Pulpotomy versus pulpectomy in the treatment of vital pulp exposure in primary incisors. A systematic review and meta-analysis. [version 3; peer review: 2 approved, 1 approved with reservations]

Lamia Gadallah\textsuperscript{1}, Mahmoud Hamdy\textsuperscript{2}, Adel El Bardissy\textsuperscript{2}, Mohamed Abou El Yazeed\textsuperscript{1}

\textsuperscript{1}Orthodontics and Pediatric Dentistry Department, National Research Centre, Egypt, Nasr city, Cairo, 11727, Egypt
\textsuperscript{2}Pediatric Dentistry and Dental Public Health, Faculty of Dentistry, Cairo University, Cairo, Cairo, Egypt

Abstract

\textbf{Background:} Early childhood caries is a serious public health problem. When caries extend to involve the pulp, various forms of pulp treatment are tried to stimulate tooth repair. Although pulpotomy is the treatment of choice for vital primary tooth pulp exposure but there is a trend among many dentists to perform pulpectomies in vital primary incisors. This study aimed to assess the effect of pulpotomy and pulpectomy in treatment of carious vital pulp exposure in primary incisors.

\textbf{Methods:} We searched Pubmed and Cochrane library databases up to March, 2018, OpenGrey for grey literature and ClinicalTrials.gov for ongoing trials. Randomized controlled trials were included and assessed with Cochrane risk of bias tool. Primary outcomes were clinical failure and radiological failure. The effect sizes were calculated as risk ratios with 95%CI using the Mantel-Haenszel method.

\textbf{Results:} Four trials were identified for qualitative assessment, only three trials were included in meta-analysis after exclusion of one trial due to its high risk of bias. The pooled results of the longest follow up period for clinical failure showed no statistically significant difference between pulpotomy and pulpectomy. The relative risk (RR) was e 2.69, 95% CI 0.76 to 9.58 for clinical failure. For radiographic failure, the sensitivity analysis showed RR 0.45, 95% CI 0.25 to 0.83 with a higher risk for radiographic failure in pulpectomy. The evidence was limited by the small number of trials included in the meta-analysis.

\textbf{Conclusions:} Both pulpotomy and pulpectomy can be used successfully in the treatment of vital pulp exposure in primary incisors.

Open Peer Review

\textbf{Reviewer Status}

\begin{tabular}{|c|c|c|c|}
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\textbf{Invited Reviewers} & 1 & 2 & 3 \\
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version 3 & report & report & report \\
(version) 25 Jun 2019 &  \\
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version 2 &  \\
(version) 20 Dec 2018 &  \\
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version 1 & report & report &  \\
26 Sep 2018 &  \\
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1. Mariem O. Wassel, Ain Shams University, Cairo, Egypt
2. Armin Shirvani, Shahid Beheshti University of Medical Sciences, Tehran, Iran
3. Marcela Baraúna Magno, Federal University of Rio de Janeiro, Rio de Janeiro, Brazil

Any reports and responses or comments on the article can be found at the end of the article.
incisors. Further high quality studies comparing between pulpotomy and pulpectomy in primary incisors with longer follow up period till exfoliation time are needed.

**Keywords**
Pulpotomy, Pulpectomy, Root Canal Therapy, Primary Incisor, Vital Pulp Exposure
Amendments from Version 2
In this version we edited the abstract to add the risk of bias tool and effect measures used. We corrected two lines in the search strategy and added the radiographic assessment in the Conclusion section.
See referee reports

Introduction
Dental caries is an international public health challenge, especially among young children. Early childhood caries (ECC) is a serious public health problem in both developing and industrialized countries. ECC has been considered to be at epidemic proportions in the developing world. Treatment of ECC can be accomplished through different types of intervention. When caries extend to involve the pulp, various forms of pulp treatment have been used to treat and/or remove the pulp or to stimulate tooth repair. The choice of technique is as important as the choice between the different pharmacotherapeutic agents which are used in the treatment of primary teeth.

Although pulpotomy is the treatment of choice for vital primary tooth pulp exposure throughout the pediatric dental literature, the current trend amongst many dentists is to perform pulpectomies for the pulp treatment of carious vital primary anterior teeth.

The most common materials used for pulpotomy are formocresol, ferric sulfate, also calcium hydroxide has been used, but with less long term success and more recently mineral trioxide aggregate (MTA) which is much more expensive.

According to the latest recommendations of the American Academy of Pediatric Dentistry, both formocresol and MTA are strongly recommended to be used in pulpotomy with moderate quality of evidence. Ferric sulfate, laser and sodium hypochlorite are conditionally recommended but as for calcium hydroxide there is a recommendation against its use in pulpotomy.

In pulpectomy, a resorbable material such as nonreinforced zinc/oxide eugenol (ZOE), a combination paste of iodoform and calcium hydroxide (Vitapex, Metapex) or a combination paste of zinc oxide and eugenol, iodoform and calcium hydroxide (Endoflas) are used to fill the canals.

According to a Cochrane systematic review, there was no conclusive evidence supporting the superiority of one material for use in pulpectomy. Zinc oxide and eugenol, Metapex and Endoflas were found to be equally effective while with low quality of evidence zinc oxide and eugenol may be better than Vitapex but further research is required for confirmation.

There are few studies that have compared pulpectomies with pulpotomies in vital primary incisors. Therefore the claim that pulpotomies don’t work in primary anterior teeth is not supported by high-quality evidence from research. Moreover two studies recently showed that there was no significant difference in success rates of pulpotomies and pulpectomies in the pulp treatment of asymptomatic vital primary incisors.

We therefore aimed to determine in patients with carious vital pulp exposure in primary incisors if pulpotomy is better than pulpectomy in terms of pain, soft tissue pathology, tooth mobility, pathological root resorption, periapical radiolucency, pulp canal obliteration and tooth survival based on evidence from randomized controlled trials.

Methods
Criteria for considering studies for this review
Types of participants. Children with carious vital pulp exposure in primary incisors.

Types of interventions. Pulpotomy and pulpectomy (root canal treatment) techniques with different medicaments.

