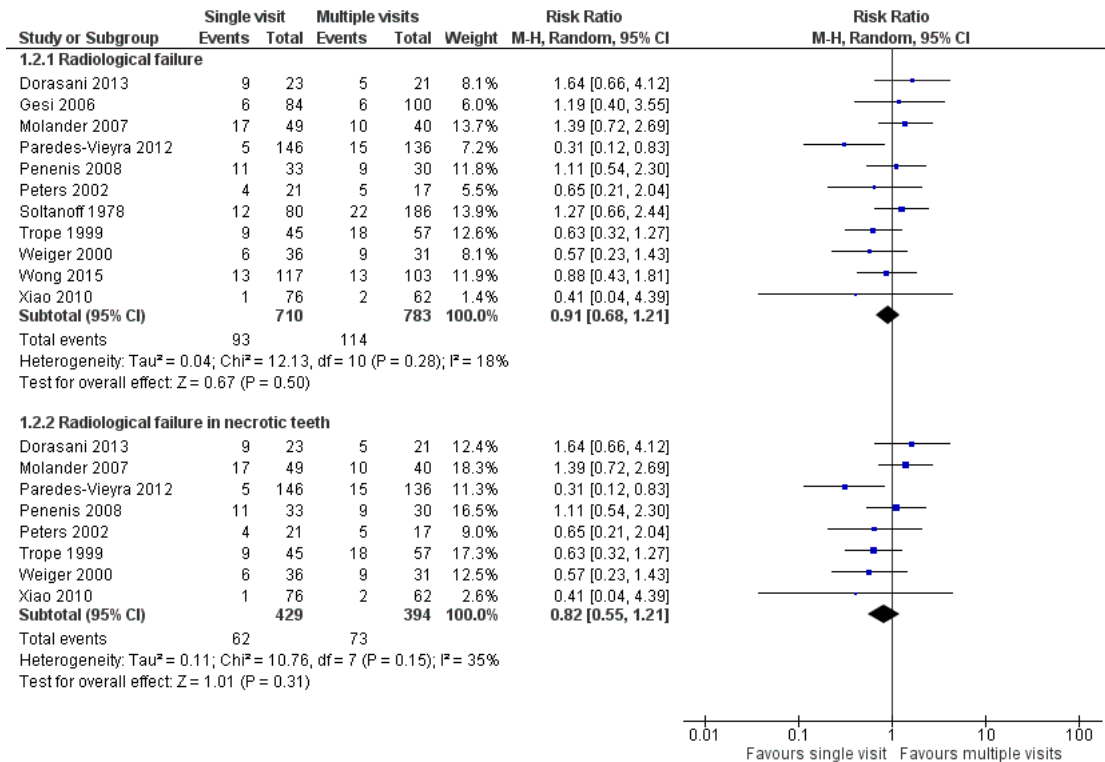
Tooth extraction due to endodontic problems (binary, yes/no)

Only one study reported data on tooth extraction due to endodontic problems ([Wong 2015](#)); there was no evidence of a difference between single- and multiple-visit treatment (odds ratio (OR) 0.44, 95% confidence interval (CI) 0.04 to 4.78; 220 participants) ([Analysis 1.1](#)). The quality of the evidence was very low ([Summary of findings for the main comparison](#)).

Radiological failure after at least one year, i.e. the presence of any periapical radiolucency (binary, yes/no)

We combined results from 11 studies ([Dorasani 2013](#); [Gesi 2006](#); [Molander 2007](#); [Penenis 2008](#); [Paredes-Vieyra 2012](#); [Peters 2002](#); [Soltanoff 1978](#); [Trope 1999](#); [Weiger 2000](#); [Wong 2015](#); [Xiao 2010](#)), that included 1467 participants with 1493 teeth ([Analysis 1.2](#); [Figure 4](#)). The follow-up was one year ([Dorasani 2013](#); [Penenis 2008](#); [Trope 1999](#)), 18 months ([Wong 2015](#)), two years ([Molander 2007](#); [Paredes-Vieyra 2012](#); [Soltanoff 1978](#); [Xiao 2010](#)), or more than three years ([Gesi 2006](#); [Peters 2002](#); [Weiger 2000](#)). The radiological failure rate was based on binary data, that is, radiological healing versus lack of such healing; scores including more than two values were dichotomised according to the methods indicated in [Table 1](#). The studies, when pooled together, irrespective of the follow-up duration, showed no difference in terms of radiological failure between the treatments and were homogeneous (RR 0.91, 95% CI 0.68 to 1.21; 1493 participants, 11 studies, $I^2 = 18\%$). The quality of the evidence related to this outcome, as assessed using the GRADE method and presented in [Summary of findings for the main comparison](#) was low ([Atkins 2004](#)).

Figure 4. Forest plot of comparison: I Primary outcomes, outcome: I.2 Radiological failure



Necrotic teeth

For eight studies (Dorasani 2013; Molander 2007; Paredes-Vieyra 2012; Penenis 2008; Peters 2002; Trope 1999; Weiger 2000; Xiao 2010), it was possible to analyse radiological failure for necrotic teeth only. Meta-analysis of these studies showed no difference between participants treated in a single visit versus those treated in multiple visits, with low statistical heterogeneity (RR 0.82, 95% CI 0.55 to 1.21; 823 participants, 8 studies, I² = 35%; Figure 4).

Vital teeth

There was one study reporting the rate of radiological failure in teeth vital at enrolment (Gesi 2006). This study found no difference between participants treated in single-visit RoCT in compar-

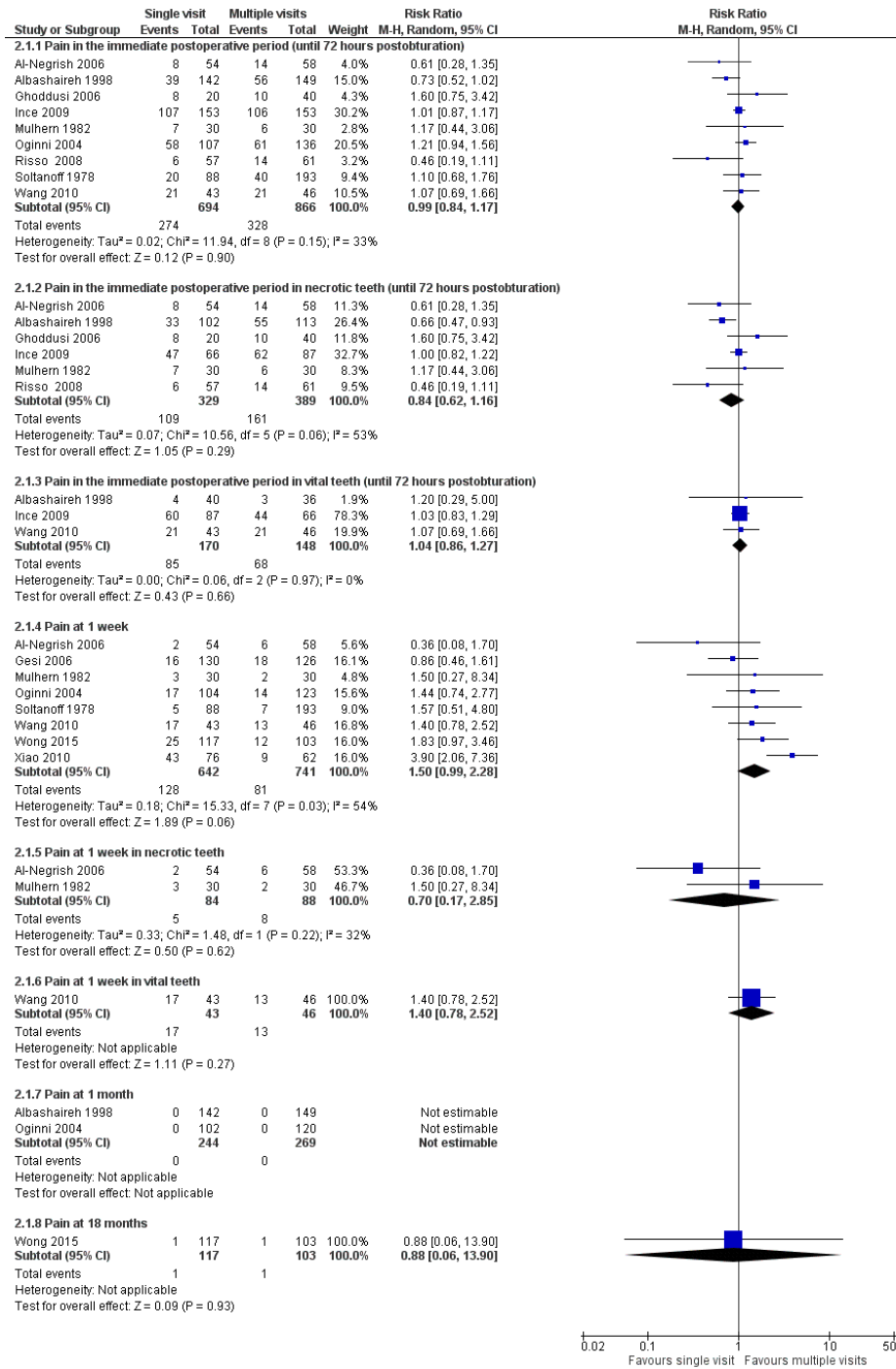
ison to those treated in multiple visits (RR 0.80, 95% CI 0.59 to 1.07; 184 participants).

Secondary outcomes

Pain in the immediate postoperative period (up to 72 hours postobturation)

Nine studies, with a total of 1560 participants, reported pain incidence 72 hours after canal obturation as a dichotomous outcome (Al-Negrish 2006; Albashaireh 1998; Ghoddsi 2006; Ince 2009; Mulhern 1982; Oginni 2004; Risso 2008; Soltanoff 1978; Wang 2010). No difference was found between groups, with low statistical heterogeneity (RR 0.99, 95% CI 0.84 to 1.17; I² = 33%; Figure 5; Analysis 2.1).

Figure 5. Forest plot of comparison: 2 Secondary outcomes, outcome: 2.1 Pain (dichotomous)



The evidence related to this outcome, as assessed using the GRADE method and presented in 'Summary of findings for the main comparison, is moderate quality.

Gesi 2006, Ghoddusi 2006, Mulhern 1982, and Xiao 2010 reported pain incidence in the interappointment period of the multiple-visit RoCT; such data are not included in the meta-analysis, as they cannot be compared with a similar outcome of the single-visit approach.

Four studies with 414 participants evaluated postoperative pain as a continuous outcome (DiRenzo 2002; Patil 2016; Singh 2012; Wang 2010). In order to perform a meta-analysis, we standardised the results of the VAS in two studies from a 170 mm scale to a 100 mm scale (DiRenzo 2002; Patil 2016). No difference was found between the two treatments, with no statistical heterogeneity (mean difference (MD) 0.12, 95% CI -0.59 to 0.82; $I^2 = 0\%$; Analysis 2.2).

Results from Yoldas 2004 were excluded from meta-analysis of postoperative pain because the data were not stratified according to time of onset. In this study, single- and multiple-visit RoCT showed no significant difference in incidence of pain. The incidence of pain was greatest during the first 48 hours after obturation, and then decreased steadily in the subsequent seven days. Of the 227 participants enrolled in the study, 68 had symptomatic and 159 had asymptomatic teeth. When data were analysed to consider the presence of symptoms before RoCT, postoperative pain was found significantly more often in participants with symptomatic teeth.

Necrotic teeth

Six studies with 718 participants reported immediate pain as a dichotomous outcome in necrotic teeth (Albashaireh 1998; Al-Negrish 2006; Ghoddusi 2006; Ince 2009; Mulhern 1982; Risso 2008). The pooled estimate from the studies showed no differences between treatments in participants with necrotic teeth (RR 0.84, 95% CI 0.62 to 1.16; $I^2 = 53\%$; Figure 5).

Three studies (DiRenzo 2002; Patil 2016; Singh 2012), including a total of 158 participants, evaluated pain in the immediate postoperative period in non-vital teeth, as a continuous outcome, and found no difference between the two treatment groups, with no statistical heterogeneity (MD -0.21, 95% CI -0.44 to 0.03; $I^2 = 0\%$).

Vital teeth

Three studies analysed immediate postoperative pain as a dichotomous outcome in teeth vital at enrolment, for a total of 318 participants (Albashaireh 1998; Ince 2009; Wang 2010). There was no difference in the pain incidence immediately after RoCT between treatment groups, with no statistical heterogeneity (RR 1.04, 95%

CI 0.86 to 1.27; $I^2 = 0\%$; Figure 5). One study (30 participants) reported no difference between treatments for postoperative pain as a continuous outcome in vital teeth (Patil 2016).

Pain at one week

Eight studies reported pain at one week as a dichotomous outcome in a total of 1383 participants and suggested that participants treated in a single visit were more likely to experience pain than those treated over multiple visits (Al-Negrish 2006; Gesi 2006; Mulhern 1982; Oginni 2004; Soltanoff 1978; Wang 2010; Wong 2015; Xiao 2010), though the difference is not statistically significant and the meta-analysis showed substantial statistical heterogeneity (RR 1.50, 95% CI 0.99 to 2.28; $I^2 = 54\%$; Figure 5). The evidence related to this outcome, as assessed using the GRADE method and presented in Summary of findings for the main comparison, is low-quality.

Pain at one week in necrotic teeth

Al-Negrish 2006 and Mulhern 1982 evaluated the incidence of pain one week after the RoCT in non-vital teeth and found no difference between participants treated in a single visit in comparison to those treated in multiple visits, with low statistical heterogeneity (RR 0.70, 95% CI 0.17 to 2.85; 172 participants, 2 studies, $I^2 = 32\%$; Figure 5).

Pain at one week in vital teeth

Wang 2010 was the only study reporting the incidence of pain at one week after RoCT in teeth vital at enrolment. This study found no difference in this outcome between participants treated in one single visit in comparison to those treated in multiple visits (RR 1.40, 95% CI 0.78 to 2.52; Figure 5).

Pain at one month

Only two studies reported pain at one month after canal obturation as a dichotomous outcome (Albashaireh 1998; Oginni 2004). In both studies, no participant had persistent pain at one month. Thus, a meta-analysis of the studies was not possible.

Pain at 18 months

Wong 2015 was the only study reporting incidence of pain at 18 months after RoCT. This study found no difference in this outcome between participants treated in one single visit in comparison to those treated in multiple visits (RR 0.88, 95% CI 0.06 to 13.90; 220 participants).

Swelling or flare-up

We considered all studies reporting flare-up as swelling and those where flare-up was defined as swelling (with or without moderate or intense pain). For definitions of flare-up, see Table 2. We included four studies with 281 participants in a meta-analysis (DiRenzo 2002; Ghoddsi 2006; Mulhern 1982; Wang 2010), which did not find a difference between single and multiple visits (RR 1.36, 95% CI 0.66 to 2.81; $I^2 = 0\%$). The evidence related to this outcome, as assessed using the GRADE method and presented in Summary of findings for the main comparison, is low quality.

Swelling or flare-up in necrotic teeth

Two studies reported data on this outcome in necrotic teeth for a total of 120 participants (Ghoddsi 2006; Mulhern 1982), showing no difference between participants treated in a single visit versus those treated in multiple visits, with no statistical heterogeneity (RR 1.52, 95% CI 0.70 to 3.31; $I^2 = 0\%$).

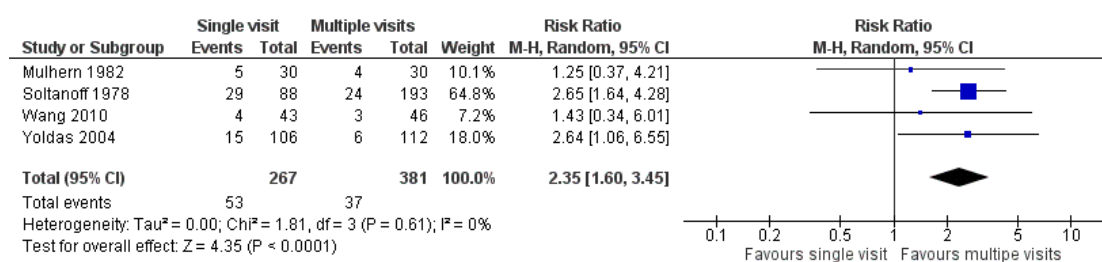
Swelling or flare-up in vital teeth

Wang 2010 was the only study reporting the incidence of swelling or flare-up in teeth vital at enrolment. This study found no difference between participants treated in one single visit in comparison to those treated in multiple visits (RR 1.07, 95% CI 0.07 to 16.57; 89 participants).

Painkiller use

Use of painkillers was reported by four studies (Mulhern 1982; Soltanoff 1978; Wang 2010; Yoldas 2004), including a total of 648 participants. Pooled estimates from all studies showed that the use of painkillers after RoCT was more common among participants undergoing the single-visit approach (RR 2.35, 95% CI 1.60 to 3.45; Figure 6). The studies were homogeneous ($\text{Chi}^2 = 1.81$, $\text{df} = 3$ ($P = 0.61$); $I^2 = 0\%$). The evidence related to this outcome, as assessed using the GRADE method and presented in Summary of findings for the main comparison, is moderate quality.

Figure 6. Forest plot of comparison: 2 Secondary outcomes, outcome: 2.4 Painkiller use



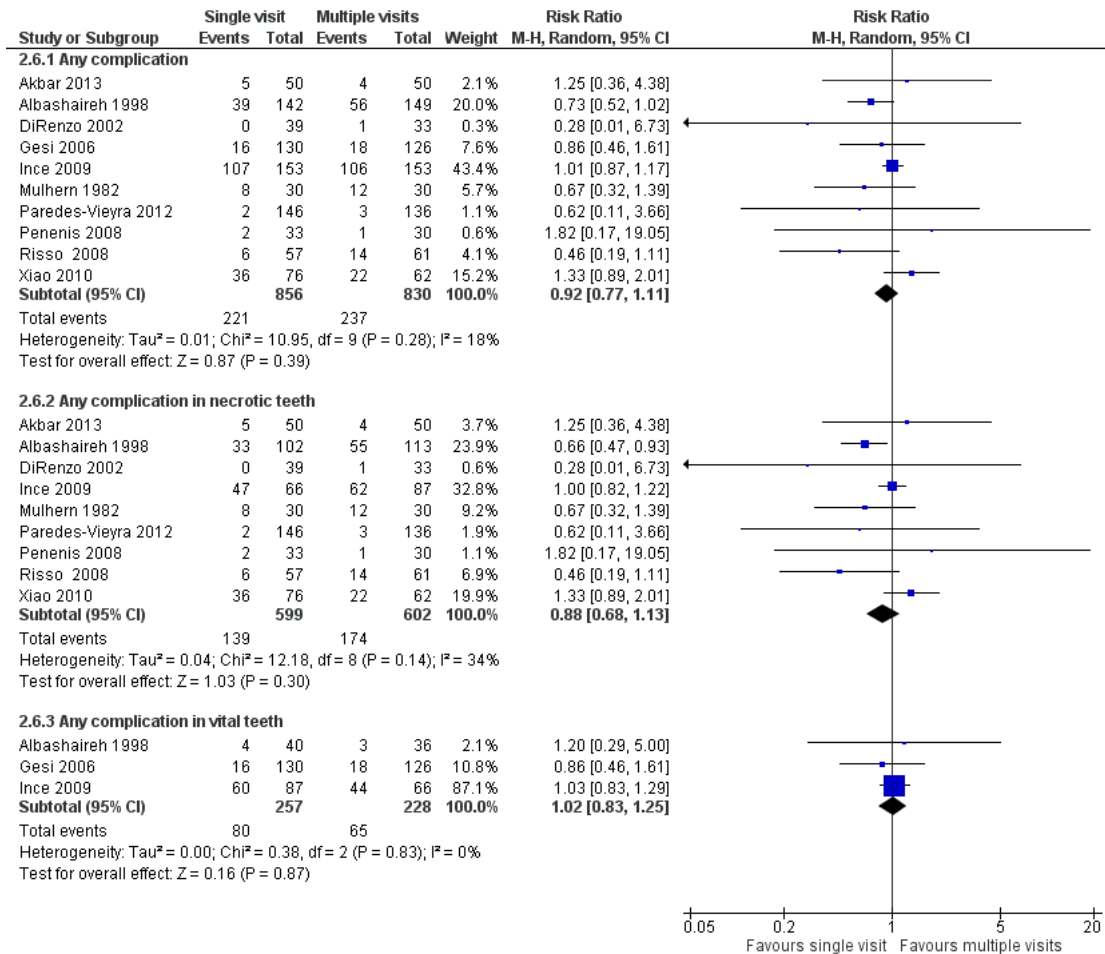
Necrotic teeth

Mulhern 1982 reported the use of painkillers in necrotic teeth, showing no difference between participants treated in a single visit versus those treated in multiple visits (RR 1.25, 95% CI 0.37 to 4.21; 60 participants).

Sinus track or fistula formation

Only two studies reported persistent sinus track or fistula, both of which included necrotic teeth only (Paredes-Vieyra 2012; Penis 2008). The studies were homogeneous and did not find a difference between single and multiple visits (RR 0.98, 95% CI 0.15 to 6.48; 345 participants; $I^2 = 0\%$, Figure 7). The evidence related to this outcome, as assessed using the GRADE method and presented in Summary of findings for the main comparison is low quality.

Figure 7. Forest plot of comparison: 2 Secondary outcomes, outcome: 2.6 Any complication



Any complication

We could extrapolate accurate data for 10 studies: Akbar 2013; Albashaireh 1998; DiRenzo 2002; Gesi 2006; Ince 2009; Mulhern 1982; Paredes-Vieyra 2012; Penenis 2008; Risso 2008; Xiao 2010. A total of 1686 participants were included in this meta-analysis, which did not find a difference between single and multiple visits; the studies were homogeneous (RR 0.92, 95% CI 0.77 to 1.11; I² = 18%; Figure 7). The evidence related to this outcome, as assessed using the GRADE method and presented in Summary of findings for the main comparison is moderate quality.

Any complication in necrotic teeth

We could extrapolate accurate data on necrotic teeth from nine studies (Akbar 2013; Albashaireh 1998; DiRenzo 2002; Ince

2009; Mulhern 1982; Paredes-Vieyra 2012; Penenis 2008; Risso 2008; Xiao 2010), with a total of 1201 participants. Meta-analysis showed no difference between participants treated in a single visit versus those treated in multiple visits, with low statistical heterogeneity (RR 0.88, 95% CI 0.68 to 1.13; I² = 34%; Figure 7).

Any complication in vital teeth

There were three studies (Albashaireh 1998; Gesi 2006; Ince 2009), including 485 participants, available to analyse the frequencies of any complication. No difference in frequency of complications was found between participants treated in a single visit versus those treated in multiple visits, with no statistical heterogeneity (RR 1.02, 95% CI 0.83 to 1.25; I² = 0%; Figure 7).

DISCUSSION

Summary of main results

This review included 25 randomised controlled trials (RCTs) with a total of 3780 participants undergoing root canal treatment (RoCT) in a single visit or over multiple visits. Only one study reported data on dental extraction due to endodontic problems (Wong 2015).

The results of this review show that, at present, there is no evidence of a substantial advantage of one of the two approaches of RoCT we compared.

Endodontic success indicators can be short- or long-term. The short-term indicators concern the absence of any postoperative discomfort, the most important short-term outcome of RoCT. Pain perception is highly subjective and modulated by multiple physical and psychological factors, and the measurement of pain is fraught with hazards and opportunities for errors. The level of discomfort must be rated in categories arranged in advance and exactly described (for example, slight pain: the tooth involved was slightly painful for a time, regardless of duration, but no need existed to take analgesics). From the results obtained in this review, there is evidence that the incidence of postobturation pain (evaluated at different times and using both dichotomous and continuous data) is similar in single- and multiple-visit RoCT, although pain at one week and painkiller use might be less in people undergoing multiple-visit RoCT. It is possible that in the single-visit approach the working time is longer, causing a more severe acute inflammatory response. Another factor could be the beneficial effect of the intracanal medication in the between-visit interval.

There is no evidence of a difference in the incidence of short-term swelling between people undergoing the single and multiple-visit approach. Among the four studies considering this outcome (DiRenzo 2002; Ghoddsu 2006; Mulhern 1982; Wang 2010), one evaluated teeth with vital pulp only (Wang 2010). Only two participants (one from the single-visit group and one from the multiple-visit group) experienced slight swelling and flare-up that required a visit to the clinic the day after obturation. The Ghoddsu 2006 study is of particular interest. In fact, when no interappointment canal medication was employed, the incidence of swelling was very similar in the two groups, while when calcium hydroxide was left in the canals between visits, the multiple-visit treatment performed much better. Such a difference may have occurred because normal saline solution was used as the sole irrigant during RoCT. Thus, with the single-visit approach, nothing with any antibacterial activity was included in the RoCT. While with the multiple-visit treatment employing interappointment medication, antibacterial activity was provided by calcium hydroxide.

Only two studies reported data on sinus track or fistula formation and from the data available there is no evidence of a difference between RoCT performed in a single visit versus multiple visits (Paredes-Vieyra 2012; Penenis 2008).

Long-term success is based mainly on the healing of periapical

lesions, whenever present, and the prevention of new lesions. The healing rate can be established by radiographic interpretation, a method very dependent on human visual perception. There is no evidence of a difference, however, in terms of radiological healing between people treated in a single visit when compared with those undergoing a multiple-visit approach.

We pooled data from 10 studies to assess the incidence of any complication reported by participants in the short-term (Akbar 2013; Albashaireh 1998; DiRenzo 2002; Gesi 2006; Ince 2009; Mulhern 1982; Paredes-Vieyra 2012; Penenis 2008; Risso 2008; Xiao 2010), finding no evidence of a difference between the two groups.

We performed subgroup analysis for vital and necrotic teeth when data were available. In the previous version of this systematic review (Figini 2007), a meta-analysis of a small number of studies suggested a possible difference in radiological healing among necrotic teeth, in favour of single-visit RoCT; this result was not confirmed in this update, where a larger number of studies were included in the meta-analysis.

Thus, the effectiveness of single- and multiple-visit RoCT does not seem to be substantially different. Most short- and long-term complications are similar in terms of frequency, although patients undergoing single-visit RoCT may experience a higher level of pain at one week after RoCT and are more likely to take analgesics.

Overall completeness and applicability of evidence

The available evidence is from a range of countries and is applicable to healthy patients aged over 10 years. Identified trials did not include patients with depressed immune systems, patients with other illnesses, or elderly patients. The results of this review may or may not be generalisable to these groups, which would be expected to have different rates of short- and long-term endodontic success. The majority of the trials were conducted in public structures (hospitals or university clinics) and only one trial was performed in a private practice. This may influence the generalisability of our results, with particular regard to the number of operators involved in the RoCT or their clinical skill, as those in the included studies varied from expert endodontists working in referral centres to postgraduate students. Whether results would be similar for general dental practitioners is unclear.

About half of the studies (11/25) did not report the use of rubber dam during the RoCT procedure. Although the use of rubber dam is considered part of the RoCT, and for this reason several authors may have not reported its use during the procedure, this might represent a limit in terms of external validity.

Quality of the evidence

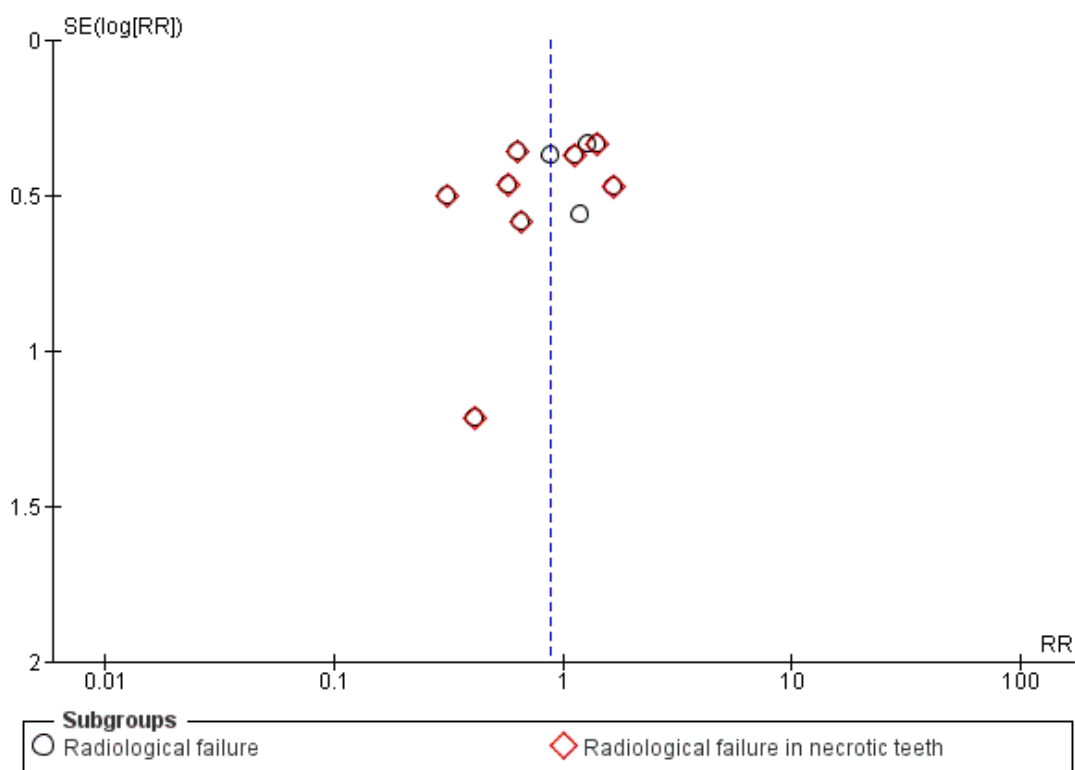
On the basis of the criteria used in the critical appraisal of the studies, only three of the 25 included studies resulted in a low risk of bias (Molander 2007; Singh 2012; Wang 2010). Less than 25% of the studies were at low risk of selection bias (i.e. with inadequate random sequence generation and/or allocation concealment). We judged the risk of attrition bias, detection bias, reporting bias, and other bias, as low in more than 75% of the studies. When assessed using the GRADE method (Atkins 2004), evidence on immediate pain (dichotomous), painkiller use and any complication, resulted in moderate quality; radiological failure, swelling or flare-up, pain at one week and persistent sinus tract or fistula resulted in low

quality; extraction due to endodontic problems resulted in very low quality (Summary of findings for the main comparison).

Potential biases in the review process

We have taken steps to minimise the bias in every step of the review. We searched databases, and trial registries with no language limitations, to identify all the relevant reports. We tried to contact the study authors for missing data through e-mails. We did not detect publication bias on the basis of the funnel plots analysis (Figure 8).

Figure 8. Funnel plot of comparison: 2 Signs, outcome: 2.3 Radiological failure



Agreements and disagreements with other studies or reviews

We found four other systematic reviews that compared pain and long-term radiological healing between single- and multiple-visit RoCT. The Sathorn 2005 review focused on postoperative pain and flare-up, coming to the same conclusions as our present re-

view. Su 2011 and Zhang 2015 analysed the differences between the two approaches, in teeth with infected root canals only. The results of the two reviews did not differ significantly from those of our subgroup analysis in necrotic teeth. In the systematic review by Su 2011, when short-term postobturation pain was analysed, a benefit for patients undergoing single-visit RoCT was evident.

This difference from our result reporting no statistical difference can be explained by the difference in inclusion criteria (necrotic teeth only), which led to a different set of studies being combined. The review by Wong 2014 concluded that the studies reported in the literature showed that neither single-visit endodontic treatment nor multiple-visit treatment could guarantee the absence of postoperative pain. It found that neither single-visit endodontic treatment nor multiple-visit treatment had superior results in terms of healing or success rate, which our review also found.

AUTHORS' CONCLUSIONS

Implications for practice

There is no evidence to suggest that one treatment regimen (single-visit or multiple-visit root canal treatment (RoCT)) is better than the other. Neither can prevent all short- and long-term complications. On the basis of the available moderate-quality evidence, it seems likely that the benefit of a single-visit treatment, in terms of time and convenience, for both patient and dentist, has the cost of a higher frequency of late postoperative pain (and as a consequence, painkiller use).

Implications for research

As use of rotary nickel titanium instruments is increasing, a well-designed RCT comparing single-visit and multiple-visit RoCT, both performed with such instruments, would be an important contribution. It would be very helpful for clinicians that researchers include tooth loss as a primary outcome in new studies, even reporting if none occurs.

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* *Indicates the major publication for the study*

CHARACTERISTICS OF STUDIES

Characteristics of included studies *[ordered by study ID]*

Akbar 2013

Methods	Study design: randomised parallel group clinical trial. Conducted in Pakistan. Number of centres: 1. Recruitment period: not reported. Funding source: not reported.
Participants	Inclusion criteria: one asymptomatic molar tooth with periapical radiolucency Exclusion criteria: not reported. Diagnostic criteria for pulpal or periapical disease: periapical Rx of the involved tooth and patient's history 100 participants randomised: 51 males, 49 females, aged between 12 and 40 years Group 1: 50 randomised (1 participant 1 tooth), 50 analysed. Group 2: 50 randomised (1 participant 1 tooth), 50 analysed.
Interventions	Group 1: single visit. Group 2: multiple visits (2 visits, not reported how many days after the first the second visit was performed, root canal medication with calcium hydroxide) Number of operators not reported. Rubber dam isolation. Use of magnification loupes not reported. Canal shaping: step-back technique by K files and gates Glidden-drills. Canal irrigation: 2 ml of 2.5% sodium hypochlorite. Working length determined by Rx. Obturation with gutta-percha cones and a zinc oxide eugenol sealer in lateral condensation technique
Outcomes	<ul style="list-style-type: none"> • Flare-up Measured after obturation and daily for 7 days and defined as moderate to severe pain, or moderate to severe swelling that begins 12 to 48 hours after treatment and lasts at least 48 hours
Notes	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "the patients were randomly assigned into two groups"
Allocation concealment (selection bias)	Unclear risk	Comment: insufficient information to permit judgment.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes

Akbar 2013 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Low risk	100/100 of participants enrolled (100%) were analysed.
Selective reporting (reporting bias)	Low risk	Comment: important outcomes and adverse effects reported as planned
Other bias	Low risk	Comment: no other source of bias identified.

Al-Negrish 2006

Methods	Study design: quasi-randomised parallel group clinical trial Conducted in Jordan. Number of centres: 1. Recruitment period: not reported. Funding source: not reported.
Participants	Inclusion criteria: one asymptomatic necrotic central incisor Exclusion criteria: any evidence of periapical radiolucent lesion, teeth tender to touch, with intracanal calcification or incompletely formed apices, retreatments, teeth with pulpal sensitivity and vitality Diagnostic criteria for pulpal or periapical disease: Rx signs, pulp testing, presence or absence of haemorrhage upon access opening 120 participants randomised: 66 female and 54 males, aged between 15 to 45 years Group 1: 60 quasi-randomised; 54 analysed. Group 2: 60 quasi randomised; 58 analysed.
Interventions	Group 1: single visit. Group 2: multiple visits (2 visits, second appointment 7 days later the first, root canal medication with calcium hydroxide paste with a dry sterile cotton pledget and temporary filling restoration for 7 days) Single operator. Rubber dam isolation. Use of magnification loupes not reported. Canal shaping: step-back technique with conventional K files and gates. Irrigation: 2.5% sodium hypochlorite. Working length determined by Rx. Obturation with gutta-percha and a zinc oxide eugenol sealer (Tubliseal, Kerr) in lateral condensation
Outcomes	<ul style="list-style-type: none"> • Pain Pain after 2-day postobturation period and after 7-day postobturation period was reported with a 4-grade scale: 1 = no pain, 2 = slight pain, 3 = moderate pain, 4 = severe pain. In the present review, the scale was dichotomised: no pain versus pain (slight, moderate or severe) <ul style="list-style-type: none"> • Flare-up Proportion of participants experiencing moderate to severe pain evaluated after 2 and 7 days (see Table 2).
Notes	
Risk of bias	

Al-Negrish 2006 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Comment: quasi-random method.
Allocation concealment (selection bias)	High risk	Comment: no allocation concealment was possible as patients were alternatively assigned to treatments
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	112/120 patients (93.3%) of patients who entered the study were included in the final analysis. Eight patients (6 females and 2 males, 6 from Group 1 and 2 from Group 2) were excluded from the analysis of the results as they failed to attend postoperative visits
Selective reporting (reporting bias)	Low risk	Comment: important outcomes and adverse effects reported as planned
Other bias	Low risk	Comment: no other source of bias identified.