Types of outcome measures
Primary outcomes. We defined two primary outcomes: clinical failure and radiological failure.

Secondary outcomes. According to the core set of component outcomes as specified by Smail-Faugeron et al., these secondary outcomes were considered:

Secondary clinical outcomes: pain, soft tissue pathology (gingival swelling, fistulous tract), pathological mobility and tooth survival.

Secondary radiographical outcomes: pathological radiolucency, pathological root resorption, pulp canal obliteration.

Types of studies
Randomized controlled trials comparing pulpotomy and pulpectomy techniques in the treatment of carious vital pulp exposure in primary incisors were included.

Search methods
Electronic search. We searched the electronic databases as the Cochrane library to 1/3/2018 and PubMed to 1/3/2018. We developed detailed search strategies for each database searched. We placed no restrictions on the date of publication when searching the electronic databases. The search strategy included appropriate keywords, and Mesh terms when applicable; combined with Boolean operators “AND”, “OR” and “NOT” as shown in Table 1 and Table 2. We also searched OpenGrey for grey literature and ClinicalTrials.gov for ongoing trials.

Hand searching. We hand searched the reference lists of the included full text articles.

Data collection and analysis
Selection of studies. Two review authors (LG and AEB) independently scanned the titles of all reports identified by the search to determine whether the studies were relevant. They independently scanned selected abstracts to determine whether the study was relevant. After obtaining the full report for all relevant articles, they independently scanned the full reports and
### Table 1. Index terms used in search with synonyms.

<table>
<thead>
<tr>
<th>PICO item</th>
<th>Item</th>
<th>Synonyms</th>
</tr>
</thead>
<tbody>
<tr>
<td>P</td>
<td>Patients with carious vital pulp exposure in primary anterior teeth</td>
<td>Primary teeth (tooth) Deciduous teeth (tooth) Milky teeth (tooth) Baby teeth (tooth) Incisor(s) Anterior teeth (tooth) Vital teeth (tooth) Pulpally exposed teeth (tooth)</td>
</tr>
<tr>
<td>I</td>
<td>pulpotomy</td>
<td>Pulpotomy Pulpotomies Vital pulp therapy Dental pulp exposure</td>
</tr>
<tr>
<td>C</td>
<td>pulpectomy</td>
<td>Pulpectomy Pulpectomies Root canal therapy</td>
</tr>
</tbody>
</table>

### Table 2. Detailed search strategy for Pubmed and Cochrane library.

<table>
<thead>
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<th>Item</th>
<th>Pubmed 11/6</th>
<th>Cochrane 11/6</th>
</tr>
</thead>
<tbody>
<tr>
<td>#1</td>
<td>Tooth, deciduous (Mesh Term)</td>
<td>10215 411</td>
</tr>
<tr>
<td>#2</td>
<td>Incisor</td>
<td>22967 1012</td>
</tr>
<tr>
<td>#3</td>
<td>Incisors</td>
<td>28128</td>
</tr>
<tr>
<td>#4</td>
<td>Anterior teeth</td>
<td>11087 700</td>
</tr>
<tr>
<td>#5</td>
<td>Anterior tooth</td>
<td>8907</td>
</tr>
<tr>
<td>#6</td>
<td>Vital teeth</td>
<td>2445 339</td>
</tr>
<tr>
<td>#7</td>
<td>Vital tooth</td>
<td>1923</td>
</tr>
<tr>
<td>#8</td>
<td>Pulpally exposed teeth</td>
<td>5 4</td>
</tr>
<tr>
<td>#9</td>
<td>Pulpally exposed tooth</td>
<td>4</td>
</tr>
<tr>
<td>#10</td>
<td>#2 OR #3 OR #4 OR #5</td>
<td>34393 1495</td>
</tr>
<tr>
<td>#11</td>
<td>#6 OR #7 OR #8 OR #9</td>
<td>2449 341</td>
</tr>
<tr>
<td>#12</td>
<td>#1 AND #10 AND #11</td>
<td>22 6</td>
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<tr>
<td>#13</td>
<td>Pulpotomy</td>
<td>1416 152</td>
</tr>
<tr>
<td>#14</td>
<td>Pulpotomies</td>
<td>1442</td>
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<tr>
<td>#15</td>
<td>Vital pulp therapy</td>
<td>722 88</td>
</tr>
<tr>
<td>#16</td>
<td>Dental pulp exposure</td>
<td>1564 120</td>
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<tr>
<td>#17</td>
<td>#13 OR #14 OR #15 OR #16</td>
<td>331 282</td>
</tr>
<tr>
<td>#18</td>
<td>#12 AND #17</td>
<td>11 4</td>
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<tr>
<td>#19</td>
<td>Pulpectomy</td>
<td>1277 117</td>
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<td>#20</td>
<td>Pulpectomies</td>
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<td>#21</td>
<td>Root canal therapy</td>
<td>18604 696</td>
</tr>
<tr>
<td>#22</td>
<td>#19 OR #20 OR #21</td>
<td>19281 779</td>
</tr>
<tr>
<td>#23</td>
<td>#12 AND #22</td>
<td>8 4</td>
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<tr>
<td>#24</td>
<td>#18 AND #23</td>
<td>13 4</td>
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</table>
completed the data extraction form to determine whether the article should be included or excluded. Disagreements at each stage were resolved by discussion. If this did not resolve the disagreement a third author was invited to settle the disagreement. This did not occur over the course of this study.

Data extraction and management. Two review authors (LG and AEB) collected the data independently using a specially designed data extraction form (Dataset 1).

For each trial, the following data were recorded: the year of publication, the country where the trial took place, detailed description of methodology, sample size, mean age of participants, duration of follow-up and reported outcomes. We contacted the authors of randomized controlled trials for missing information if needed.

Assessment of risk of bias in included studies. Two review authors (LG and AEB) independently assessed the risk of bias in the included trials. The assessment was according to Cochrane risk of bias tool for quality assessment of randomized controlled trials described in the Cochrane Handbook for Systematic Reviews of Interventions 5.1.0 (updated March 2011)\textsuperscript{18}. Any disagreements between the two authors were resolved by discussion. If this did not resolve the disagreement a third author was invited to settle the disagreement. This did not occur over the course of this study.