Albashaireh 1998

Methods	Study design: quasi-randomised parallel group, clinical trial Conducted in Jordan. Number of centres: 1. Recruitment period: not reported. Funding source: not reported.
Participants	Inclusion criteria: one tooth for each participant. Exclusion criteria: teeth tender to touch, with extensive intracanal calcification and incompletely formed apices Diagnostic criteria for pulpal or periapical disease: pulpal vitality and sensitivity (pulp testing and direct presence or absence of haemorrhage), presence of periapical radiolucency in periapical radiographs 300 participants randomised (sex not reported, aged between 15 to 65 years) Group 1: 150 quasi-randomised; 142 analysed. Group 2: 150 quasi randomised; 149 analysed.
Interventions	Group 1: single visit. Group 2: multiple visits (2 visits, no intra-appointment medicament was placed, but a dry sterile cotton pledget sealed in pulp-chamber with a temporary filling restoration)

	One operator. Rubber dam isolation. Use of magnification loupes and working length not reported. Canal shaping with step-back technique, obturation with gutta-percha and a calcium hydroxide-based root canal sealer (Sealapex) with lateral condensation technique. Irrigation with 2.6% sodium hypochlorite solution	
Outcomes	<ul style="list-style-type: none"> • Pain Incidence and degree of pain at the 1st and 30th postobturation day was reported on a 4-grade scale: 1 = no pain, 2 = slight pain, 3 = moderate pain, 4 = severe pain. In the present review, the scale was dichotomised: no pain versus pain (slight, moderate, or severe)	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Comment: quasi-random method.
Allocation concealment (selection bias)	High risk	Comment: no allocation concealment was possible as participants were alternatively assigned to treatments
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	291/300 (97%) of participants who entered the study were included in the final analysis 9 participants (8 from Group 1 and 1 from Group 2) were excluded from the analysis of the results as 5 failed to attend postoperative visits, 3 required surgery 2 weeks after obturation and 1 had the involved tooth extracted elsewhere
Selective reporting (reporting bias)	Low risk	Comment: important outcomes and adverse effects reported as planned
Other bias	Low risk	Comment: no other source of bias identified.

DiRenzo 2002

Methods	Study design: randomised two arm parallel group clinical trial Conducted in USA. Number of Centres: 1. Recruitment period: not reported. Funding source: not reported.
Participants	Inclusion criteria: mature vital and non-vital permanent maxillary and mandibular molars requiring root canal therapy Exclusion criteria: pregnancy, use of antibiotics or corticosteroids at the time of treatment, immunocompromised states, subjects under 18 years old Diagnostic criteria for pulpal or periapical disease: not reported 80 participants randomised (sex, age and ethnic group not specified, over 18 years of age) Group 1: randomised 46; analysed 39. Group 2: randomised 34; analysed 33.
Interventions	Group 1: Single visit. Group 2: Multiple visits (2 visits, in the interappointment period the teeth were closed with a sterile dry cotton pellet and Cavit temporary restorative cement) Two operators (postgraduate students). Rubber dam isolation. Use of magnification loupes not reported. Canal shaping with hand files and nickel titanium rotary files. Irrigation with 2.5% sodium hypochlorite. Working length determined by an electronic apex locator and 2 or more angled radiographs. Obturation with gutta-percha and Roth 811 sealer in lateral condensation
Outcomes	<ul style="list-style-type: none"> • Pain A modified VAS was used to measure pain at 6, 12, 24, 48 hours after the first appointment <ul style="list-style-type: none"> • Flare-up Defined as swelling that needs antibiotics and narcotic analgesics (see Additional Table 2).
Notes	Data on pain were not available and it was not possible to extract them from the tables, for such reason data on pain were not available for meta-analysis

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "random assignment by coin toss".
Allocation concealment (selection bias)	Unclear risk	Comment: it is unclear by whom and when the coin toss was performed and how the result was communicate to the operators
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes

DiRenzo 2002 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Low risk	90% of patients (72/80) who entered the study were included in the final analysis 5 patients (Group 1) dropped out because of their inability to complete the treatment in 1 appointment, 3 patients (2 group 1, 1 group 2) did not return the VAS form
Selective reporting (reporting bias)	Low risk	VAS pain measurements reported.
Other bias	Low risk	Comment: no other source of bias identified.

Dorasani 2013

Methods	Study design: randomised two arm parallel group clinical trial Conducted in: India. Number of centres: 1. Recruitment period: not reported. Funding source: not reported.
Participants	Inclusion criteria: <ul style="list-style-type: none"> patients aged between 18 and 62 years, only single rooted teeth with Vertucci's type I configuration, teeth with radiographic evidence of periapical pathology (PAI \geq 3) and pulpal necrosis. Exclusion criteria: <ul style="list-style-type: none"> any systemic disease, pregnancy, use of antibiotics, corticosteroids or anti-inflammatory drugs prior to time of treatment, necessity of antibiotic premedication for dental treatment, previously accessed tooth, grossly decayed tooth (difficulty in rubber dam isolation), teeth with calcified canals and weeping canals. Diagnostic criteria for pulpal or periapical disease: <ul style="list-style-type: none"> not reported. 64 teeth from 57 patients (30 males, 27 females, mean age 40 years, range 18-62 years) were randomised. 6 patients contributed with more than 1 tooth: 5 patients with 2 teeth and 1 patient with 3 teeth Group 1: 34 teeth randomised, 23 analysed. Group 2: 30 teeth randomised, 21 analysed.
Interventions	Group 1: Single visit. Group 2: Multiple visits (2 visits, the second one 7 days after the first; in the interappointment period a paste carrier was used to carry calcium hydroxide medicament in the root canal and temporarily restored with Cavit) Single operator. Rubber dam isolation. Use of magnification loupes not reported. Canal shaping: hand instrumentation with flexo-files using balanced force technique and crow-down technique. Irrigation: 3% sodium hypochlorite and saline. Working length checked with apex locator and confirmed by Rx. RC-help was used as a lubricant during filling. Obturation with gutta-percha cones and pulp canal sealer (Kerr) with lateral condensation

Dorasani 2013 (Continued)

Outcomes	<p>Clinical assessment:</p> <ul style="list-style-type: none"> presence of clinical signs and symptoms at 12 months (spontaneous pain, presence of sinus tract, swelling, mobility, periodontal probing depths greater than baseline measurements, or sensitivity to percussion or palpation). <p>Radiographic assessment:</p> <ul style="list-style-type: none"> change in apical bone density at 12 months using PAI. Evaluation at 3 and 6 months was also performed. 	
Notes	The patients who had taken medication for any systemic illness during the follow-up period were excluded from the study	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Comment: insufficient information to permit judgment.
Allocation concealment (selection bias)	Unclear risk	Comment: insufficient information to permit judgment.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Comment: insufficient information to permit judgment.
Incomplete outcome data (attrition bias) All outcomes	High risk	44/64 (70%) of teeth included in the study were reported in the study. Two treatment failures before the 12-month examination (1 in G1 and 1 in G2) 18 teeth lost at follow-up
Selective reporting (reporting bias)	Low risk	Comment: important outcomes and adverse effects reported as planned
Other bias	High risk	The number of teeth exceeds the number of patients and it is not specified whether patients were randomised a number of time equal to the number of teeth undergoing treatment

Methods	Study design: Randomised parallel group, clinical trial. Conducted in Italy. Number of Centres: 2, Private dental practices in Pisa and Pistoia, Italy Recruitment period: 24 months. Funding source: not reported.	
Participants	256 participants (141 females, 115 males, age and ethnic group not reported) Inclusion criteria: patients with teeth with painful and non-painful vital pulp, with bleeding upon access of the pulpal chamber Exclusion criteria: patients with physical or mental disability, patients that took pain medications or in treatment with antibiotics for systemic or local infection Diagnostic criteria for pulpal or periapical disease: vitality testing and thermal and mechanical stimulation Group 1: randomised 130; included 130. Group 2: randomised 126; included 126.	
Interventions	Group 1: Single visit. Group 2: Multiple visits (2 visits, in the interappointment period calcium hydroxide was employed as intracanal medication and Coltosol as temporary cement) Single operator. Rubber dam. Use of magnification loupes not specified. Canal shaping: hand instrumentation with flexo-files using balanced force technique and crow-down technique. Irrigation: 3% sodium hypochlorite. Working length established by Rx. Obturation with gutta-percha and pulp canal sealer (Kerr) with lateral condensation	
Outcomes	<ul style="list-style-type: none"> • Pain <p>Evaluated at 1 week after canal obturation by clinical examination and by a verbal rating scale to assess pain experience. Participants with multivisit treatments were asked to evaluate their pain after 1 week for each visit. We considered pain-related data only after canal obturation. A verbal rating scale graded 0 to 3 was used. Participants were asked to indicate 0 for no, 1 for mild, 2 for moderate and 3 for severe pain. Teeth were also tapped for percussion sensitivity. We considered only 2 categories: pain (mild, moderate, severe) and no pain</p> <ul style="list-style-type: none"> • Healing <p>(Follow-up until 3 years): 2 endodontists, experienced in radiographic assessment of endodontic treatments, neither of whom was the operator and both masked to the assigned treatment group, carried out the analysis of the radiographs. Parameters were presence or absence of periapical radiolucency (radiographic lesion) (see Table 1). In 2- or multi-rooted teeth, the tooth was classified according to the diagnosis of the worst root</p>	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "by a simple randomisation procedure (toss of a coin) patients were allocated in group 1 or 2"

Gesi 2006 (Continued)

Allocation concealment (selection bias)	Unclear risk	Unclear by whom and when the coin toss was performed and how the result was communicate to the operators
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Healing: Quote:“ two endodontists, well experienced in radiographic assessment of endodontic treatments, neither of whom was the operator and both masked to the assigned treatment group, carried out the analysis of the radiographs”
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of patients who entered the study were included in the final analysis of the outcome 'pain.' 71.8% (244/256) of patients who entered the study were included in the final analysis of the outcome 'healing at 3 years follow-up'
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Ghoddusi 2006

Methods	Study design: three-arm randomised parallel group, clinical trial Conducted in Iran. Number of Centres: 1. Endodontics Department of Mashad Dental School, Iran Recruitment period: not reported. Funding source: a grant from the vice chancellor of research of Mashad University of Medical Sciences, Iran
Participants	69 patients enrolled (39 females and 30 males, not specified age range and ethnic group) 60 were randomised Inclusion criteria: patients with pulpally necrotised teeth referred to the Endodontics Department of Mashad Dental School Exclusion criteria: patients taking some medication for systemic conditions. Diagnostic criteria for pulpal or periapical disease: thermal and electrical pulp test, pulp cavity test (direct presence or absence of haemorrhage), presence of periapical radiolucency in periapical radiographs Group 1: randomised 20, analysed 20. Group 2: randomised 20, analysed 20. Group 2: randomised 20, analysed 20.
Interventions	Group 1: Single visit. Group 2: Multiple visits without any interappointment dressing (after the first appointment the canal was left empty, the treatment was completed after 1 week)

Ghoddusi 2006 (Continued)

	<p>Group 3: Multiple visits with interappointment dressing (the root canal space was filled with a diluted mixture of calcium hydroxide powder and aquapura water, the treatment was completed after 1 week)</p> <p>Rubber dam isolation used. Use of magnification loupes not specified. Canal shaping with hand files (K files). Irrigation with saline solution. Working length evaluated by Rx, obturation with gutta-percha in lateral condensation</p>	
Outcomes	<ul style="list-style-type: none"> • Pain <p>Incidence and degree of pain in the immediate canal postobturation until 72 hours was reported as: 1 = no pain, 2 = mild pain, 3 = moderate pain, 4 = severe pain. We considered only 2 categories: no pain, and pain (mild, moderate and severe pain)</p> <ul style="list-style-type: none"> • Flare-up (swelling) 	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote "The patients were randomly assigned to the three groups"
Allocation concealment (selection bias)	Unclear risk	Not specified.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of patients who entered the study were included in the final analysis
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Ince 2009

Methods	<p>Study design: randomised parallel group, clinical trial.</p> <p>Conducted in Turkey.</p> <p>Number of Centres: more than one. Quote: "...patients who attended participating dental clinics..."</p> <p>Recruitment period: not reported.</p> <p>Funding source: not reported.</p>
Participants	<p>306 participants (200 males, 106 females, 18 to 60 (?) years old - average 45 years), 153 vital teeth and 153 non-vital teeth</p> <p>Inclusion criteria: patients with one tooth only that required RoCT, between 18 to 60</p>

	<p>years of age, in good health</p> <p>Exclusion criteria: patients who had previously taken analgesics or antibiotics</p> <p>Diagnostic criteria for pulp vitality: electric test with pulp testing device</p> <p>Group 1: randomised 153; included 153.</p> <p>Group 2: randomised 153; included 153.</p>
Interventions	<p>G1: single visit.</p> <p>G2 : multiple visits (2 visits, second appointment 7 days after the first, no interappointment medication, a sterile cotton pellet was placed in the pulp-chamber and the access cavity was sealed with quick-setting zinc oxide eugenol cement)</p> <p>Two experienced clinicians performed the treatments.</p> <p>Rubber dam isolation - use of magnification loupes not specified</p> <p>Canal shaping: step-back technique, hand files and Gates-Glidden drills. Irrigation: 2 ml of 5% sodium hypochlorite; working length determined radiographically; root canals were filled with gutta-percha points sealed with AH-26 root canal sealer using lateral condensation technique</p>
Outcomes	Pain: preoperative (absence or presence of pain), postoperative at 3 days after initial appointment (absence or presence of pain; degree of pain: none, slight, moderate, severe)
Notes	No data are reported in text regarding: 1. how many centres have been involved in the study; 2. the exact age of the participants included in the study (Table 1 reports a different range of age from that declared in the material and method section); 3. data about secondary outcomes (radiographic and clinical data at follow-up) that are not reported in the text. An e-mail was sent to the corresponding author but no answer was obtained

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported.
Allocation concealment (selection bias)	Unclear risk	Quote: "The case and the operator distribution were blinded..."
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of patients were analysed.
Selective reporting (reporting bias)	Low risk	Outcome reported.
Other bias	Low risk	Comment: no other source of bias identified.

Molander 2007

Methods	<p>Study design: randomised parallel-group, clinical trial. Conducted in Sweden. Number of Centres: 1. Clinic of Endodontics, Public Dental Health Service, Gothenburg, Sweden Recruitment period: not reported. Founding Source: not reported.</p>
Participants	<p>101 teeth, 94 participants (47 women, 47 men, mean age 55 years) Inclusion criteria: asymptomatic teeth with necrotic pulps and apical periodontitis Exclusion criteria: not specified. Diagnostic criteria for periapical disease: radiography. G1: 53 teeth randomised, analysed 49. G2: 48 teeth randomised, analysed 40.</p>
Interventions	<p>G1: single visit. G2 : multiple visits (2 visits, second appointment 7 days after the first, calcium hydroxide was used as interappointment medication using a lentulo spiral and access cavity sealed with Coltosol) Four endodontic specialists performed the treatments. Rubber dam isolation, use of magnification loupes and working length not specified Canal shaping: nickel titanium instruments for rotary and/or hand use, Irrigation: 0.5% sodium hypochlorite; root canals were filled with gutta-percha cones using cold lateral condensation technique including rosin chloroform as sealer</p>
Outcomes	<p>Healing (radiographic and clinical) at 2 years. Two examiners independently evaluated all the RX (previously coded blind and organised in a random order). Observer used a strict definition of periapical disease and reported a positive finding only when absolute certain. In case of disagreement joint re-evaluation was performed. The size of periapical radiolucency was assessed by measuring with a ruler its largest horizontal and vertical width Teeth with symptoms of persisting periapical inflammation: not healed Cases with unchanged/increased size of periapical radiolucency: not healed Teeth with reduced size of periapical rarefaction (sum of horizontal and vertical reduction > or = 2 mm): uncertain Teeth with complete restitution of periodontal contours: healed Teeth with more than 1 root, the least favourable outcome was register</p>
Notes	<p>CONSORT Clinical trial. 3 out of 12 participants lost to follow-up died.</p>

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Randomisation was performed before the clinical examination using the minimisation method. Two randomisation factors were considered: tooth group and size of periapical lesion"

Molander 2007 (Continued)

Allocation concealment (selection bias)	Low risk	Quote: “Patients were randomly assigned to one or two-visit treatment”
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote: “Two examiners independently evaluated all the Rx exams, previously coded blind and organised in a random order”
Incomplete outcome data (attrition bias) All outcomes	Low risk	89/101 teeth analysed (12 teeth lost, 12%) . Quote “The loss did not alter the results”
Selective reporting (reporting bias)	Low risk	Outcome reported.
Other bias	Low risk	Comment: no other source of bias identified.

Mulhern 1982

Methods	Study design: randomised parallel group, clinical trial. Conducted in USA. Number of centres: not reported. Recruitment period: not reported. Funding source: a Grant-in-Aid of research from the Endowment and Memorial Foundation of the American Association of Endodontists
Participants	60 participants (31 females, 29 males, range age from 13 to 75 years, ethnic group reported: 1 Asian, 42 White and 17 Black) Inclusion criteria: non-surgical endodontic treatment of asymptomatic mature single-rooted teeth with necrotic pulps Exclusion criteria: patients with severe medical conditions, using corticosteroids or anti-inflammatory drugs and/or recent or active antibiotics therapy. Diagnostic criteria for pulpal or periapical disease: Rx and vitality test Group 1: randomised 30, analysed 30. Group 2: randomised 30, analysed 30
Interventions	Group 1: single visit. Group 2: multiple visits (3 visits: in the interappointment period no medication was used, only a dry pledget of cotton with a double cement system of Cavit G and zinc oxyphosphate cement in the coronal access cavity was employed) 2 operators (graduate endodontic students). Rubber dam. Use of magnification loupes and canal shaping not detailed. Irrigation: 2.5% sodium hypochlorite. Working length not reported. Obturation with lateral condensation was performed using gutta-percha and Kerr Tubli-Seal
Outcomes	<ul style="list-style-type: none"> • Pain Evaluated at 48 hours after treatment (by a questionnaire) and at 1 week (clinical examination). Participants with multivisit treatment were asked to complete a questionnaire for each visit

Mulhern 1982 (Continued)

	<ul style="list-style-type: none"> • Painkiller use • Flare-up (swelling) (see Table 2). 	
Notes	Patients in the experimental group received free treatment, whereas those in the control group were charged the usual clinic fee for treatment	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Quote "The teeth were randomly assigned to group 1 (single visit) or 2 (multiple visits)"
Allocation concealment (selection bias)	Unclear risk	Not specified.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of patients who entered the study were included in the final analysis
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Oginni 2004

Methods	<p>Study design: randomised parallel group, clinical trial. Conducted in Nigeria. Number of Centres: 1. Restorative Dentistry Department, Obafemi Awolowo University Teaching Hospitals Complex, Ile-Ife, Nigeria Recruitment period: 12 months. Funding source: not reported.</p>
Participants	<p>255 patients (sex, range or mean age not reported) were enrolled in the study for a total of 283 teeth (1.11 teeth per patient); 227 teeth were included in the final analysis about pain and flare-up. For patients with more than 1 tooth requiring treatment, the treatment of each tooth was separated by a period of at least 4 weeks Inclusion criteria: all patients referred to the Department of Restorative Dentistry for root canal therapy Exclusion criteria: patients that did not turn up after the first appointment (incomplete treatment) Diagnostic criteria for pulpal or periapical disease: the pulp vitality was determined by an electric pulp tester in combination with the presence of pulpal haemorrhage Group 1: randomised 129 107 teeth evaluated (1st postobturation day); 104 (7th pos-</p>

	tobturation day); 102 (30th postobturation day) Group 2: randomised 154, 136 teeth evaluated (1st postobturation day); 123 (7th postobturation day); 120 (30th postobturation day)	
Interventions	Group 1: single visit. Group 2: multiple visits (medication and number of visits in the multiple-visit treatment not reported) Rubber dam isolation, use of magnification loupes, canal shaping, irrigation, working length not reported. The root canals were obturated with multiple gutta-percha cones and a zinc oxide-eugenol based sealer, using the lateral condensation technique	
Outcomes	<ul style="list-style-type: none"> • Pain Incidence and degree of pain at the 1st, 7th and 30th days postobturation. Pain was recorded as none, slight or moderate/severe. We considered only 2 categories: pain (slight and moderate\severe), and no pain • Flare-up Defined as either patient reporting pain not controlled with over-the-counter medication or increasing swelling or both (see Table 2). 	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Comment: insufficient information to permit judgement.
Allocation concealment (selection bias)	Unclear risk	Comment: insufficient information to permit judgement.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	56 teeth (25 from single-visit group, 31 from multiple-visit group) were excluded from the study due to non availability of participants at postobturation recall visits 85.86% of patients who entered the study were included in the final analysis about pain and flare-up in the 1st day; 80.21% of patients who entered the study were included in the final analysis of pain and flare-up on the 7th day; 78.4% of patients who entered the study were included in the final analysis at 30th day

Oginni 2004 (Continued)

Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	High risk	It is not explicitly stated that participants with more than one tooth needing treatment were randomised the same number of times

Paredes-Vieyra 2012

Methods	<p>Study design: randomised parallel group, clinical trial. Conducted in Mexico. Number of centres: 1. School of Dentistry, Universidad Autonoma de Baja California, Tijuana, Baja California, Mexico Study period: February 2009 to December 2011. Funding source: not reported.</p>
Participants	<p>287 participants (149 females and 138 males, age range: 18 to 60 years, mean age 55) and 300 teeth were enrolled; 282 teeth were included in the final analysis. For participants with more than 1 tooth requiring treatment, the treatment of each tooth was separated by a period of at least 4 weeks</p> <p>Inclusion criteria: radiographic evidence of apical periodontitis (minimum size 2.0 mm) and a diagnosis of pulpal necrosis confirmed by negative response to hot and cold tests, acceptance of the aims and requirements of the study, patients in good health, all teeth had nonvital pulps and apical periodontitis with or without a sinus tract, a negative response to hot and cold pulp sensitivity tests, presence of enough coronal tooth structure for rubber dam isolation, no prior endodontic treatment on the involved tooth, no analgesics or antibiotics used before the clinical procedures</p> <p>Exclusion criteria: patients who did not meet inclusion requirements, patients who did not provide authorisation for participation, patients younger than 16 years of age, patients who were pregnant, history of antibiotic use within the past month, patients who were diabetic, patients whose tooth had been previously accessed or endodontically treated</p> <p>Diagnostic criteria for pulpal or periapical disease: radiographic evidence of apical periodontitis (minimum size ≥ 2.0 mm x 2.0 mm) and a diagnosis of pulpal necrosis confirmed by negative response to hot and cold tests</p> <p>Group 1: randomised 155 teeth, analysed after a 2-year follow-up 146 teeth Group 2: randomised 145 teeth, analysed after a 2-year follow-up 136 teeth</p>
Interventions	<p>All treatment sessions were approximately 50 minutes in length to allow for acceptable time for the completion of treatment for 1 or 2 visits</p> <p>All treatment was performed by the author.</p> <p>Rubber dam isolation. The tooth was disinfected with 5.25% sodium hypochlorite. All caries were removed and endodontic access cavities made with sterile high-speed carbide #331 and Zekrya Endo burs. The working length was established with the Root ZX Electronic Apex Locator and confirmed radiographically</p> <p>The canals were negotiated and enlarged with hand instruments until reaching an ISO #20 at the working length. The coronal portions of the canals were flared with sizes 2 to 3 Gates Glidden burs</p> <p>Canals were then irrigated with 2.0 ml 5.25% sodium hypochlorite. LightSpeedLSX</p>

	<p>rotary instruments were used to complete the canal preparation to a size #60 for the anteriors and premolars and to a size #45 to #55 for molars. RC prep was used as a lubricant. After completion of canal instrumentation, all canals were irrigated with 2.5 ml 17% EDTA for 30 seconds followed by a final irrigation with 5.0 ml 5.25% sodium hypochlorite using the EndoVac irrigation system</p> <p>Group 1: Single visit: the canals were dried with sterile paper points and obturated at the same appointment by using lateral condensation of gutta-percha and Sealapex sealer. Access cavities of anterior teeth were etched and restored with Fuji IX. For posterior teeth, a build-up restoration was placed by using the same etching technique and Fuji IX</p> <p>Group 2: multiple visits (2 visits: the second appointment was scheduled at least 1 week after the initial appointment). The canals were dried and calcium hydroxide powder was placed with an amalgam carrier and condensed with a size 9 posterior Schilder plugger. The access cavities were sealed with Cavit, and the quality of the calcium hydroxide powder filling was checked radiographically with post-treatment radiographs. At the second appointment, the calcium hydroxide was removed with hand instruments, and copious irrigation with 5.25% sodium hypochlorite followed by 2.5 ml 17% EDTA and a final rinse of 5.0 ml 5.25% sodium hypochlorite using the EndoVac irrigation system was performed. For complete removal of the calcium hydroxide, the canals were dried with sterile paper points, and obturation was performed with the same technique described for the 1-visit group and post-treatment</p>
<p>Outcomes</p>	<ul style="list-style-type: none"> • Radiographic healing assessment <p>The primary outcome measure for this study was classified by using a modification of the Strindberg study used for radiographic healing assessment. Teeth with symptoms of persisting periapical inflammation were scored as not healed as were the cases with periapical radiolucencies that remained unchanged or increased in size</p> <p>Secondary outcome measures:</p> <ul style="list-style-type: none"> • Presence of clinical symptoms or abnormal findings (i.e. spontaneous pain, presence of sinus tract, swelling, mobility, periodontal probing depths greater than baseline measurements, or sensitivity to percussion or palpation). <p>Outcomes were evaluated at 2 year follow-up.</p>
<p>Notes</p>	<p>CONSORT report.</p> <p>Financial incentive to return for follow-up for clinical and radiograph examination</p> <p>Discrepancy between data on participants treated for more then one tooth and total number of teeth: authors stated that "Twenty-one patients contributed more than 1 tooth", but since the number of patients enrolled is 287 and the number of teeth enrolled is 300, it seems that no more than 13 patients could have contributed more than 1 tooth</p>
<p>Risk of bias</p>	
<p>Bias</p>	<p>Authors' judgement</p>
<p>Random sequence generation (selection bias)</p>	<p>Low risk</p> <p>The participant was randomly assigned to either the 1-visit or 2-visit group by using a sequence of random numbers generated by one of the investigators by a computer programme</p>

Paredes-Vieyra 2012 (Continued)

Allocation concealment (selection bias)	Unclear risk	Comment: insufficient information to permit judgment.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote: radiographic images were coded and stored and evaluated blindly and independently by 2 experienced endodontists
Incomplete outcome data (attrition bias) All outcomes	Low risk	Only 18 (9 in each group) out of 300 (6%) teeth lost at follow-up
Selective reporting (reporting bias)	Low risk	Outcomes reported.
Other bias	High risk	It is not explicitly stated that patients with more than one tooth needing treatment were randomised the same number of times

Patil 2016

Methods	<p>Study design: randomised parallel group, open-label clinical trial Conducted in India. Number of centres: 1. Department of Conservative Dentistry and Endodontics, KLE University Study period: not reported. Funding source: not reported.</p>
Participants	<p>66 participants (gender of patients and age range not specified) and 66 teeth were enrolled Inclusion criteria: acceptance of the aims and requirements of the study, vital and non-vital maxillary central incisors that needed endodontic treatment and teeth in which initial master file (K-type) binds at the apex was of ISO size #45 or less Exclusion criteria: patients with any systemic diseases, pregnant patients, patients younger than 15 years of age and older than 50 years, patients who had been taking antibiotics, non steroidal anti-inflammatory drugs or corticosteroids at the time of treatment, patients with acute apical periodontitis, acute apical abscess and weeping canals, necrotic painful teeth with absence of sinus tract for drainage, retreatment cases, teeth with calcified canals, teeth with periapical radiolucencies of diameter greater than 0.5 cm (5 mm) Group 1: randomised 33 patients, analysed 32. Group 2: randomised 33 patients, analysed 33.</p>
Interventions	<p>All treatment was performed by a single operator. Rubber dam isolation. All caries were removed and endodontic access cavity prepared and canal patency was checked with a size 15K file. Then orifice openers taper 0.12 and 0.10 were used for enlarging the coronal and middle third of the canal, at speed of 350 rpm. RC-prep was used as a lubricant and 2.5% NaOCl saline as irrigants. The working length was established with K-file using apex locator and confirmed radiographically Instrumentation was carried out using 0.06 taper K3 nickel titanium rotary files in crown-down manner along with copious irrigation using 2.5% sodium hypochlorite and saline</p>

	<p>Group 1: single visit: the canals were dried with sterile paper points and obturated at the same appointment by using lateral condensation of gutta-percha and AH plus sealer. Temporary restoration was done</p> <p>Group 2: multiple visits (2 visits: the second appointment was scheduled at 1 week after the initial appointment). The canals were dried and double sealed with cavit and phosphate cement. Final obturation was made with a similar method and materials as used in Group 1</p>	
Outcomes	<ul style="list-style-type: none"> • Postoperative pain <p>Using a modified Heft-Parker VAS (0-170) at 6, 12, 24, and 48 hours. After one week from obturation, final clinical evaluation for pain was done with the vertical percussion method</p>	
Notes		
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Biased coin randomization".
Allocation concealment (selection bias)	Low risk	Quote: "Tossing coin, allocation and sequence was operated by a post-graduate student"
Blinding of outcome assessment (detection bias) All outcomes	High risk	Quote: "unblinded/open label".
Incomplete outcome data (attrition bias) All outcomes	Low risk	65/66 participants who entered the study were included in the final analysis
Selective reporting (reporting bias)	High risk	One of the outcomes mentioned in the M&M section (pain by vertical percussion method at one week) was not reported among the results
Other bias	Low risk	Comment: no other source of bias identified.

Penenis 2008

Methods	<p>Study design: randomised parallel group, clinical trial. Conducted in USA. Number of centres: 1. Postgraduate Endodontics Clinic, University of Illinois, USA Recruitment period: 3 years (August 2003 to May 2006). Funding source: research grant from the American Association of Endodontist Foundation</p>	
Participants	<p>97 participants enrolled, analysed 63 (29 males, 34 females, mean age 54 years, range 18-91) Inclusion criteria: necrotic teeth with radiographic evidence of apical periodontitis (minimum size > or = 2.0 mm x 2.0 mm) Exclusion criteria: patients younger than 18 year, pregnant, positive history of antibiotic use in the previous month, needed antibiotic for dental treatments, diabetic or with tooth previously treated Diagnostic criteria for pulpal or periapical disease: radiography and vital testing performed with cold and electric pulp test Group 1: 49 randomised, analysed 33 for PAI at 12 months; analysed 35 for sinus tract formation Group 2: 48 randomised, analysed 30 for PAI at 12 months, analysed 31 for sinus tract formation</p>	
Interventions	<p>G1: single visit. G2: multiple visits (the second at least two weeks but no more than 4 weeks after the first, in the interappointment period a paste made by mixing calcium hydroxide powder with 2% of chlorhexidine liquid was filled in the canals using a lentulo spiral) Multiple operators. Rubber dam and dental operating microscope use reported. Canal shaping: 0.06 taper K3 nickel titanium crown-down technique with RC prep as lubricant. Irrigation: 5.25% sodium hypochloride. Working length established using Root ZX electronic apex locator. Obturation with gutta-percha using warm vertical condensation technique</p>	
Outcomes	<ul style="list-style-type: none"> • Healing <p>Change in apical bone density at 1 year, the PAI was used to evaluate radiographic healing Secondary outcomes measures:</p> <ul style="list-style-type: none"> • Presence of clinical symptoms or abnormal findings at 12 months (i.e. spontaneous pain, presence of sinus tract, swelling, mobility, periodontal probing depths greater than baseline measurements, or sensitivity to percussion or palpation) and proportion of teeth in each group that could be considered improved (decreased PAI score) or healed (PAI < or = 2). 	
Notes	<p>CONSORT Clinical Trial; financial incentive to join the study</p>	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Patients were randomly assigned to the one-visit or two-visit group by using a block of random numbers generated by one of the investigators"

Penenis 2008 (Continued)

Allocation concealment (selection bias)	Low risk	Quote: "Neither the postgraduate clinician nor the patient was aware of the group assignment before agreeing to participate in the study"
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote: "The clinical and radiographic examination was performed by an endodontic resident unaware of the patient's group assignment or baseline presentation"
Incomplete outcome data (attrition bias) All outcomes	High risk	63/97 (65%) participants (33 G1, 30 G2) were analysed at 1 year follow-up 16 participants lost in group 1: 2 elected not participate in follow-up visits, 12 unable to contact, 2 treatment failures 18 participants lost in group 2: 1 elected not participate in follow-up visits, 16 unable to contact, 1 treatment failure
Selective reporting (reporting bias)	Low risk	Outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Peters 2002

Methods	Study design: quasi-randomised, parallel-group, clinical trial Conducted in The Netherlands. Number of Centres: 1. Academic Centre for Dentistry , Amsterdam, The Netherlands Recruitment period: not reported. Funding source: not reported.
Participants	39 participants (19 females and 20 males, mean age 40 years, range 19 to 86 years; ethnic groups not specified) Inclusion criteria: root with 1 canal, teeth asymptomatic that did not respond to sensitivity testing and never had endodontic treatment, root that showed radiographic evidence of periapical bone loss Exclusion criteria: maxillary molars, patients < 19 and > 86 years old Diagnostic criteria for pulpal or periapical disease: Rx evaluated with PAI score, sensitivity testing Group 1: randomised 21, analysed 21. Group 2: randomised 18, analysed 17.
Interventions	Group 1: single visit. Group 2: multiple visits (2, the second appointment 4 weeks later. In this group in the interappointment period the canals were dressed with a thick mix of calcium hydroxide in sterile saline and the cavity access filled with 2 layers of Cavit and a glass ionomer

Peters 2002 (Continued)

	restoration) 1 operator (endodontist). Use of rubber dam isolation and magnification loupes reported. Canal shaping: hand instrumentation by double flare technique. Irrigation: 2% sodium hypochlorite. Working length evaluated by Rx and electronic apex locator. Obturation: gutta-percha and AH 26 sealer in lateral condensation	
Outcomes	<ul style="list-style-type: none"> • Healing Follow-up 4.5 years. Routine evaluation during follow-up: 3, 12, 24 months to 4.5 years. The authors evaluated the treatment outcome as: score A (success: the width and contour of the periodontal ligament is normal, or there is a slight radiolucent zone around excess filling material); score B (uncertain: the radiolucency is clearly decreased but additional follow-up is not available); score C (failure: there is an unchanged, increased or new periradicular radiolucency). We considered only 2 categories: success (score A) and failure (score B and C) (see Table 1)	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Quote: "The teeth were randomly divided into two treatment groups, every second patient was assigned to group 2". Quasi-random method
Allocation concealment (selection bias)	High risk	Alternative assignment (randomly and consecutively, quasi-random method)
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote "Three experienced endodontists who had not been involved in the treatment or follow-up appointments were asked to analysed the radiographs"
Incomplete outcome data (attrition bias) All outcomes	Low risk	97.44% of participants who entered the study were included in the final analysis. 1 participant lost because his series of radiographs for imperfections of radiographic technique was excluded
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Methods	Study design: quasi-randomised parallel-group, clinical trial Conducted in India. Number of centres: not reported. Recruitment period: not reported. Funding source: not reported.	
Participants	148 participants (range 18 to 50 years; males-female not specified, ethnic groups not specified) Inclusion criteria: root with single uncomplicated canal with full formed apex (anterior teeth), non vital teeth: negative test of pulpal sensitivity by thermal stimuli prior to anaesthesia and no bleeding response on access to the pulp Exclusion criteria: patients with multiple teeth that required treatment, non-restorable teeth, teeth affected by periodontal diseases, patients < 18, affected by systemic diseases, affected by severe pain or acute periapical abscesses, using antibiotics or corticosteroids Diagnostic criteria for pulpal or periapical disease: Rx evaluated with PAI score, sensitivity testing Group 1: randomised 74, analysed 74. Group 2: randomised 74, analysed 74.	
Interventions	Group 1: single visit. Group 2: multiple visits (2, the second appointment 1 week later the first. In this group in the interappointment period the canal was sealed with a sterile dry cotton pellet and a temporary filling material) Use of rubber dam isolation, use of magnification loupe not specified. Canal shaping: combination of hand files (K files) and ProTaper, engine-driven rotary nickel titanium files using hybrid technique. Irrigation: 2.5% sodium hypochlorite. Working length evaluated by electronic apex locator and 2 or more angled radiographs. Obturation: gutta-percha cones and resin sealer using lateral condensation technique	
Outcomes	<ul style="list-style-type: none"> • Pain (Postobturation) using a VAS (0 to 4: 0 = no pain, 1 = slight pain/discomfort, 2 = moderate pain relieved by analgesics, 3 = moderate to severe pain not completely relieved by analgesics, 4 = severe pain/swelling not relieved by analgesics and required unscheduled visit) form filled by participants at 6, 24, 48 hours and 7 days after treatment • Painkiller use • Clinical examination after 1 week 	
Notes		
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Quote: "the patients were randomly assigned to either the one-visit or two-visit by using a set of random numbers generated by one of the investigators"

Rao 2014 (Continued)

Allocation concealment (selection bias)	Unclear risk	Alternative assignment (randomly and consecutively, quasi-random method)
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of participants (148/148) who entered the study were included in the final analysis
Selective reporting (reporting bias)	Unclear risk	Not specified.
Other bias	Low risk	Comment: no other source of bias identified.