Measures of treatment effect. Estimated effect size was calculated as risk ratios with 95% Confidence Interval (CI) for dichotomous outcomes. The unit of analysis was the tooth, because teeth were randomly assigned to intervention.

Data synthesis. The effect sizes and associated 95% confidence intervals were calculated using the Mantel-Haenszel method. If the results from trials were homogenous then fixed effect model was preferred. The statistical analysis was performed with the Review Manager program v.5.3 (RevMan)\textsuperscript{19}.

Results

By searching the different databases, 14 references were identified after removal of the duplicates. By scanning the titles and abstracts, 4 studies were included as shown in the Prisma flow diagram Figure 1 and data extraction was performed (completed PRISMA checklist is available as Supplementary File 1).

Figure 1. PRISMA Flow diagram. A list of included articles is shown in Table 3.
The trials were. trial 9.21.9. Following caries removal and pulp exposure, the trial was completed in one session, whether they were performed under general or local anesthesia.

**Results of studies**

**Clinical failure.** Clinical failure was reported to be 2% and 3% for pulpotomy at 12 months and 18 months respectively, while for the pulpectomy there were none at 12 months, and 1% at the 18 months follow up period in the Nguyen et al. trial. No clinical failures were reported for neither pulpotomy nor pulpectomy at 12 months, and 1% at the 18 months follow up period in the Nguyen et al. trial.

<table>
<thead>
<tr>
<th>Study characteristics of included studies</th>
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<tbody>
<tr>
<td><strong>Year of publication, setting and operators.</strong></td>
<td>The trials were published between 2004 and 2017. Two trials were in Canada by Nguyen et al., and Casas et al., one trial in the United states of America by Howley et al. and one in Iran by Aminabadi et al. Operators were dentists in the four trials.</td>
</tr>
<tr>
<td><strong>Participants.</strong></td>
<td>The age range of participants with carious vital pulp exposures in primary incisors varied varying from 18 to 60 months.</td>
</tr>
<tr>
<td><strong>Number of arms.</strong></td>
<td>All four trials by Nguyen et al., Howley et al., Aminabadi et al., and Casas et al. were two-arm studies.</td>
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<tr>
<td><strong>Duration of follow up.</strong></td>
<td>Follow up was at 12 and 24 months in two trials, by Aminabadi et al., and Casas et al. Follow up was to 23 months at three intervals: 5–9, 10–14, and 15–23 months in the trial by Howley et al. and at 12 and 18 months in the trial by Nguyen et al. Anesthesia. Three trials, Nguyen et al., Howley et al., and Casas et al., were under general anesthesia and one trial was under local anesthesia by Aminabadi et al. Rubber dam. All four trials used rubber dam isolation. Treatment. The following treatments were compared in the included trials: The Casas et al. trial used ferric sulphate in pulpotomy compared with zinc oxide and eugenol in pulpectomy. Formocresol in pulpotomy was compared with zinc oxide and eugenol in pulpectomy in the Aminabadi et al. trial. Formocresol in pulpotomy was compared with vitapex (calcium hydroxide/iodoform paste) in pulpectomy in the Howley et al. trial. Ferric sulfate and mineral trioxide aggregate in pulpotomy was compared with zinc oxide and eugenol in pulpectomy in the Nguyen et al. trial. Medicaments Pulpectomy Two trials by Howley et al. and Aminabadi et al. used formocresol following hemostasis with a cotton pellet and zinc oxide and eugenol as capping material. One trial used ferric sulphate to achieve hemostasis and zinc oxide and eugenol as capping material, Casas et al., and one trial used ferric sulfate to achieve hemostasis, with mineral trioxide aggregate as a capping material, Nguyen et al. Pulpectomy Zinc oxide and eugenol was used in three trials by Nguyen et al., Aminabadi et al., and Casas et al. One trial, Howley et al., used vitapex (calcium hydroxide/iodoform paste). Pulp access. Following caries removal and pulp exposure, the pulp chamber was accessed using a sterile no. 56 bur in a water-cooled high-speed handpiece and was refined using a sterile round bur in a slow-speed handpiece in three trials, Nguyen et al., Aminabadi et al., and Casas et al. In the trial by Howley et al. the pulp chamber was unroofed using a no. 330 sterile bur in a water-cooled high-speed handpiece then the access was refined using a sterile round bur in a slow-speed handpiece. Pulpotomy. The coronal pulp was amputated using a sharp spoon excavator in two trials by Howley et al. and Aminabadi et al. and was amputated using a sterile low-speed round bur in another two trials by Nguyen et al. and Casas et al. Root canal treatment. In three trials Nguyen et al., Howley et al. and Casas et al. pulp tissue was removed en bloc using two or more endodontic files (Hedström files or K files), if the first attempt was unsuccessful, the procedure was repeated until all of the pulp tissue was removed. In the fourth trial by Aminabadi et al. an endodontic K file was introduced to the working length after a periapical radiograph was taken, and most of the pulp tissue was removed completely on the first attempt. If the first attempt was unsuccessful, the procedure was repeated and canals were generally enlarged three sizes past the initial file. Irrigation. The irrigation solution was sterile water in Nguyen et al. and saline in the trials by Howley et al. and Aminabadi et al., while in the fourth trial by Casas et al. the irrigating solution was unidentified. Final restoration. Resin restorations was performed in three trials, Nguyen et al., Aminabadi et al., and Casas et al. Full coverage crown whether stainless steel crown (SCC) or SCC with white esthetic veneer were used in Howley et al. trial. Number of visits. In the four trials by Nguyen et al., Howley et al., Aminabadi et al., and Casas et al. the intervention was completed in one session, whether they were performed under general or local anesthesia.</td>
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</table>

<table>
<thead>
<tr>
<th>Table 3. List of included articles.</th>
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</thead>
<tbody>
<tr>
<td>Nguyen et al. 2017</td>
<td>Pubmed</td>
</tr>
<tr>
<td>Howley et al. 2012</td>
<td>Pubmed</td>
</tr>
<tr>
<td>Aminabadi et al. 2008</td>
<td>Pubmed</td>
</tr>
<tr>
<td>Casas et al. 2004</td>
<td>Pubmed</td>
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</tbody>
</table>
pulpectomy in this 23 months trial by Howley et al.\(^9\). The clinical failure rate was 13.1% for pulpotomy and 4.4% for pulpectomy at 2 years follow up for Aminabadi et al.\(^{20}\). Clinical failure was 22% in the pulpectomy group, with no clinical failures in the pulpotomy group in the trial by Casas et al.\(^{21}\).

**Radiographical failure.** Radiographical failure was reported to be 3% and 7% in the pulpotomy group at 12 and 18 months respectively, while for the pulpectomy group it was 8% at both 12 months and 18 months follow ups in the Nguyen et al. trial\(^9\). Cumulative radiographical results showed failure in 11% in the pulpotomy group, and 27% in pulpectomy group in the Howley et al. trial\(^9\). Radiographical failure was 23.9% in the pulpotomy group, and 8.6% in pulpectomy group at 2 years follow up as reported by Aminabadi et al.\(^{20}\). Radiographical failure was 41% in the pulpotomy group and 18% in pulpectomy group at 2 years follow up trial by Casas et al.\(^{21}\).

**Overall failure.** One incisor was lost early and counted as a failure in the pulpotomy group in the Howley et al. trial\(^9\). One tooth had to be extracted postoperatively (2.2%) in pulpotomy group in the Aminabadi et al. trial\(^{20}\).

**Pain.** One tooth with spontaneous pain (1%) was reported in pulpotomy group in Nguyen et al. trial\(^9\). No pain was reported in either group in Howley et al. trial\(^9\). Two teeth were reported as having spontaneous pain (4.4%) in the pulpotomy group, and one (2.2%) in the pulpectomy group in Aminabadi et al.\(^{20}\). No pain was reported in either group in the Casas et al. trial\(^{21}\).

**Soft tissue pathology.** Two teeth showed fistula (2%) in the pulpotomy group, and one tooth had soft tissue swelling (1%) in pulpectomy group in the Nguyen et al. trial\(^9\). No soft tissue pathology was reported in either group in Howley et al.\(^9\). The presence of fistula was reported in 3 teeth (6.6%) in the pulpotomy group in Aminabadi et al. trial\(^{20}\). The presence of gingival swelling or parulis in 9 teeth (22%) in pulpotomy group was reported by Casas et al.\(^{21}\).

**Pathological mobility.** Pathologic mobility was not reported for any tooth in three trials, Howley et al.\(^9\), Aminabadi et al.\(^{20}\) and Casas et al.\(^{21}\). Only one tooth with pathological mobility was reported by Nguyen et al.\(^{15}\).

**Pathological radiolucency.** The odds ratio for periapical radiolucency was 177.55; 95% CI 20.29 to 1554.01 (P<0.0001; chi-square test) in the Nguyen et al. trial\(^9\). After 23 months follow up, 7 teeth (23%) showed frank periapical radiolucency in the pulpotomy group, while only 1 tooth (3%) showed frank periapical radiolucency in the pulpectomy group in Howley et al. trial\(^9\). At 2 years follow up, 5 teeth (11.11%) showed periapical radiolucency in the pulpotomy group, while only one tooth (2.17%) in the pulpectomy group showed periapical radiolucency in the Aminabadi et al. trial\(^{20}\). At 2 years follow up, 7 teeth (58%) showed periapical radiolucency in the pulpotomy group, and 3 teeth (27%) in the pulpectomy group in Casas et al.\(^{21}\).

**Pathological root resorption.** The odds ratio for external root resorption was 136.41;95% CI 15.02 to 1238.27 (P<0.0001; chi-square test) in Nguyen et al. trial\(^9\). After 23 months follow up, 2 teeth (7%) showed large external root resorption in the pulpotomy group, and 4 teeth (14%) in the pulpectomy group, while for internal resorption, one tooth (3%) showed perforating internal root resorption in the pulpotomy group in the Howley et al. trial\(^9\).

At 2 years follow up, pathologic external or internal root resorption occurred in 6 teeth (13.3%) of the pulpotomy group and in 2 teeth (4.34%) of the pulpectomy group in the Aminabadi et al. trial\(^{20}\).

At 2 years follow up pathologic external root resorption occurred in 4 teeth (33%) in the pulpotomy group and in 3 teeth (27%) in the pulpectomy group, internal resorption was observed in 17% of the pulpotomy group in Casas et al. trial\(^{21}\).

**Pulp canal obliteration.** At 23 months, pulp canal obliteration was seen in 18 teeth (60%) in the pulpotomy group in Howley et al.\(^9\). At 2 years follow up, no teeth showed pulp canal obliteration in Aminabadi et al. trial\(^{20}\). At 2 years follow up, 3 teeth (25%) showed pulp canal obliteration in the pulpotomy group in the Casas et al. trial\(^{21}\).

**Tooth survival.** The survival rate was 0.94 (95% CI equals 0.89 to 0.97) for pulpotomy and 0.97 for pulpectomy at 18 months in Nguyen et al. trial\(^9\). The survival rate was 63% for pulpotomy and 85% for pulpectomy at 2 years follow up in Casas et al. trial\(^{21}\).

**Risk of bias in included studies.** The risk of bias of included studies is shown in Table 4, Table 5, Table 6, and Table 7. The overall risk of bias was low in two trials, Nguyen et al.\(^{15}\) and Aminabadi et al.\(^{20}\). The risk of bias was unclear for clinical assessment in two trials, Howley et al.\(^9\) and Casas et al.\(^{21}\), and also for random sequence generation in Casas et al.\(^{21}\). One trial, Casas et al.\(^{21}\), showed high risk of bias due to high percentage of dropped out cases. The risk of bias of included studies is also summarized in Figure 2.