Risso 2008

Methods	<p>Study design: randomised parallel-group, clinical trial. Conducted in Brazil. Number of centres: 1. Endodontics Clinic of the School of Dentistry of Federal University of Rio de Janeiro, Brazil Recruitment period: not reported. Funding source: Brazilian Grant from CAPES and FAPER.</p>
Participants	<p>121 participants enrolled, 118 included in the final analysis (48 males, 70 females, age between 11 and 18 years, mean age 13.6 years) Inclusion criteria: healthy, no analgesics, antibiotics or anti-inflammatory during the 10 days prior to the beginning of the treatment, age between 11-18 years, lower first or second permanent molar presenting complete root formation and necrotic pulp with or without symptoms, absence of periodontal disease, pulp calcification or acute dento-alveolar abscesses Exclusion criteria: age < 11 or > 18 years, teeth with haemorrhage in the canal during medication Diagnostic criteria for pulp necrosis were determined with pulp testing (Cold test), percussion-palpation, examination and direct observation of the presence or absence of haemorrhage in the canal Group 1: randomised not specified, analysed 57. Group 2: randomised not specified, analysed 61.</p>
Interventions	<p>G1: single visit. G2: multiple visit (2 visits, the second appointment 10-12 days after the first, in the interappointment period the canals were medicated with calcium hydroxide paste and sterile distilled water with lentulo spiral and the complete filling of the canal was verified through periapical radiography. A dry-sterile cotton pad was sealed in the pulp chamber with a minimum of 3 mm thickness temporary filling restoration (cavit) Single operator (first author, RPA). Rubber dam isolation, magnification loupes not reported</p>

Risso 2008 (Continued)

	Canal shaping: middle coronal preflaring with hand instruments: (initial passive instrumentation and Gates-Glidden burs). Ideal working length determined by an electronic apex locator and periapical radiography. Irrigation: 5.25% sodium hypochlorite solution, then 10% citric acid and 5.25% sodium hypochlorite solution and then 5% sodium thiosulfate. Obturation with gutta-percha cones using a lateral compaction filling technique as well as zinc oxide. Eugenol-based root canal sealer. Teeth filled with light cured resin	
Outcomes	<ul style="list-style-type: none"> • Pain Preoperative and postoperative measured with VAS (0 to 5: no = 0, yes = 1-5) in a 10-day period (6-12-24 hours during the first day and then every 24 hours during the 9 following days). VAS equal to 4 or 5 were considered flare-up	
Notes		
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Randomisation was performed using a random numbers listed in a table"
Allocation concealment (selection bias)	Unclear risk	Not reported.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	121 participants enrolled, 118/121 (97.5%) evaluated in the final analysis (3 participants were excluded because they missed the obturation visit)
Selective reporting (reporting bias)	High risk	Painkiller use not reported in the results.
Other bias	Low risk	Comment: no other source of bias identified.

Singh 2012

Methods	<p>Study design: randomised parallel-group, clinical trial.</p> <p>Conducted in India.</p> <p>Number of centres: 1. Department of Conservative Dentistry and endodontics, Darshan dental College, Udaipur, Rajasthan, India</p> <p>Recruitment period: not reported.</p> <p>Founding source: not reported.</p>
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Participants	<p>200 participants, 188 included in the final analysis (105 males, 83 females, aged between 20 and 60 years)</p> <p>Inclusion criteria: participants over 18 years of age, healthy. Only one tooth with a single root of each patient was included.</p> <p>Exclusion criteria: pregnant patients, patients taking antibiotics or corticosteroids at the time of treatment, affected by complicating systemic diseases or immunocompromised, below 18 years of age. Any tooth with periodontal disease or periapical radiolucency of more than 0.5 cm was excluded</p> <p>Diagnostic criteria for pulp vitality were based on the results of electric pulp tester and by direct clinical observation of haemorrhage in the canal, without considering the clinical diagnosis as being normal pulp, reversible pulpitis or irreversible pulpitis. All teeth had completely formed foramina and no calcified canals, which were preliminary evaluated by preoperative periapical radiographs</p> <p>Group 1: 100 participants, analysed 94.</p> <p>Group 2: 100 participants, analysed 94.</p>
Interventions	<p>G1: single visit.</p> <p>G2: multiple visit (2 visits, the second appointment 7 days after the first, no intracanal medication in the interappointment period but a dry-sterile cotton pellet was placed in the pulp chamber and the access was sealed with a temporary filling restoration, Cavit-G)</p> <p>Rubber dam isolation or use of magnification loupes not reported. Working length determined by electronic apex locator</p> <p>Canal shaping: combination of hand files and ProTaper engine-driven rotary nickel titanium files. RC prep used as lubricant. Irrigation: 2.5% sodium hypochlorite. Obturation: ProTaper universal gutta-percha and AH plus sealer using lateral compaction technique and restored with temporary restorative material, Cavit-G</p>
Outcomes	<ul style="list-style-type: none"> • Pain <p>Preoperative and postobturation at 6, 12, 24 and 48 hours after obturation) measured with Heft Parker VAS</p>
Notes	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote "Patients were randomly assigned to either one-or two-visit treatment by biased coin randomisation, a dynamic randomisation method, which was specially designed to get the same number in both groups and the sequence..."
Allocation concealment (selection bias)	Low risk	Quote: "... tossing coin and allocation were operated by a graduate student who was blind to the nature of the study"

Singh 2012 (Continued)

Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	12/200 participants (6%, 6 from each group) were excluded from the study because they did not attend the recall visits
Selective reporting (reporting bias)	Low risk	Outcome reported.
Other bias	Low risk	Comment: no other source of bias identified.

Soltanoff 1978

Methods	Study design: randomised, parallel group clinical trial. Conducted in USA. Number of centres: 1. Department of Endodontics, New Jersey Dental School, USA Recruitment period: 20 years. Funding source: not reported.
Participants	330 participants (sex, range or mean age, ethnic group not reported) Inclusion and exclusion criteria not reported. Diagnostic criteria for pulpally or periapical disease not specified Group 1: 135 randomised; analysed 88 (pain) and 80 (radiographical healing) Group 2: 195 randomised; analysed 193 (pain) and 186 (radiographical healing)
Interventions	Group 1: single visit. Group 2: multiple visits (the medication and the total number of visits were not specified) Rubber dam, use of magnification loupes and working length not reported. Canal shaping using root canal files In both groups, sterile saline solution was used as irrigation, the canals were filled with gutta-percha cones and Ostby's Kloroperka as the cementing medium for lateral condensation
Outcomes	<ul style="list-style-type: none"> • Pain Incidence, severity and duration: less than 1 day, 1 to 3 days, 4 to 7 days, more than 1 week. Pain was categorised as no pain, mild pain, moderate pain, severe pain. We considered only 2 categories: no pain and pain (mild, moderate, severe pain) <ul style="list-style-type: none"> • Painkiller use • Healing Observed radiographically in periods ranging from 6 months to 2 years postoperatively. The criteria for success or failure were: healed (success) and non-healed (failure)
Notes	
<i>Risk of bias</i>	

Soltanoff 1978 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Cases selected at random are the basis of this report"
Allocation concealment (selection bias)	Unclear risk	Not reported.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Not reported.
Incomplete outcome data (attrition bias) All outcomes	Low risk	85.1% of participants who entered the study were included in the final analysis about pain and use of painkillers, 80.6% of participants who entered the study were included in the final analysis about healing
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Trope 1999

Methods	Study design: randomised, parallel group clinical trial. Conducted in USA. Number of Centres: 1. School of Dentistry, University of North Carolina, USA Recruitment period: not reported. Funding source: not reported.
Participants	81 participants (54 females and 27 males had a mean age of 44.6 years, with a range of 19 to 79) with 102 teeth (61 participants had a single teeth to treat, 18 had 2, 2 had 3) Inclusion criteria: presence of radiographically demonstrable apical periodontitis on a single-rooted tooth or on 1 root with a single canal in a multi-rooted tooth Exclusion criteria: patients with diagnosis of diabetes, HIV infection or other immunocompromising disease, patients < 16 or > 80 years old and teeth with 2/3 of the root canal treated before enrolment Group 1: 45 single visits. Group 2: 57 multiple visits.
Interventions	Group 1: single visit. Group 2: multiple visits without any dressing (the instrumentation was completed at the first appointment, the canal was left empty, the treatment was completed after 1 week). 1 operator, 9 observers (4 graduate oral and maxillofacial radiology residents, 2 graduate endodontic residents, 1 oral epidemiologist, 1 general dentist, 1 experienced endodontist) to evaluate radiographs using the PAI scoring system Rubber dam isolation used. Use of magnification loupes and canal shaping not specified in a satisfactory way. Irrigation with 2.5% sodium hypochlorite. Working length evaluated

Trope 1999 (Continued)

	by Rx, obturation with gutta-percha and Roth 801 sealer in lateral condensation	
Outcomes	<ul style="list-style-type: none"> • Healing Follow-up 52 weeks. The criteria for success or failure were the following: success (PAI 1 or 2), failure (PAI 3, 4, 5) (see Table 1)	
Notes		
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "The patients were assigned to a treatment group by the throwing of a die"
Allocation concealment (selection bias)	Low risk	Unclear who performed the throw of a dice.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote "The 9 observers that participated in the study (...) were all blinded to the treatment groups and aims of the study"
Incomplete outcome data (attrition bias) All outcomes	Low risk	100% of patients who entered the study were included in the final analysis
Selective reporting (reporting bias)	Low risk	Planned outcomes reported.
Other bias	High risk	It is not explicitly stated that participants with more than one tooth needing treatment, were randomised the same number of times

Wang 2010

Methods	<p>Study design: Randomised parallel-group, clinical study.</p> <p>Conducted in: China.</p> <p>Number of Centres: 1. Endodontic Centre, West China Dental Hospital Sichuan University, China</p> <p>Recruitment period: 3 years.</p> <p>Founding source: not reported.</p>
Participants	<p>100 patients (56 women, 44 men, aged 20 to 60 years).</p> <p>Inclusion criteria: Patients over 18 years of age with vital single-rooted teeth with pre-operative vital pulps. One tooth only for each participant</p> <p>Exclusion criteria: Pregnant patients, patients taking antibiotics or corticosteroids at the time of treatment, with immuno-compromised, complicating systemic disease or under 18 years of age. Any tooth with periodontal disease or periapical pathosis was also excluded</p> <p>Diagnostic criteria for pulpal vitality based on electric pulp tester and thermal test.</p>

	<p>Vitality of pulps was confirmed by direct clinical observation of haemorrhage in the canal. All teeth had completely formed foramina and no calcified canals, evaluated with preoperative radiography</p> <p>Group 1: 50 participants randomised, analysed 43.</p> <p>Group 2: 50 participants randomised, analysed 46.</p>
Interventions	<p>Group 1: One visit.</p> <p>Group 2: Multiple visit (2 appointments, the second 1 week later the first, intermedication with calcium hydroxide paste, a sterile dry cotton pellet and 3.0 mm of Caviton)</p> <p>Two experienced operators. Rubber dam isolation or use of magnification loupes not reported</p> <p>Canal shaping: combination of hand files and ProTaper engine-driven rotary nickel titanium files. Irrigation: 2.5% sodium hypochlorite. Working length verified using Root ZX II apex locator. Obturation: ProTaper universal gutta-percha and AH plus sealer, using a lateral compaction technique</p>
Outcomes	<ul style="list-style-type: none"> • Pain (Verbal descriptor scale-VDS: preoperative and postoperative at 6-24-48 h and 1 week after the treatment). Pain categorised as no pain, slight pain, moderate pain, strong pain, severe pain, maximum pain • Flare-up and swelling
Notes	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote "Patients were randomly assigned to either one-or two-visit treatment by biased coin randomisation, a dynamic randomisation method, which was specially design to get the same number in both groups and the sequence..."
Allocation concealment (selection bias)	Low risk	Quote: "...tossing coin and allocation were operated by someone who was a graduate student and was not aware of the nature of the study"
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	11/100 (11%) were excluded from the study. 11 patients (7 from Group 1 and 4 from Group 2) were excluded from the analysis: 7 (5 from G1 and 2 from G2) did not at-

Wang 2010 (Continued)

		tend the recall visits, 2 (1 from G1 and 1 from G2) failed to return the forms used to monitor postobturation pain, and 2 (1 from G1 and 1 from G2) had more than one root canal
Selective reporting (reporting bias)	Low risk	Outcome reported.
Other bias	Low risk	Comment: no other source of bias identified.

Weiger 2000

Methods	Study design: Randomised, parallel group clinical trial. Conducted in: Germany. Number of Centres: 1. Department of Conservative Dentistry, University of Tübingen, Germany Recruitment period: Duration of the study: 5 years. Funding source: not reported.
Participants	73 participants recruited, 67 entered final analysis (37 females and 30 males, mean age 38 years range: 11 to 84) Inclusion criteria: teeth with periapical lesion radiographically demonstrated and where the vitality test was negative; in each patients only 1 tooth was selected Exclusion criteria: teeth having pockets communicated with the lesion, teeth treated previously, patients that had taken antibiotics 4 weeks prior to the treatment Diagnostic criteria for pulpally or periapical disease: Rx and vitality test Group 1: randomised not specified, analysed 36. Group 2: randomised not specified, analysed 31.
Interventions	Group 1: Single visit. Group 2: Multiple visits (2 visits, the interappointment medication used was calcium hydroxide mixed with sterile physiological saline, that was left in the canals for 7-47 days. The cavity access was filled by a temporary cement) 2 operators (experienced endodontists). Use of rubber dam isolation. Use of magnification loupes not reported. Canal shaping: K- files and Gates Glidden used in step-back technique. Irrigation: 1% sodium hypochlorite. Working length determined by Rx. Obturation: gutta-percha with Sealapex in lateral condensation
Outcomes	<ul style="list-style-type: none"> • Healing Follow-up 5 years. The criteria for success or failure were the following in the paper: complete healing, incomplete healing, no healing. The radiographs were judged by both dentists involved in the study by using a magnifying glass and a light box. The operators did not know whether the tooth belonged to the 1-visit or the 2-visit group. In case of disagreement a joint decision was made. We considered only 2 categories: success (complete healing) and failure (incomplete healing and no healing)
Notes	

Weiger 2000 (Continued)

<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: " The allocation of the tooth modality followed the method of minimization to balance the two groups of teeth with regard to the criterion tooth type"
Allocation concealment (selection bias)	Unclear risk	Not reported.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote: "The radiographs were judged by both dentists involved in the study, the operators did not know whether the tooth belonged to the one-visit or the two-visits group"
Incomplete outcome data (attrition bias) All outcomes	Low risk	91.7% of participants who entered the study were included in the final analysis 6 patients lost at follow-up (5 did not return at recall appointments, 1 deceased prior to the first scheduled recall rate)
Selective reporting (reporting bias)	Low risk	Outcome reported.
Other bias	Low risk	Comment: no other source of bias identified.

Wong 2015

Methods	Study design: Randomised, parallel group clinical trial. Conducted in: China. Number of Centres: 1. Health Service Dental Clinic of the University of Hong Kong Recruitment period: 30 months. Funding source: not reported.
Participants	256 teeth from 228 participants: 115 in group 1 and 113 in group 2. A total of 194 participants with 220 teeth entered final analysis (85 males, 135 females, aged from 46.38 ± 14.06) Inclusion criteria: participants without history of periodontitis, tooth that required primary endodontic treatment was periodontally healthy and at least half of the coronal structure had to be remaining Exclusion criteria: teeth with pulpotomy, participants who had severe acute pulpitis with facial swelling or systemic infection, severe systemic disease, increased stress on the temporomandibular joint musculature or increased psychological stress Group 1: randomised 128 teeth from 115 participants, analysed 117 teeth from 105 participants

	Group 2: randomised 128 teeth from 113 participants, analysed 103 teeth from 89 participants	
Interventions	<p>Two general dentists carried out the endodontic treatments. Rubber dam isolation. One of the operator was trained to use a magnifying loupe (2.5x). The two dentists received a calibration workshop prior to this clinical trial to standardise the instrumentation and obturation technique</p> <p>The root canals were cleaned and shaped using Ni-Ti rotary files. A 5.25 % sodium hypochlorite was used for irrigation. Temporary restoration was made using a resin-modified zinc oxide and eugenol cement until obturation. All teeth were obturated using a core-carrier technique. The total chairside time was recorded by the dental assistant. The treated teeth were restored with silver amalgam or composite resin. All patients were reviewed 1 week after obturation, and were advised to have indirect extracoronal restoration (partial or full veneer) to avoid failure due to extra-coronal leakage or tooth fracture</p> <p>Group 1: One visit. Group 2: Multiple visits (2 or 3 appointments, depending on the complexity of the treatment, the second one scheduled one week after the first). A non-setting 5 % calcium hydroxide paste was used as interappointment medication</p>	
Outcomes	<ul style="list-style-type: none"> • Success or failure of endodontic treatment (Success was graded as no clinical signs/symptoms and no radiographic radiolucency found in the periapical radiograph) • Pain at 1 week after obturation: 10-point Likert scale (0 to 10) • Radiological assessment according to Chu 2005, based on absence-presence of periapical radiolucency (see Table 1). Multiple-rooted teeth with different periapical statuses at different roots were classified according to the most severe periapical condition. 	
Notes	The clinical trial was registered in the Chinese Clinical Trial Registry of the World Health Organization (ChiCTR-IOR-15006117)	
<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "The treated teeth were randomly assigned to either single-visit or multiple-visit treatments using the random-number generating function of a calculator"
Allocation concealment (selection bias)	Low risk	Quote: "patients were randomly assigned by the receptionist for endodontic treatment"
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Quote: "A number unknown to the operators and the independent assessor was given to each treated tooth for clinical and radiographic assessment, data entry and analy-

Wong 2015 (Continued)

		sis”
Incomplete outcome data (attrition bias) All outcomes	High risk	36 teeth (14.6) from 34 patients (14.1%) were lost at follow at follow-up
Selective reporting (reporting bias)	Low risk	Outcomes reported.
Other bias	Low risk	Comment: no other source of bias identified.