**Synthesis of results.** Only three trials were included in the meta-analysis by Nguyen et al.\(^{15}\) Howley et al.\(^9\) and Aminabadi et al.\(^{20}\) as one trial by Casas et al.\(^{21}\) was excluded due to its high risk of bias. Data were extractable from all three RCTs totaling 338 teeth.

The data of the longest follow up period was included in the meta-analysis, this was 24 months for Aminabadi et al.\(^{20}\), 18 months for Nguyen et al.\(^{15}\), and at 15 to 23 months for Howley et al.\(^9\).

**Clinical failure**

At the longest follow up period the pooled results showed no statistically significant difference in clinical failure for pulpotomy compared to pulpectomy (RR 2.69, 95% CI 0.76 to 9.58) as shown in Figure 3.

The Howley et al. study was excluded from the meta-analysis of clinical failure as there were no events in both arms\(^{15}\).
Table 4. Risk of bias in Nguyen et al. 2017.

<table>
<thead>
<tr>
<th>Bias</th>
<th>Authors’ judgement</th>
<th>Support for judgement</th>
</tr>
</thead>
<tbody>
<tr>
<td>Random sequence generation (selection bias)</td>
<td>Low risk</td>
<td>Computer-generated simple random numbers sequence with a one to one allocation ratio.</td>
</tr>
<tr>
<td>Allocation concealment (selection bias)</td>
<td>Low risk</td>
<td>Quote: Allocation occurred after induction of general anesthesia to ensure allocation concealment. The pediatric dentist, nurse, or assistant directed subjects at the time of dental surgery to the appropriate treatment group they had been assigned to by the investigator.</td>
</tr>
<tr>
<td>Blinding of participants and personnel (performance bias)</td>
<td>Low risk</td>
<td>Quote: All other contributors were blinded to generation and implementation of the treatment assignment</td>
</tr>
<tr>
<td>Blinding of clinical outcomes assessment</td>
<td>Low risk</td>
<td>Quote: A single investigator, who did not perform any pulp therapy or participate in radiographic evaluation, performed all clinical assessments.</td>
</tr>
<tr>
<td>Blinding of radiological outcomes assessment</td>
<td>Low risk</td>
<td>Quote: Two experienced pediatric dentists who did not participate in protocol development or treatment performed radiographic assessments. It is not possible to blind the assessors due to the nature of treatment received.</td>
</tr>
<tr>
<td>Incomplete outcome data (attrition bias)</td>
<td>Low risk</td>
<td>9% drop out of 172 incisors at 12 months, due to loss to follow-up (n=13) and due to trauma (n=2). 21% drop out of 172 incisors at 18 months, due to loss to follow-up or dropout (n =31), exfoliation (n = 3), and trauma (n = 3).</td>
</tr>
<tr>
<td>Reporting bias</td>
<td>Low risk</td>
<td>We revised the protocol that was registered with the ClinicalTrials.gov Protocol and Registration System (ID no. NCT02019563)</td>
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</tbody>
</table>

Table 5. Risk of bias in Howley et al. 2012.

<table>
<thead>
<tr>
<th>Bias</th>
<th>Authors’ judgement</th>
<th>Support for judgement</th>
</tr>
</thead>
<tbody>
<tr>
<td>Random sequence generation (selection bias)</td>
<td>Low risk</td>
<td>An incisor in each pair was randomly assigned by a coin toss.</td>
</tr>
<tr>
<td>Allocation concealment (selection bias)</td>
<td>Low risk</td>
<td>Quote: An incisor in each pair was randomly assigned, by a coin toss to either the experimental group or the control group with the contralateral- paired incisor being designated to the other treatment group</td>
</tr>
<tr>
<td>Blinding of participants and personnel (performance bias)</td>
<td>Low risk</td>
<td>It is not possible to blind the operator and the participant blinding is ineffective on outcomes</td>
</tr>
<tr>
<td>Blinding of clinical outcomes assessment</td>
<td>unclear risk</td>
<td>Insufficient information to make a clear judgement</td>
</tr>
<tr>
<td>Blinding of radiological outcomes assessment</td>
<td>Low risk</td>
<td>Quote: The radiographs were evaluated independently by 2 standardized and calibrated examiners who were not otherwise involved in the study. It is not possible to blind the assessors due to the nature of treatment received.</td>
</tr>
<tr>
<td>Incomplete outcome data (attrition bias)</td>
<td>Low risk</td>
<td>29 study patients, 3 patients failed to return for follow-up.</td>
</tr>
</tbody>
</table>

Radiographic failure

The pooled results showed RR 0.79, 95% CI 0.46 to 1.21 in radiographical failure for pulpotomy compared to pulpectomy as shown in Figure 4 but considerable heterogeneity was presented in the meta-analysis of radiographic failure due to the presence of one outlying study of Aminabadi et al. so we performed the analysis both with and without this study as part of a sensitivity analysis as shown in Table 8.

<table>
<thead>
<tr>
<th>Bias</th>
<th>Authors’ judgement</th>
<th>Support for judgement</th>
</tr>
</thead>
<tbody>
<tr>
<td>Random sequence generation (selection bias)</td>
<td>Low risk</td>
<td>Coin tossing</td>
</tr>
<tr>
<td>Allocation concealment (selection bias)</td>
<td>Low risk</td>
<td>Quote: For each patient, by coin tossing, if one tooth was randomly assigned for formocresol pulpotomy then root canal therapy (RCT) was performed on the other incisor.</td>
</tr>
<tr>
<td>Blinding of participants and personnel (performance bias)</td>
<td>Low risk</td>
<td>It is not possible to blind the operator and the participant blinding is ineffective on outcomes</td>
</tr>
<tr>
<td>Blinding of clinical outcomes assessment</td>
<td>Low risk</td>
<td>Quote: Two clinicians who did not perform any treatments analyzed the clinical and radiographic outcomes</td>
</tr>
<tr>
<td>Blinding of radiological outcomes assessment</td>
<td>Low risk</td>
<td>Quote: Two clinicians who did not perform any treatments analyzed the clinical and radiographic outcomes. It is not possible to blind the Assessors due to the nature of treatment received.</td>
</tr>
<tr>
<td>Incomplete outcome data (attrition bias)</td>
<td>Low risk</td>
<td>4 subjects were dropped out of 50 patients.</td>
</tr>
</tbody>
</table>