Xiao 2010

Methods	Study design: Randomised, parallel group clinical trial. Conducted in: China. Number of Centres: 1. Shijitan Hospital of Beijing, Beijing, China Recruitment period: February 2005 to February 2006. Funding source: not reported.
Participants	138 teeth from 86 participants: 76 in group 1 and 62 in group 2 Not clear the exact number of males and females who were enrolled in the study Inclusion criteria: tooth with pulp necrosis, acute or chronic periapical periodontitis and had only one root canal Exclusion criteria: tooth with out-of-control intra-root canal exudation, periapical periodontitis with space infection, retreatment, resorption of the alveolar bone which exceeded 1/2 of the root, patients with systemic disease and that could not keep follow-up Group 1: randomised 76, analysed 76 (mean age 49.6±14.7). Group 2: randomised 62, analysed 62 (mean age 45.7±13.7).
Interventions	Group 1: One visit. Group 2: Multiple visit (2 appointments, the second 1 week later the first, interappointment medication with calcium hydroxide paste) Use of rubber dam and magnification loupes not specified. Root canal was prepared with ProTaper, work length was detected by Root-ZX, irrigated with 2% sodium hypochlorite. The restorative materials were Cortisomol and gutta-percha
Outcomes	<ul style="list-style-type: none"> ● Pain Evaluated before treatment, immediately after treatment and 7 days after treatment <ul style="list-style-type: none"> ● Cured No symptom, well-functioned, no clinical sign, PAI level was 1 to 2 <ul style="list-style-type: none"> ● Relieved No symptom, well-functioned, no clinical sign, PAI level stayed the same or decreased <ul style="list-style-type: none"> ● Failed Symptoms, with red or swelling of the periapical area, sinus formation, tenderness and periapical indisposed after percussion. PAI level increased
Notes	Healing: authors evaluated the treatment outcome as: cured (no symptom, well-functioned, no clinical sign, PAI level was 1-2), relieved (no symptom, well-functioned, no clinical sign, PAI level stayed the same or decreased), failed (with symptom, with red

Xiao 2010 (Continued)

	or swelling of the periapical area, sinus formation, tenderness and periapical indisposed after percussion. PAI level increased) Pain postobturation as reported by participant, measured by VAS and than dichotomised, and after percussion test immediately after treatment and 7 days after the treatment
Risk of bias	
Bias	Authors' judgement
Random sequence generation (selection bias)	Low risk
Allocation concealment (selection bias)	Low risk
Blinding of outcome assessment (detection bias) All outcomes	Low risk
Incomplete outcome data (attrition bias) All outcomes	Low risk
Selective reporting (reporting bias)	Low risk
Other bias	High risk
	Quote: all the participants were allocated by tossing coins. Quote: all the participants were allocated by tossing coins. Quote: the x-rays of pretreatment, 6 months, 1 year and 2 years were numbered and were assessed by 2 radiologists and a clinician blindly. Thus, only one component (x-ray evaluation) of the composite outcome (healing) was blinded to the assessors 100% of participants who entered the study were included in the final analysis Outcomes reported. Participants with more that one tooth needing RoCT were treated alternatively with the two methods

Yoldas 2004

Methods	Study design: Randomised, parallel group clinical trial. Conducted in: Turkey. Number of Centres: 1. Department of Conservative Dentistry and Endodontics, Faculty of Dentistry, University of Cukurova, Turkey Recruitment period: 2 years. Funding source: not reported.
Participants	227 participants (sex and ethnic group not reported; age over 18 years) Inclusion criteria: teeth with inadequate root canal filling Exclusion criteria: patients with complicating systemic disease, severe pain or acute apical abscess or both, under 18 years of age, use of antibiotics or corticosteroids, multiple teeth requiring retreatment, root canals that could not be treated well with initial RoCT Diagnostic criteria for pulpal or periapical disease: evaluation of periapical status with Rx evaluated by 1 author according to PAI

	<p>Group 1: randomised 115, analysed 106. Group 2: randomised 112, analysed 112. 9 patients (7 from Group 1 and 2 from Group 2) were excluded from the study because did not participate to recall visits</p>
Interventions	<p>Group 1: Single visit. Group 2: Multiple visits (2 visits, the second appointment 7 days after the first). The canals in the interappointment period were medicated with calcium hydroxide chlorhexidine paste and closed with a sterile dry cotton pellet and a temporary restorative material (Cavit) for 7 days) 3 operators. Use of rubber dam and magnification loupes not specified. Canal shaping with Gates Glidden, hand files nickel titanium rotary instruments with step-back technique. Irrigation: 2.5% sodium hypochlorite. Working length: determined by apex locator and periapical radiograph. Obturation: gutta-percha and AH 26 sealer with lateral condensation</p>
Outcomes	<ul style="list-style-type: none"> • Pain (1 week after initial appointment the patients were recalled and asked about the occurrence of postoperative pain): the level of discomfort was rated as follows: no pain, mild pain, moderate pain, severe pain. We considered only 2 categories: no pain, pain (mild, moderate, severe) • Flare-up Participants with severe postoperative pain or occurrence of swelling or both (see Table 2). • Painkiller use
Notes	

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Quote: "randomisation of assignment into the treatment groups was made by 1 author according to the PAI, the tooth type and patient symptoms"
Allocation concealment (selection bias)	High risk	Assignment by 1 of the authors.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Comment: not applicable, radiographic assessment was not among outcomes
Incomplete outcome data (attrition bias) All outcomes	Low risk	218/227 (96%) of participants who entered the study were included in the final analysis
Selective reporting (reporting bias)	Low risk	Outcomes reported.

Yoldas 2004 (Continued)

Other bias	Low risk	Comment: no other source of bias identified.
------------	----------	--

AH-26 and AH-plus: root canal filling and sealer materials

EDTA: ethylene-diamine-tetraacetic acid

ISO: International Organization for Standardization

M&M: material and methods

PAI: periapical index

RC-prep: chemo-mechanical preparation for root canals

RC-help: lubricating paste for root canals

Rx: radiological evaluation

VAS: visual analogue scale

Root-ZX: apex locator

Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
ElMubarak 2010	This study is not randomised or quasi-randomised.
Fava 1989	This study is not randomised or quasi-randomised.
Fava 1994	This study design was unclear. We sent an e-mail to authors asking for more details about their randomisation method but we did not consider the answer satisfactory to consider the paper randomised or quasi-randomised
Friedman 1995	This study is not randomised or quasi-randomised.
Gurgel-Filho 2007	This study evaluates only the microbiological aspects of RoCT
Holland 2003	This study evaluates single versus multiple-visit RoCT in dogs
Jabeen 2014	This study is not randomised or quasi-randomised.
Kabaktchieva 2013	The study included children and primary teeth.
Kvist 2004	This study evaluates only the microbiological aspects of RoCT
Ng 2004	This study is not randomised or quasi-randomised.
Orhan 2010	The study included children and primary teeth.
Papworth B 1998	This study is not randomised or quasi-randomised.

(Continued)

Prashanth 2011	The study is not randomised or quasi-randomised.
Roane 1983	This study is not randomised or quasi-randomised.
Silveira 2007	This study evaluates single versus multiple-visit RoCT in dogs
Trusewicz 2005	This study considers only the microbiological aspects of RoCT
Vera 2012	This study evaluates only the microbiological aspects of RoCT
Waltimo 2005	The study does not include any of the outcomes considered in the review
Walton 1992	This study is not randomised or quasi-randomised.
Xavier 2013	This study evaluates only the microbiological aspects of RoCT

RoCT: root canal treatment

DATA AND ANALYSES

Comparison 1. Single versus multiple visits for endodontic treatment of permanent teeth - primary outcomes

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Extraction due to endodontic problems	1		Risk Ratio (M-H, Random, 95% CI)	Totals not selected
2 Radiological failure	11		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
2.1 Radiological failure	11	1493	Risk Ratio (M-H, Random, 95% CI)	0.91 [0.68, 1.21]
2.2 Radiological failure in necrotic teeth	8	823	Risk Ratio (M-H, Random, 95% CI)	0.82 [0.55, 1.21]

Comparison 2. Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Pain (dichotomous)	12		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
1.1 Pain in the immediate postoperative period (until 72 hours postobturation)	9	1560	Risk Ratio (M-H, Random, 95% CI)	0.99 [0.84, 1.17]
1.2 Pain in the immediate postoperative period in necrotic teeth (until 72 hours postobturation)	6	718	Risk Ratio (M-H, Random, 95% CI)	0.84 [0.62, 1.16]
1.3 Pain in the immediate postoperative period in vital teeth (until 72 hours postobturation)	3	318	Risk Ratio (M-H, Random, 95% CI)	1.04 [0.86, 1.27]
1.4 Pain at 1 week	8	1383	Risk Ratio (M-H, Random, 95% CI)	1.50 [0.99, 2.28]
1.5 Pain at 1 week in necrotic teeth	2	172	Risk Ratio (M-H, Random, 95% CI)	0.70 [0.17, 2.85]
1.6 Pain at 1 week in vital teeth	1	89	Risk Ratio (M-H, Random, 95% CI)	1.40 [0.78, 2.52]
1.7 Pain at 1 month	2	513	Risk Ratio (M-H, Random, 95% CI)	0.0 [0.0, 0.0]
1.8 Pain at 18 months	1	220	Risk Ratio (M-H, Random, 95% CI)	0.88 [0.06, 13.90]
2 Pain (continuous)	4		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.1 Pain in the immediate postoperative period (until 72 hours postobturation)	4	414	Mean Difference (IV, Random, 95% CI)	0.12 [-0.59, 0.82]
2.2 Pain in the immediate postoperative period in necrotic teeth (until 72 hours postobturation)	3	158	Mean Difference (IV, Random, 95% CI)	-0.21 [-0.44, 0.03]

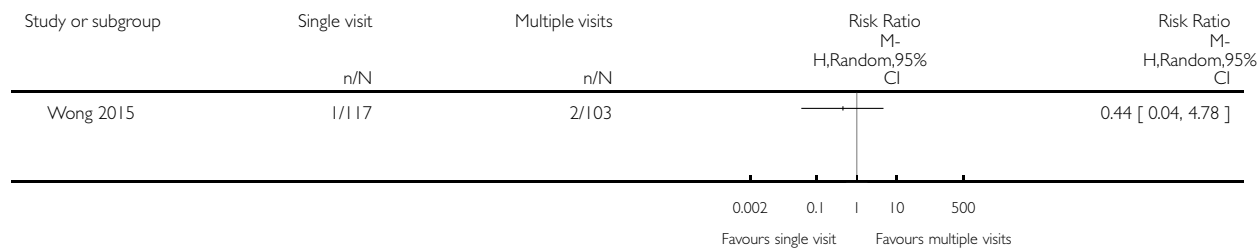
2.3 Pain in the immediate postoperative period in vital teeth (until 72 hours postobturation)	1	30	Mean Difference (IV, Random, 95% CI)	0.60 [-1.03, 2.23]
3 Swelling or flare-up	4		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
3.1 Swelling or flare-up	4	281	Risk Ratio (M-H, Random, 95% CI)	1.36 [0.66, 2.81]
3.2 Swelling or flare-up in necrotic teeth	2	120	Risk Ratio (M-H, Random, 95% CI)	1.52 [0.70, 3.31]
3.3 Swelling or flare-up in vital teeth	1	89	Risk Ratio (M-H, Random, 95% CI)	1.07 [0.07, 16.57]
4 Painkiller use	4	648	Risk Ratio (M-H, Random, 95% CI)	2.35 [1.60, 3.45]
5 Persistent sinus track or fistula	2	345	Risk Ratio (M-H, Random, 95% CI)	0.98 [0.15, 6.48]
6 Any complication	10		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
6.1 Any complication	10	1686	Risk Ratio (M-H, Random, 95% CI)	0.92 [0.77, 1.11]
6.2 Any complication in necrotic teeth	9	1201	Risk Ratio (M-H, Random, 95% CI)	0.88 [0.68, 1.13]
6.3 Any complication in vital teeth	3	485	Risk Ratio (M-H, Random, 95% CI)	1.02 [0.83, 1.25]

Analysis 1.1. Comparison 1 Single versus multiple visits for endodontic treatment of permanent teeth - primary outcomes, Outcome 1 Extraction due to endodontic problems.

Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 1 Single versus multiple visits for endodontic treatment of permanent teeth - primary outcomes

Outcome: 1 Extraction due to endodontic problems

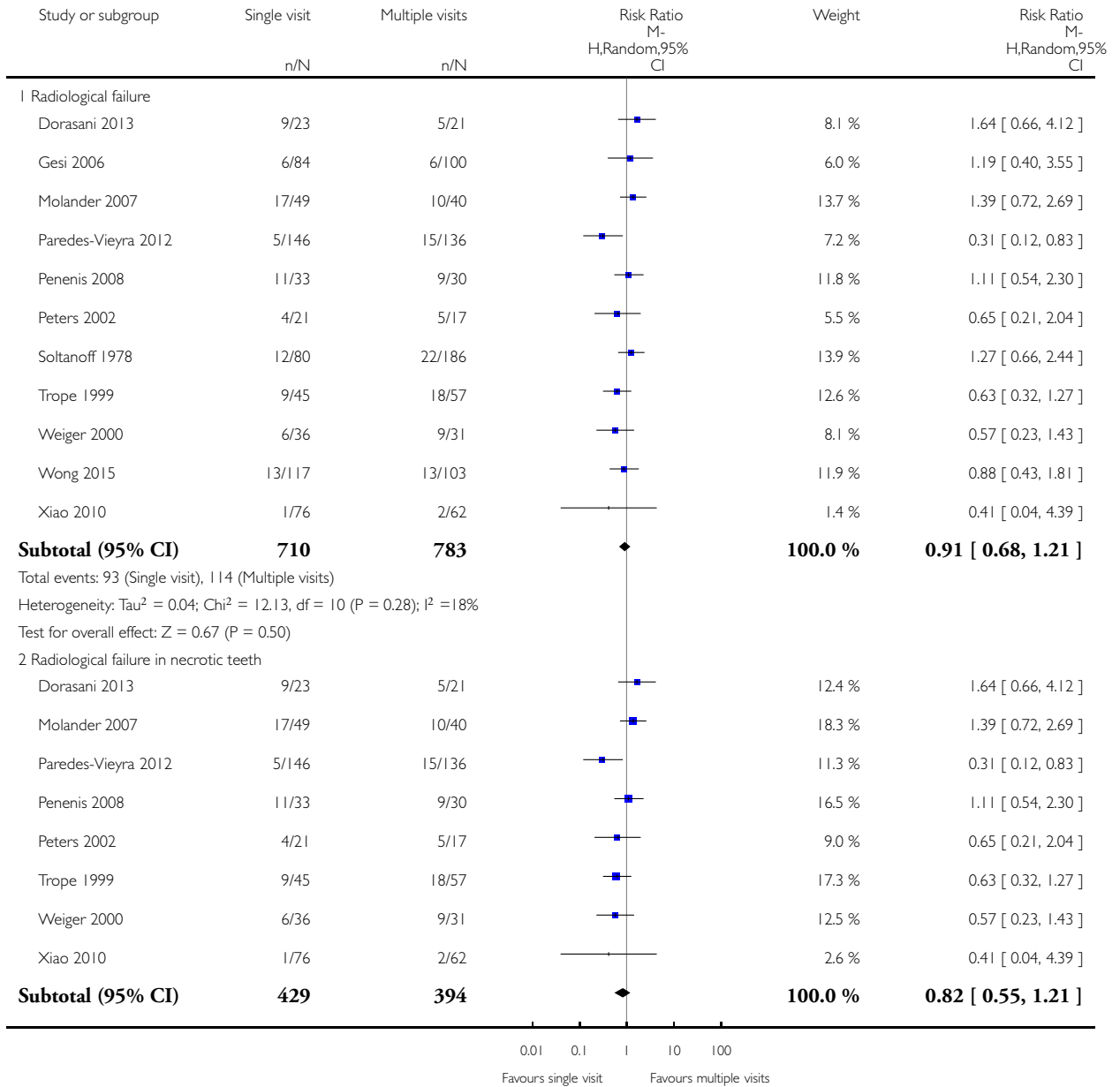


Analysis 1.2. Comparison 1 Single versus multiple visits for endodontic treatment of permanent teeth - primary outcomes, Outcome 2 Radiological failure.

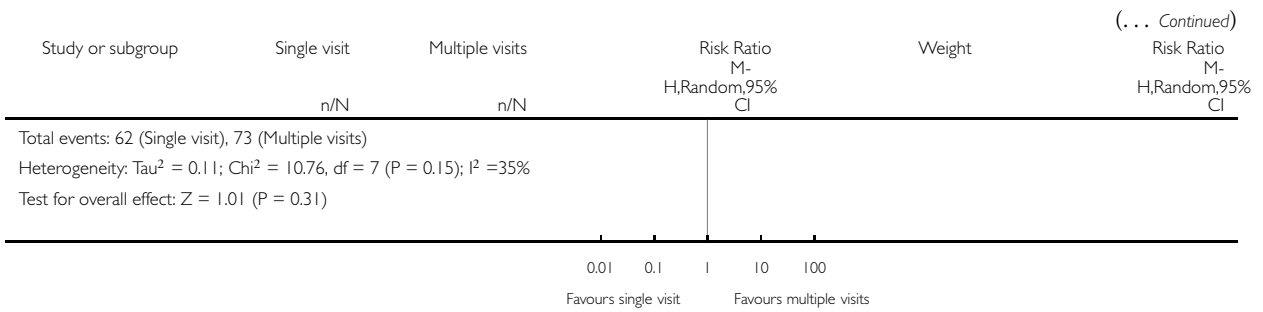
Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 1 Single versus multiple visits for endodontic treatment of permanent teeth - primary outcomes

Outcome: 2 Radiological failure



(Continued ...)

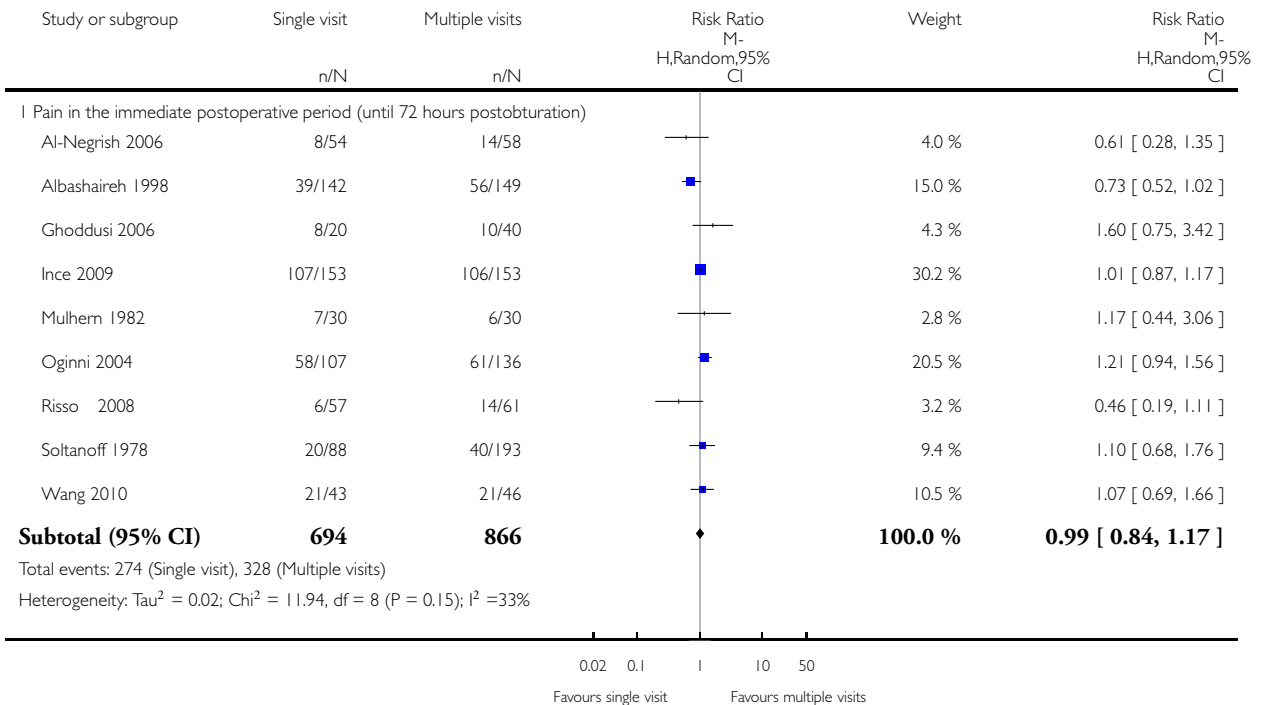


Analysis 2.1. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 1 Pain (dichotomous).