<table>
<thead>
<tr>
<th>Bias</th>
<th>Authors’ judgement</th>
<th>Support for judgement</th>
</tr>
</thead>
<tbody>
<tr>
<td>Random sequence generation (selection bias)</td>
<td>Unclear risk</td>
<td>Insufficient information to make a clear judgement</td>
</tr>
<tr>
<td>Allocation concealment (selection bias)</td>
<td>Low risk</td>
<td>Quote: Quality assurance checks were performed by 1 of the investigators (MAL), who did not provide treatment or review postoperative radiographs, to ensure that the investigators who provided treatment complied with the randomization protocol</td>
</tr>
<tr>
<td>Blinding of participants and personnel (performance bias)</td>
<td>Low risk</td>
<td>It is not possible to blind the operator and the participant blinding is ineffective on outcomes</td>
</tr>
<tr>
<td>Blinding of clinical outcomes assessment</td>
<td>Unclear risk</td>
<td>Insufficient information to make a clear judgement.</td>
</tr>
<tr>
<td>Blinding of radiological outcomes assessment</td>
<td>Low risk</td>
<td>Quote: Two independent pediatric dentists who were not otherwise involved in the investigation evaluated the radiographs. It is not possible to blind the assessors due to the nature of treatment received.</td>
</tr>
<tr>
<td>Incomplete outcome data (attrition bias)</td>
<td>High risk</td>
<td>Quote: Of the enrolled participants, 64% returned for at least 1 evaluation. 36% drop out in the pulpotomy group and 48% drop out in the root canal treatment group at 2 years follow up</td>
</tr>
</tbody>
</table>

The sensitivity analysis for radiographic failure showed RR 0.45, 95% CI 0.25 to 0.83 with statistical significant difference between pulpotomy and pulpectomy and a higher risk for radiographic failure in pulpectomy. Pain The pooled results showed no statistically significant difference in pain for pulpotomy compared to pulpectomy (RR 2.06, 95% CI 0.31 to 13.8)).
Figure 2. Risk of bias summary of included studies.

<table>
<thead>
<tr>
<th>Study or Subgroup</th>
<th>pulpotomy Events</th>
<th>pulpotomy Total</th>
<th>pulpectomy Events</th>
<th>pulpectomy Total</th>
<th>Weight</th>
<th>Risk Ratio M-H, Fixed, 95% CI</th>
</tr>
</thead>
<tbody>
<tr>
<td>Aminabadi et al. 2008</td>
<td>6</td>
<td>46</td>
<td>2</td>
<td>46</td>
<td>63.2%</td>
<td>3.00 [0.64, 14.10]</td>
</tr>
<tr>
<td>Howley et al. 2012</td>
<td>0</td>
<td>37</td>
<td>0</td>
<td>37</td>
<td>Not estimable</td>
<td></td>
</tr>
<tr>
<td>Nguyen et al. 2017</td>
<td>3</td>
<td>100</td>
<td>1</td>
<td>72</td>
<td>36.8%</td>
<td>2.16 [0.23, 20.35]</td>
</tr>
</tbody>
</table>

Total (95% CI) 183 155 100.0% Risk Ratio M-H, Fixed, 95% CI 2.63 [0.76, 9.58]

Test for overall effect: Z = 1.53 (P = 0.13)

Heterogeneity: Chi² = 0.06, df = 1 (P = 0.81); I² = 0%

Figure 3. Meta-analysis for clinical failure.

<table>
<thead>
<tr>
<th>Study or Subgroup</th>
<th>pulpotomy Events</th>
<th>pulpotomy Total</th>
<th>pulpectomy Events</th>
<th>pulpectomy Total</th>
<th>Weight</th>
<th>Risk Ratio M-H, Random, 95% CI</th>
</tr>
</thead>
<tbody>
<tr>
<td>Aminabadi, Farahani et al. 2008</td>
<td>11</td>
<td>46</td>
<td>4</td>
<td>46</td>
<td>31.5%</td>
<td>2.75 [0.94, 8.01]</td>
</tr>
<tr>
<td>Howley, Seale et al. 2012</td>
<td>4</td>
<td>37</td>
<td>10</td>
<td>37</td>
<td>31.5%</td>
<td>0.40 [0.14, 1.16]</td>
</tr>
<tr>
<td>Nguyen, Judd et al. 2017</td>
<td>10</td>
<td>100</td>
<td>15</td>
<td>72</td>
<td>37.0%</td>
<td>0.48 [0.23, 1.01]</td>
</tr>
</tbody>
</table>

Total (95% CI) 183 155 100.0% Risk Ratio M-H, Random, 95% CI 0.79 [0.25, 2.42]

Test for overall effect: Z = 0.42 (P = 0.67)

Heterogeneity: Tau² = 0.75; Chi² = 8.39, df = 2 (P = 0.02); I² = 76%

Figure 4. Meta-analysis for radiographic failure.
Soft tissue pathology
The pooled results showed no statistically significant difference in soft tissue pathology for pulpotomy compared to pulpectomy (RR 3.11, 95% CI 0.54 to 17.7).

Pathological radiolucency
The results of two trials showed considerable statistical heterogeneity so no meta-analysis were performed for this outcome.

Pathological root resorption
The pooled results of two trials showed no statistically significant difference in pathologic resorption for pulpotomy compared to pulpectomy (RR 1.5, 95% CI 0.56 to 4.04).

Discussion
Pulp therapy is performed to preserve primary teeth and maintain its developmental, esthetic, and functional capabilities. Pulpotomy and root canal therapy have been both performed as techniques for the management of asymptomatic vital primary incisors with large carious lesions where removal of caries will lead to pulp exposure. However the preference of many pediatric dentists to perform pulpectomy in primary incisors was due to the fact that they were taught to do so in their pediatric dentistry residencies and not due to evidence from high quality research. The aim of this systematic review was to compare between pulpotomy and pulpectomy clinically and radiographically in the treatment of carious vital pulp exposure in primary incisors.

Upon performing our systematic search there were only four randomized controlled trials that have compared pulpotomy and pulpectomy outcomes in vital primary incisors. After exclusion of one trial due to its high risk of bias, only three trials were left to be included in the meta-analysis. The data of the longest follow up period was included as the follow ups were close to each other ranging from 15 months to 24 months and they best reveal the efficacy of the performed techniques.

The results were calculated with risk ratio (RR) effect measure and confidence intervals (CIs). The pooled results of the clinical failure outcome showed the relative risk RR was 2.69 with 95% CI from 0.76 to 9.58, the CI including the number 1 means that there was no statistical significant difference between pulpotomy and pulpectomy cases.

For radiographic failure the RR was 0.79 with 95% CI from 0.25 to 2.42 but considerable statistical heterogeneity was detected due to the presence of one outlying study of Aminabadi et al. with results that conflict the other studies so sensitivity analysis was performed that showed that RR was 0.45 with 95% CI 0.25 to 0.83 with higher risk of radiographic failure for pulpectomy rather than pulpotomy.

Although there is no clinical diversity among the included trials that may have led to this statistical heterogeneity, we may relate it to that the criteria of radiographic assessment in Aminabadi et al. study was not clearly specified.

The exclusion of a study from a meta-analysis based on their result may introduce bias so the results must be interpreted with an appropriate degree of caution and further investigation is required. For the clinical outcomes pain and soft tissue pathology, the pooled results for these outcomes showed no statistically significant difference while pathologic mobility was only reported for one incisor in one trial.

The radiographic outcomes included periapical radiolucency and pathologic root resorption. For pathologic resorption, the pooled results showed no statistically significant difference between pulpotomy and pulpectomy while pathologic mobility was only reported for one incisor in one trial.

Pediatric dentists do not consider the radiographic pathological changes as questionable radiolucencies or limited pathological root resorptions to be an absolute indication for extraction taking.
into consideration the absence of clinical pathological signs or symptoms\(^8\).

Overall failure was reported for two pulpotomized incisors in two trials. On the other hand tooth survival although it is an important outcome, but it is not commonly reported.

This study showed that there is no statistical significant difference in clinical success rates of pulpotomy and pulpectomy with different medicaments in the treatment of carious vital pulp exposure in primary incisors while radiographically, pulpectomy may have a higher risk for radiographic failure than pulpotomy and this refutes the misconception among some pediatric dentists that pulpotomy does not work in primary incisors.

The evidence was limited by the small number of trials included in the meta-analysis. The overall risk of bias of primary studies was low for three trials, except for the unclear risk for blinding of clinical assessment which was not effective. We did not have access to all the trial protocols to assess the selective reporting bias except for only one trial.

Conclusion

There was no statistical significant difference in the clinical success rates of pulpotomies and pulpectomies in the pulp treatment of carious vital pulp exposure in primary incisors while radiographically, pulpectomy may have a higher risk for radiographic failure than pulpotomy. We recommend teaching of pulpotomy as a treatment option for vital pulp exposure in primary incisors in pediatric dentistry residency programs and further high quality studies comparing between pulpotomy and pulpectomy in primary incisors with a longer follow up period till exfoliation time.

**Data availability**

Dataset 1: Summarized extracted data from included trials. 10.5256/f1000research.16142.d218857\(^7\)

**Grant information**

The author(s) declared that no grants were involved in supporting this work.

**Author information**

LG is a Researcher Assistant at the Orthodontics and Pediatric Dentistry Department, National Research Centre, Egypt. MH is a Professor at the Pediatric Dentistry and Dental Public Health Department, Faculty of Dentistry, Cairo University, Egypt. MAEY is a Professor at the Orthodontics and Pediatric Dentistry Department, National Research Centre, Egypt.

**Supplementary material**

Supplementary File 1: Completed PRISMA checklist.

Click here to access the data.

**References**


Marcela Baraúna Magno
Department of Pediatric Dentistry and Orthodontics, School of Dentistry, Federal University of Rio de Janeiro, Rio de Janeiro, Brazil

The authors performed most of the requested changes. Minor changes are necessary.

Abstract:
- Radiographic assessment needs to be added in abstract conclusion.

Results:
- According Cochrane, yellow represents unclear. This could be generated in Revman if you justified the unclear classification in software.

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: systematic reviews, pediatric dentistry, dental materials, dental trauma, carie

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.
Marcela Baraúna Magno  
Department of Pediatric Dentistry and Orthodontics, School of Dentistry, Federal University of Rio de Janeiro, Rio de Janeiro, Brazil

Abstract:  
- The methodology of bias risk analysis and meta-analysis was lacking.  
- If you have word limitation, you can reduce the abstract background.  
- Is risk ratio correct or relative risk?  
- I was confused. How radiographically did the pulpectomy prove worse?

Methodology:  
- Differentiate the primary outcome and the secondary outcome.  
- Correct me if I am wrong, but the factors considered in the secondary outcome are those that will determine the primary outcome.  
- The search is updated...  
- Search is confusing and seems to be wrong.  
- I did not understand why at # 22 you crossed # 18 (which would be terms related to 'prior vital teeth' and 'pulpotomy') or terms related to 'pulpectomy' (# 19 or # 20).  
- The # 21 was not used at any intersection.  
- I did not understand the purpose of # 24.  
- If you want studies evaluating pulpotomy X pulpectomy, you should cross terms related to 'anterior teeth' and 'pulpotomy' and 'pulpectomy'.  
- If you did not do this kind of crossover, some terms may have been selected wrong. Please review.  
- The authors considered homogeneous until the value of i2?

Results:  
- Include the unclear symbols in figure 2.  
- Remove the Howley study, since this is not considered in analysis in figure 3.  
- Include a descriptive result for pathological radiolucency. What the studies concluded about this outcome?

Conclusion:  
- Only for clinical parameters, no?  
- Pulpectomy presented more failures than pulpotomy for radiographic parameter.