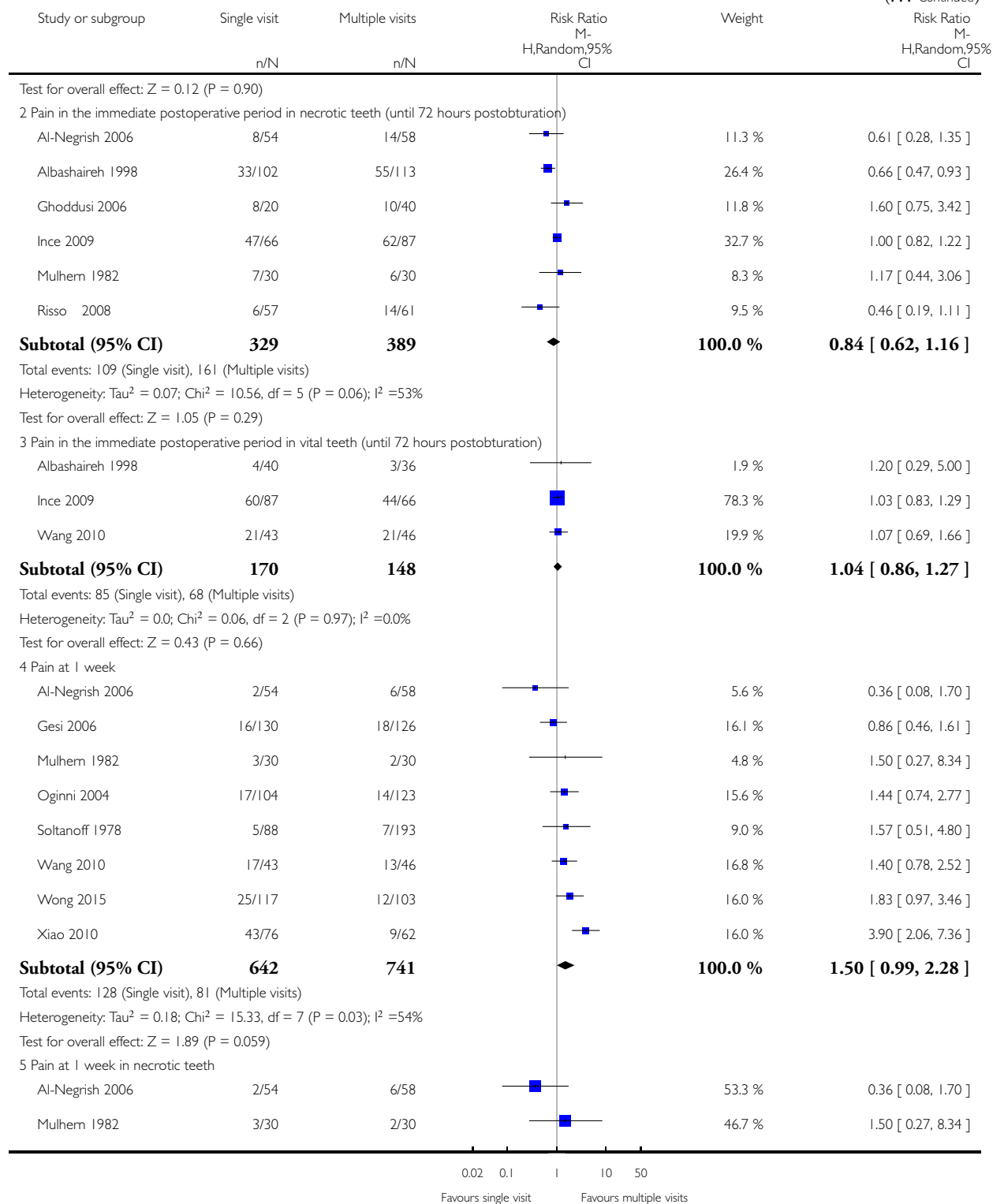
Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

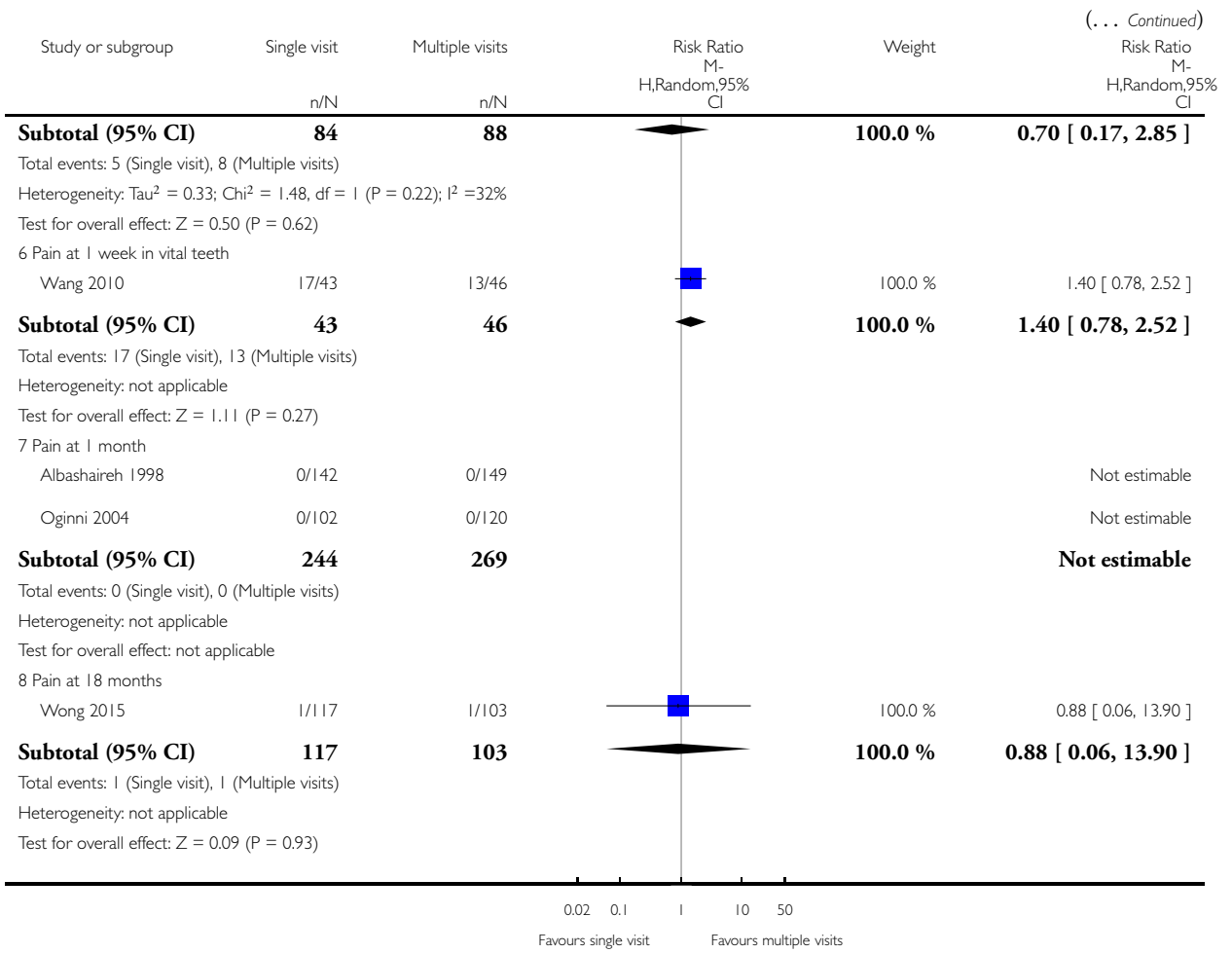
Outcome: 1 Pain (dichotomous)



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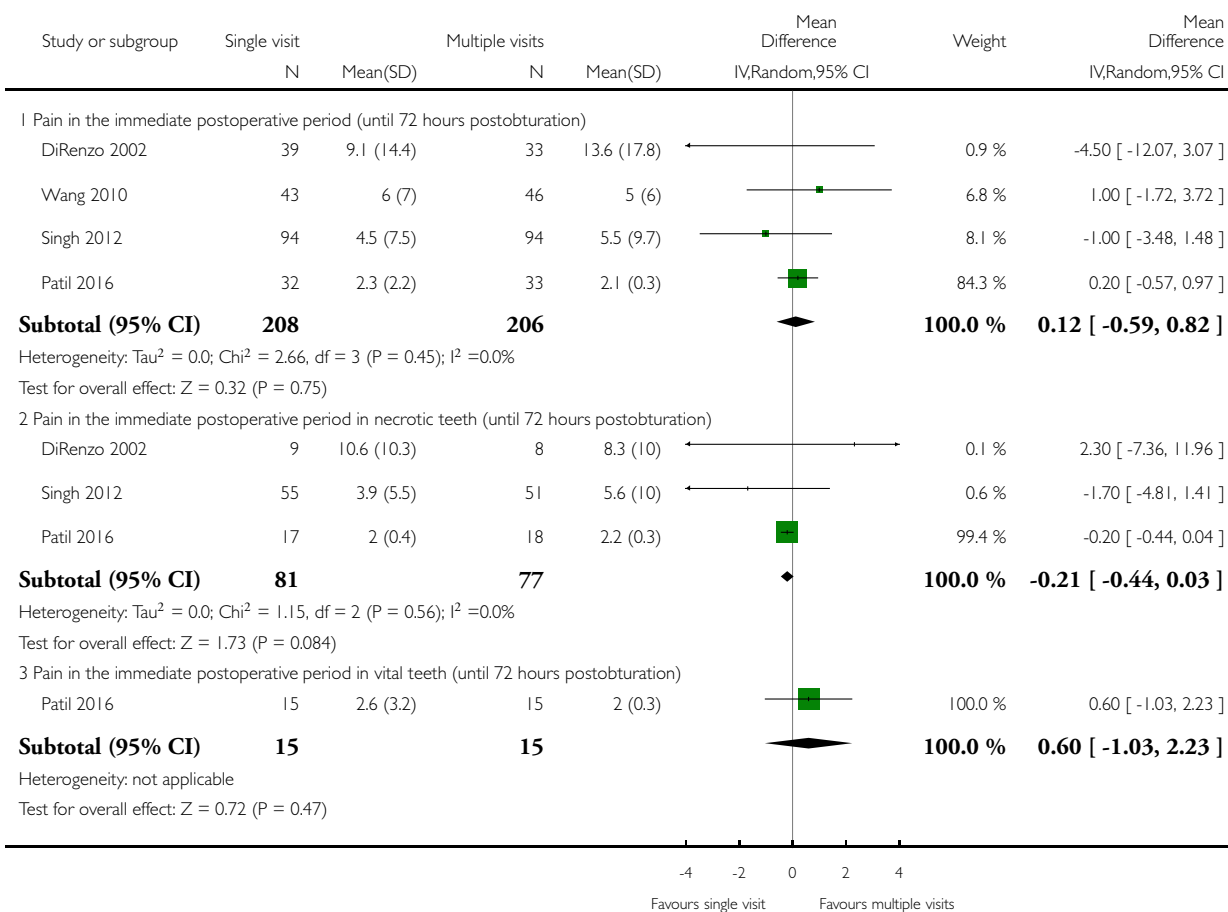


Analysis 2.2. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 2 Pain (continuous).

Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome: 2 Pain (continuous)

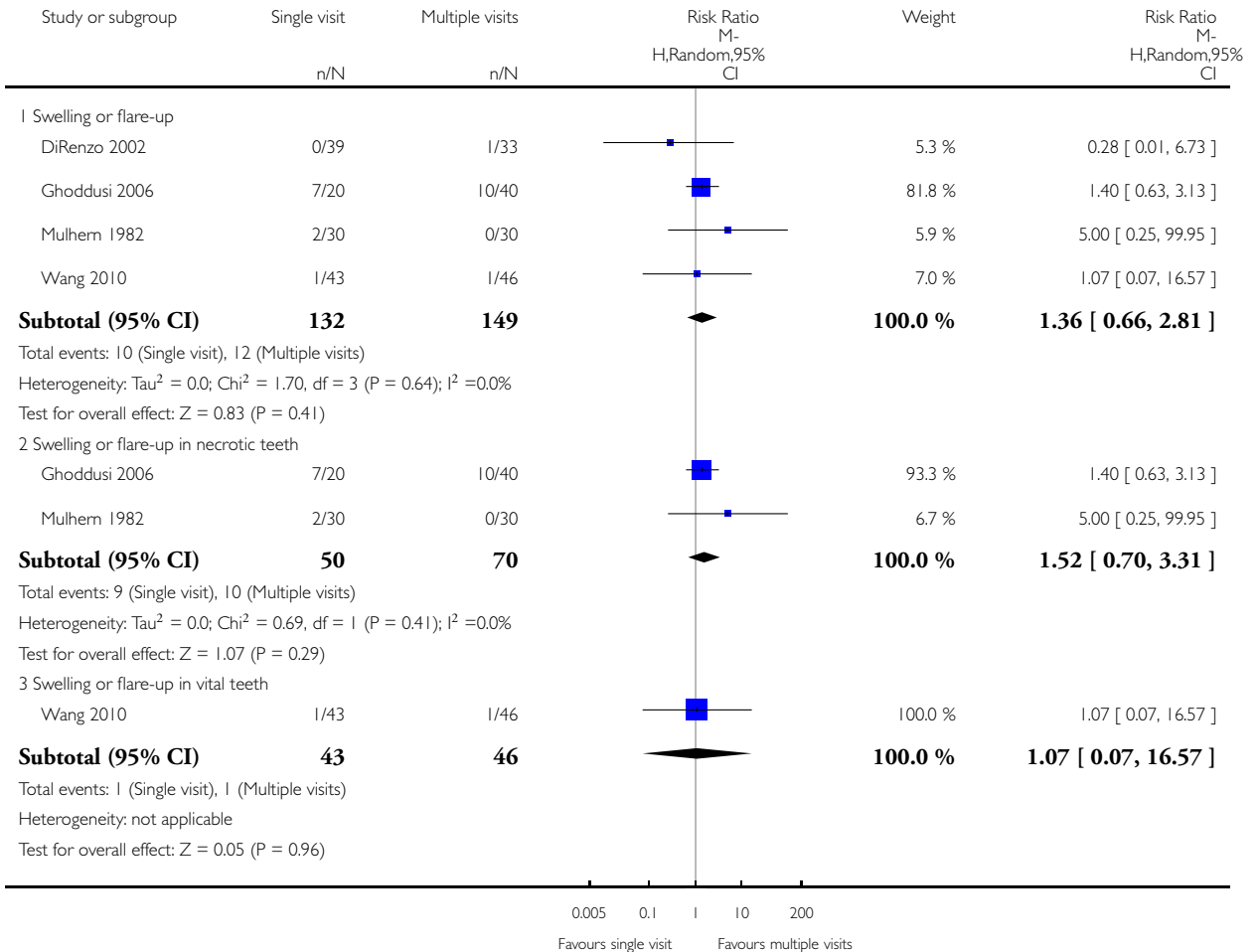


Analysis 2.3. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 3 Swelling or flare-up.

Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome: 3 Swelling or flare-up

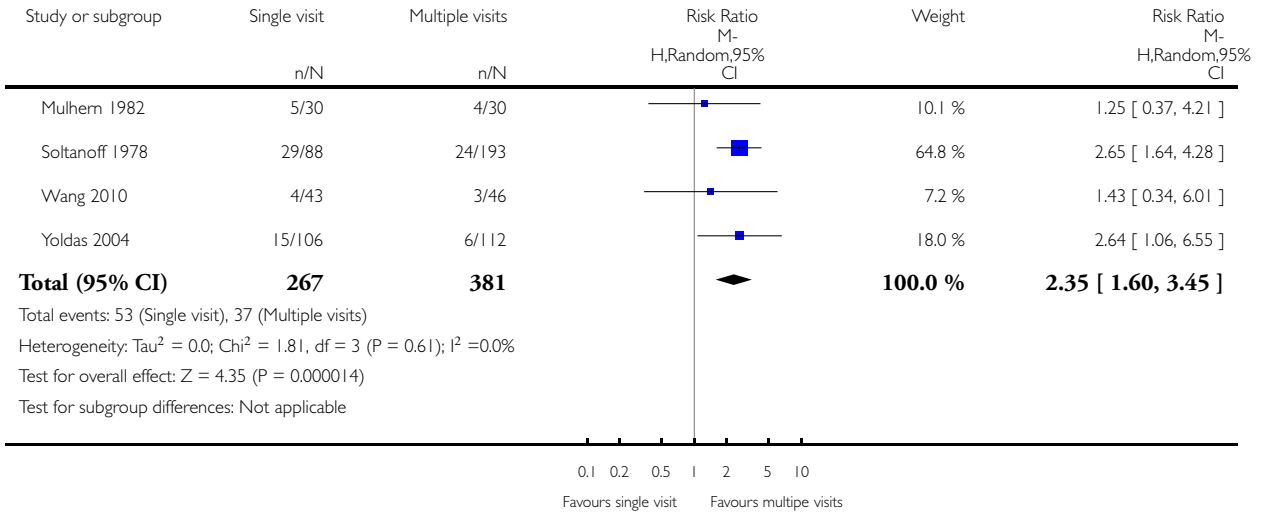


Analysis 2.4. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 4 Painkiller use.

Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome: 4 Painkiller use

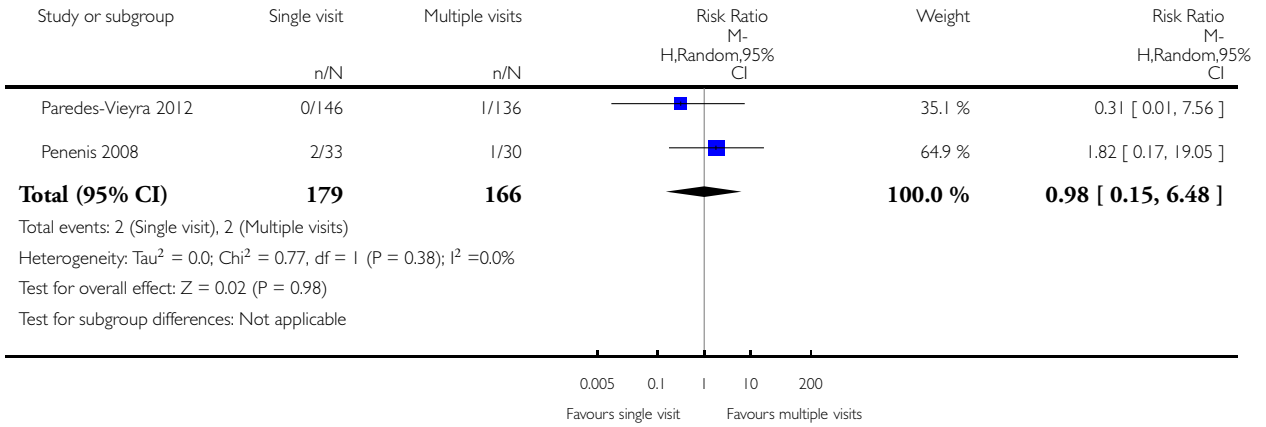


Analysis 2.5. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 5 Persistent sinus track or fistula.

Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome: 5 Persistent sinus track or fistula

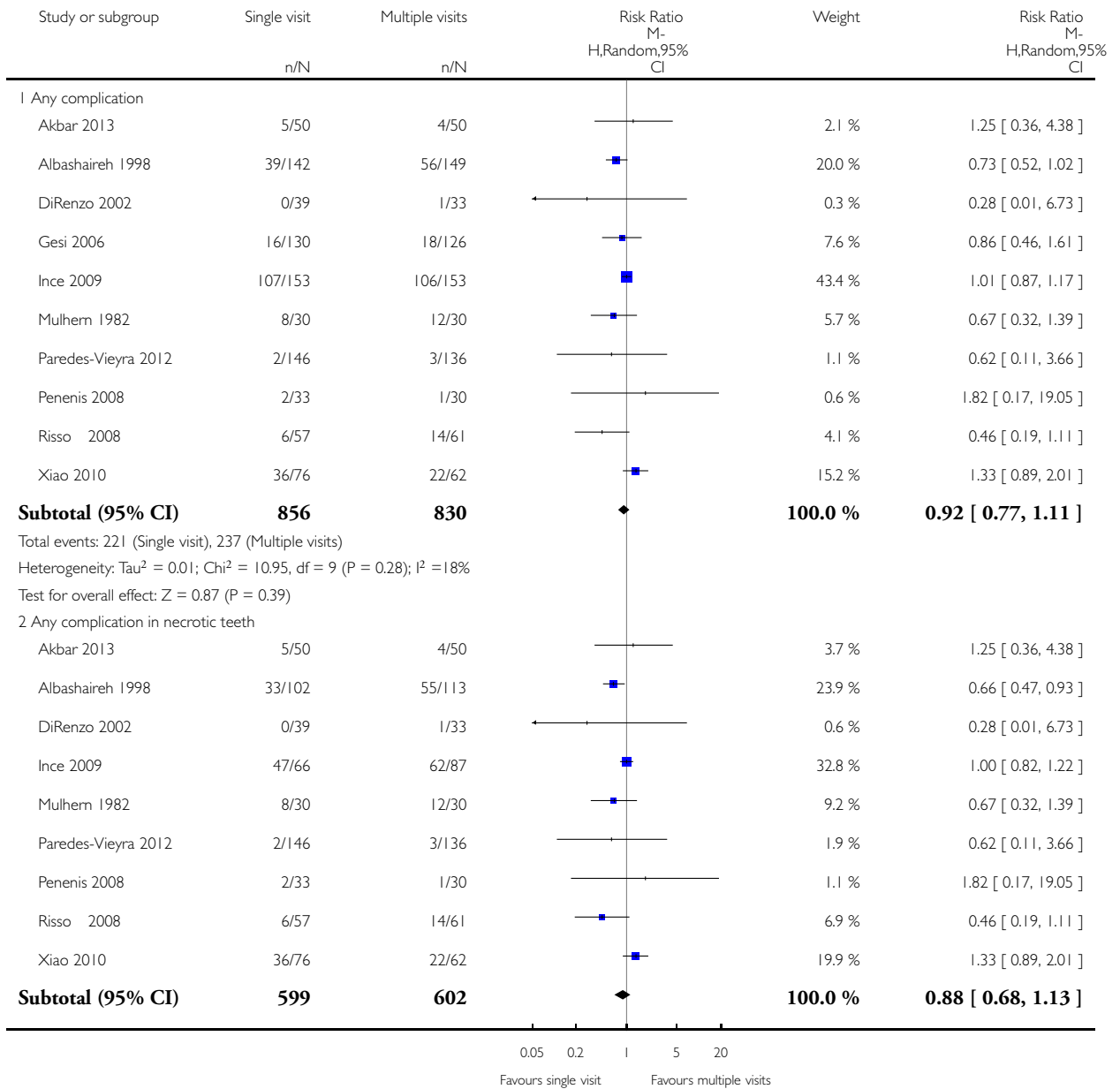


Analysis 2.6. Comparison 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes, Outcome 6 Any complication.

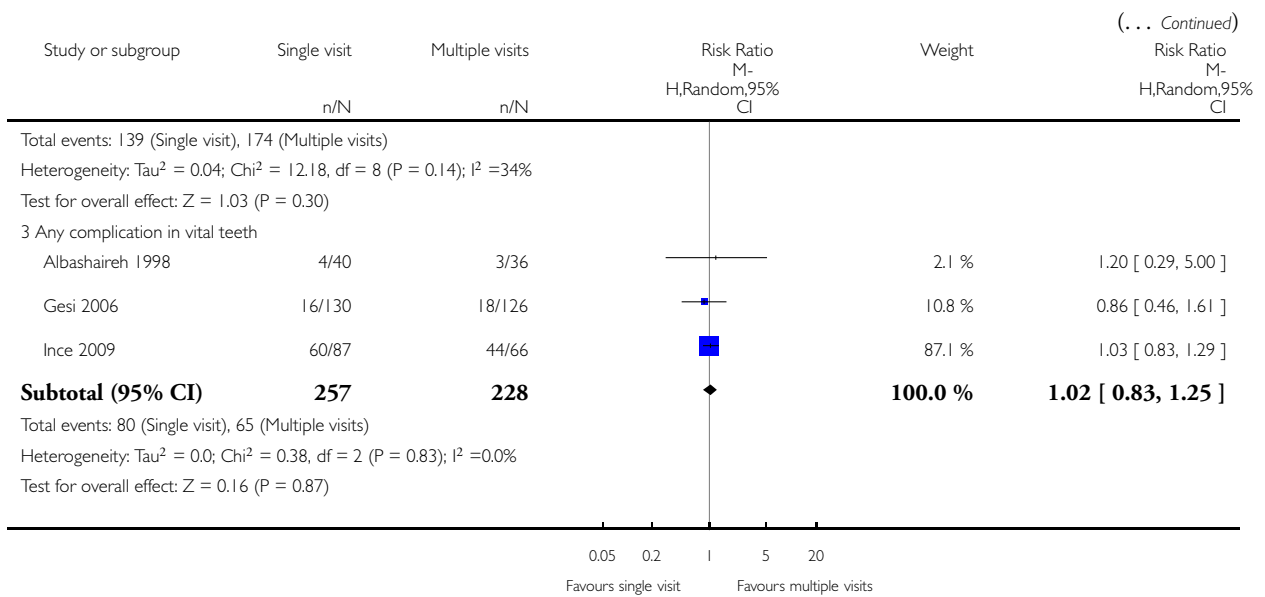
Review: Single versus multiple visits for endodontic treatment of permanent teeth

Comparison: 2 Single versus multiple visits for endodontic treatment of permanent teeth - secondary outcomes

Outcome: 6 Any complication



(Continued ...)



ADDITIONAL TABLES

Table 1. Endodontic radiological success and failure: from scales to binary outcome

Classification	Success (binary)	Failure (binary)
Trope 1999 ; Orstavik 1991 ; Orstavik 1998	PAI score 1 (normal periapical), PAI score 2 (bone structural changes)	PAI score 3 (structural changes with mineral loss), PAI score 4 (radiolucency), PAI score 5 (radiolucency with features of exacerbation)
Strinberg 1956	Success (normal to slightly thickened periodontal ligament space < 1 mm, elimination of previous rarefaction, normal lamina dura in relation to adjacent teeth, no evidence of resorption)	Questionable (increased periodontal ligament space > 1 mm and < 2 mm, stationary rarefaction or slight repair evident, increased lamina dura in relation to adjacent teeth, evidence of resorption); failure (increased width of periodontal ligament space > 2 mm, lack of osseous repair within rarefaction or increased rarefaction, lack of new lamina dura, presence of osseous rarefactions in periradicular areas where previously none existed)
Katebzadeh 2000	Healed (normal pattern of trabecular bone and normal width of periodontal ligament space)	Improved (reduction in lesion size); failed (increased or no change in the lesion size)

Table 1. Endodontic radiological success and failure: from scales to binary outcome (Continued)

Halse 1986	Healed (normal pattern of trabecular bone and normal width of periodontal ligament space)	Increased width of the periodontal space, pathological findings
Peters 2002; Kvist 2004	Success (A) the width and contour of the periodontal ligament is normal, or there is a slight radiolucent zone around apical	Uncertain (B) the radiolucency is clearly decreased but additional follow-up is not available; failure (C) there is an unchanged, increased, or new periradicular radiolucency
Weiger 2000	Complete healing: no clinical signs and symptoms, radiographically a periodontal ligament space of normal width	Incomplete healing: no clinical signs and symptoms, radiographically a reduction of the lesion in size or an unchanged lesion within an observation time of 4 years. No healing: clinical signs and symptoms indicating an acute phase of apical periodontitis and/or radiographically a persisting lesion after a follow-up time of 4 to 5 years and/or a new lesion formed at an initially uninvolved root of a multi-rooted tooth
Soltanoff 1978	Healed (by Rx but the criteria not specified in a satisfactory way)	Not healed (by Rx but the criteria not specified in a satisfactory way)
Gesi 2006	Normal periapical condition or unclear apical condition (widened apical periodontal space or diffused lamina dura)	Presence of periapical radiolucency when there was a distinct radiolucent area associated with the apical portion of the root
Petersson 1991; Chu 2005	Normal - normal appearance of the surrounding osseous structure	Apical periodontitis - periapical radiolucency observed Periapical status not classified - the quality of the radiograph was insufficient for examination of the periapical structure

PAI: periapical index
Rx: radiological evaluation

Table 2. Definition of flare-up in the included studies

Study	Definition of flare-up
Akbar 2013	Moderate to severe postoperative pain or moderate to severe swelling that begins 12 to 48 hours after treatment and lasts at least 48 hours
Al-Negrish 2006	Percentage of participants experiencing moderate to severe pain evaluated after 2 and 7 days

Table 2. Definition of flare-up in the included studies (Continued)

DiRenzo 2002	Swelling that needs antibiotics and narcotic analgesics
Ghoddusi 2006	Swelling
Mulhern 1982	Swelling
Oginni 2004	Participant report of pain not controlled with over-the-counter medication or increasing swelling or both
Risso 2008	Intensive pain (visual analogue scale > 4)
Wang 2010	Swelling
Yoldas 2004	Severe postoperative pain or occurrence of swelling or both

APPENDICES

Appendix I. Cochrane Oral Health's Trials Register search strategy

Updated searches of Cochrane Oral Health's Trials Register were undertaken using the Cochrane Register of Studies and the search strategy below:

#1 ((endodontic* OR "root canal" OR pulp* OR "root fill*"):ti,ab) AND (INREGISTER)

#2 ("single visit*" OR single-visit* OR "single appoint*" OR single-appoint* OR "single session*" OR single-session* OR "multiple visit*" OR multiple-visit* OR "multiple appoint*" OR multiple-appoint* OR "multiple session*" OR multiple-session* OR "first visit*" OR first-visit* OR "1st visit*" OR 1st-visit* OR "one visit" OR one-visit OR "first appointment" OR "one appointment" OR one-appointment OR "first session*" OR "one session" OR one-session OR "second visit*" OR "2nd visit*" OR "two visit*" OR two-visit* OR "two appointment*" OR two-appointment* OR "two session*" OR "two-session*" OR "third visit*" OR third-visit OR "three visit*" OR three-visit* OR "third appointment*" OR third-appointment* OR "three appointment*" OR three-appointment* OR "three session*" OR three-session* OR "single and multiple visit*" OR "one and two visit*"):ti,ab) AND (INREGISTER)

#3 (#1 and #2) AND (INREGISTER)

Previous searches of Cochrane Oral Health's Trials Register were undertaken using the Procite software and the search strategy below: ((endodontic* OR "root canal" OR pulp* OR "root fill*") AND ("single visit*" OR single-visit* OR "single appoint*" OR single-appoint* OR "single session*" OR single-session* OR "multiple visit*" OR multiple-visit* OR "multiple appoint*" OR multiple-appoint* OR "multiple session*" OR multiple-session* OR "first visit*" OR first-visit* OR "1st visit*" OR 1st-visit* OR "one visit" OR one-visit OR "first appointment" OR "one appointment" OR one-appointment OR "first session*" OR "one session" OR one-session OR "second visit*" OR "2nd visit*" OR "two visit*" OR two-visit* OR "two appointment*" OR two-appointment* OR "two session*" OR "two-session*" OR "third visit*" OR third-visit OR "three visit*" OR three-visit* OR "third appointment*" OR third-appointment* OR "three appointment*" OR three-appointment* OR "three session*" OR three-session* OR "single and multiple visit*" OR "one and two visit*"))

Appendix 2. Cochrane Central Register of Controlled Trials (CENTRAL) search strategy

- #1 MeSH descriptor Endodontics explode all trees
- #2 (endodontic* in All Text or pulpectom* in All Text or pulpotom* in All Text)
- #3 “root canal” in All Text
- #4 ((pulp in All Text near/4 cap* in All Text) or (pulp in All Text near/4 devital* in All Text))
- #5 (#1 or #2 or #3 or #4)
- #6 (single next visit* in All Text or single next appointment* in All Text or single next session* in All Text)
- #7 (multi* next visit* in All Text or multi* next appointment* in All Text or multi* next session* in All Text)
- #8 (first next visit* in All Text or first next appointment* in All Text or first next session* in All Text or second next visit* in All Text or second next appointment* in All Text or second next session* in All Text or third next visit* in All Text or third next appointment* in All Text or third next session* in All Text)
- #9 (1st next visit* in All Text or 1st next appointment* in All Text or 1st next session* in All Text or 2nd next visit* in All Text or 2nd next appointment* in All Text or 2nd next session* in All Text or 3rd next visit* in All Text or 3rd next appointment* in All Text or 3rd next session* in All Text)
- #10 (one next visit* in All Text or one next appointment* in All Text or one next session* in All Text or two next visit* in All Text or two next appointment* in All Text or two next session* in All Text or three next visit* in All Text or three next appointment* in All Text or three next session* in All Text)
- #11 (#6 or #7 or #8 or #9 or #10)
- #12 (#5 and #11)

Appendix 3. MEDLINE Ovid search strategy

- 1 ENDODONTICS/
- 2. exp ROOT CANAL THERAPY/
- 3. dental pulp capping/ or pulpectomy/ or pulpotomy/
- 4. (endodontic\$ or pulpectom\$ or pulpotom\$)
- 5. (root canal adj (therapy or treat\$))
- 6. ((pulp adj3 cap\$) or (pulp\$ adj3 devitali\$))
- 7. or/1-6
- 8. (single adj (visit\$ or appointment\$ or session\$))
- 9. (multi\$ adj (visit\$ or appointment\$ or session\$))
- 10. ((first or second or third) adj (visit\$ or appointment\$ or session\$))
- 11. ((1st or 2nd or 3rd) adj (visit\$ or appointment\$ or session\$))
- 12. ((one or two or three) adj (appointment\$ or visit\$ or session\$))
- 13 or/8-12
- 14. 7 and 13

Appendix 4. Embase Ovid search strategy

- 1 ENDODONTICS/
- 2. exp ROOT CANAL THERAPY/
- 3. dental pulp capping/ or pulpectomy/ or pulpotomy/
- 4. (endodontic\$ or pulpectom\$ or pulpotom\$)
- 5. (root canal adj (therapy or treat\$))
- 6. ((pulp adj3 cap\$) or (pulp\$ adj3 devitali\$))
- 7. or/1-6
- 8. (single adj (visit\$ or appointment\$ or session\$))
- 9. (multi\$ adj (visit\$ or appointment\$ or session\$))
- 10. ((first or second or third) adj (visit\$ or appointment\$ or session\$))
- 11. ((1st or 2nd or 3rd) adj (visit\$ or appointment\$ or session\$))
- 12. ((one or two or three) adj (appointment\$ or visit\$ or session\$))
- 13 or/8-12

14. 7 and 13

Appendix 5. US National Institutes of Health Ongoing Trials Register (ClinicalTrials.gov) and the WHO International Clinical Trials Registry Platform search strategy

endodontic and visit
endodontic and appointment
endodontic and session

WHAT'S NEW

Last assessed as up-to-date: 14 June 2016.

Date	Event	Description
9 November 2016	New citation required but conclusions have not changed	Our substantive conclusions have not changed. On the basis of the available evidence, it seems likely that the benefit of a single-visit treatment, in terms of time and convenience, for both patient and dentist, has the cost of a higher frequency of late postoperative pain (and as a consequence, painkiller use) One of the newly included studies reported data on tooth loss, which was no different between the two approaches (single visit and multiple visits)
14 June 2016	New search has been performed	Searches updated. Thirteen new studies added to the previous version of the review, making a total of 25

HISTORY

Protocol first published: Issue 2, 2005

Review first published: Issue 4, 2007

Date	Event	Description
31 July 2008	Amended	Converted to new review format.

CONTRIBUTIONS OF AUTHORS

- Maddalena Manfredi: main review author, participation in all phases of the review's preparation.
- Lara Figini: participation in all phases of the review's preparation.
- Massimo Gagliani: manuscript preparation, interpretation of results.
- Giovanni Lodi: group co-ordinator, contributor in all phases of the review's preparation, article retrieval, data collection, interpretation of results.

DECLARATIONS OF INTEREST

- Maddalena Manfredi: none known.
- Lara Figini: none known.
- Massimo Gagliani: none known.
- Giovanni Lodi: none known.

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Internal sources

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External sources

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- Cochrane Oral Health Group Global Alliance, Other.

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DIFFERENCES BETWEEN PROTOCOL AND REVIEW

The outcomes were modified from the protocol in the last version of the review. We added 'any complication' as a secondary outcome in this version.

INDEX TERMS

Medical Subject Headings (MeSH)

*Dentition, Permanent; Analgesics [*therapeutic use]; Anti-Bacterial Agents [therapeutic use]; Appointments and Schedules; Dental Pulp Necrosis [diagnostic imaging; *therapy]; Office Visits [*utilization]; Pain, Postoperative [etiology]; Pulpitis [diagnostic imaging; *therapy]; Radiography; Randomized Controlled Trials as Topic; Root Canal Therapy [adverse effects; *methods]; Tooth Extraction; Treatment Outcome

MeSH check words

Humans