Are the rationale for, and objectives of, the Systematic Review clearly stated?  
Yes

Are sufficient details of the methods and analysis provided to allow replication by others?  
Partly

Is the statistical analysis and its interpretation appropriate?
Yes

**Are the conclusions drawn adequately supported by the results presented in the review?**
Partly

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** systematic reviews, pediatric dentistry, dental materials, dental trauma, carie

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

---

**Author Response 20 Jun 2019**

**Lamia Gadallah**, National Research Centre, Egypt, Nasr city, Egypt

Dear Dr. Marcela,

We are honored by your reviewing to our article and we took into consideration all your valuable corrections and comments.

Firstly, in the abstract section:
The risk of bias tool and meta analysis method used was added.
The relative risk is called the risk ratio and both terms are mentioned in Cochrane Handbook for systematic reviews of interventions.
Radiographically the risk ratio value was 0.45 95%CI 0.25 to 0.83. These values are less than one which means higher risk of radiographic failure in the control(pulpectomy) group.

Secondly, in the methodology section:
The authors chose the primary outcomes according to a clinical point of view that there is a set of outcomes clinically and radiographically that collectively indicate the success or failure of the technique and you are absolutely right that those secondary outcomes are the single outcomes that composed the composite primary outcome, and we preferred to show their effect separately as secondary ones due to their less importance as single outcomes.
Yes the search was last updated in march 2018.

In the search strategy, we thank you for the two mistakes you have highlighted. By revising the saved history of search strategy, we found two editing mistakes, first line #22 is composed of #19,#20 #21 and not #18.
Line 24 is combined with AND not OR because we wanted the studies of incisors and pulpotomy with incisors and pulpectomy, that's how the actual search was performed, the mistakes were during editing tables.
The I2 is considered homogenous up till 50%.

Thirdly, the results section:
Figure 2 is according to Risk of bias summary figure in Cochrane Handbook where green represents low risk, red represents high risk and blank is unclear risk.
The Howley et al study was automatically excluded from the forest plot as there are zero events in both arms, we left it only in the table to show that zero events are not estimable.
In the periapical radiolucency outcome the two trials included were Howley et al. and Aminabadi et al, the study of Nguyen et al. was excluded due to lack of values in this outcome, only odds ratio was reported with no prevalence percentage in the control or non
exposure group so we could not convert the odds ratio to risk ratio.
Lastly in the conclusion section:
Radiographic assessment was added.

**Competing Interests:** No competing interests

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**Version 1**

Reviewer Report 03 December 2018

https://doi.org/10.5256/f1000research.17628.r41014

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Armin Shirvani

Iranian Center for Endodontic Research, Research Institute of Dental Sciences, Faculty of Medical Education, Shahid Beheshti University of Medical Sciences, Tehran, Iran

The manuscript is well structured and the format of the report on the details of the study is acceptable.

There are two points to consider in the Meta-analysis:
1. The results of Hawley's study should not have been removed because of zero in nominators. They could use 0.01 instead of 0 and continue the analysis.
2. Heterogeneity in time intervals is an important issue and if a significant heterogeneity has been detected the results of the meta-analysis would not be valid.

**Are the rationale for, and objectives of, the Systematic Review clearly stated?**
Yes

**Are sufficient details of the methods and analysis provided to allow replication by others?**
Yes

**Is the statistical analysis and its interpretation appropriate?**
No

**Are the conclusions drawn adequately supported by the results presented in the review?**
Yes

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** research methodology and statistical analysis
I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 09 Dec 2018

Lamia Gadallah, National Research Centre, Egypt, Nasr city, Egypt

Dear Dr. Armin,

Many thanks for reviewing our study and for your valuable comments. Concerning your first comment on the exclusion of Howley et al. study from the clinical failure meta-analysis, we returned to Cochrane handbook for systematic reviews of "intervention that stated that". The standard practice in meta-analysis of odds ratios and risk ratios is to exclude studies from meta-analysis where there are no events in both arms. This is because such studies do not provide any indication of either the direction or magnitude of the relative treatment effect"

Actually the Revman program that we used for meta-analysis does not accept any decimals in the events in risk ratio as dichotomous outcomes. As for the heterogeneity in the radiographic evaluation, we performed a sensitivity analysis and it was due to one outlying study so we performed the meta-analysis both with and without this study and we stated that the results should be interpreted with caution.

Competing Interests: No competing interests were disclosed.

Reviewer Report 07 November 2018

https://doi.org/10.5256/f1000research.17628.r39650

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Mariem O. Wassel

Department of Pediatric Dentistry and Dental Public Health, Faculty of Dentistry, Ain Shams University, Cairo, Egypt

The systematic review and meta-analysis regarding pulpotomy versus pulpectomy in the treatment of vital pulp exposure in primary incisors is well designed and that the methodology and discussion are well written. One question I have is: was the protocol of this review registered before carrying out the research? Some mistakes in writing the manuscript are present which are:

○ Please be sure the reference number is written right after the author name through-out the whole manuscript.

○ In the background section of the abstract: This study aimed to assess the (effectiveness) of pulpotomy and pulpectomy (pulpectomy) in treatment of carious vital pulp exposure in primary incisors.

○ In the conclusion section of the abstract; Further high quality studies comparing between
pulpotomy and pulpectomy in primary incisors with longer follow up period till exfoliation time (are needed).

Page 2, introduction section:
1. In pulpectomy, a resorbable material such as nonreinforced
2. According to a Cochrane systematic review there was no
3. Endoflas were found to be equally effective while with a low quality of evidence zinc oxide and eugenol may be better than
4. We therefore aimed to determine in patients with carious vital pulp exposure in primary incisors if pulpotomy is better than

Page 6:

Duration of follow up:
- “Follow up was at 12 and 24 months in two trials, by Aminabadi et al[ref-1] and Casas et al[2].”
- Follow up was (up) to 23 months at three intervals: 5–9, 10–14, and 15–23 months
- Anesthesia: under local anesthesia (by) Aminabadi et al[1].
- Medicaments -- Pulpotomy: please rewrite this paragraph noting that 2 trials performed formocresol pulpotomy as we don't achieve hemostasis with formocresol.
- Pulp access: In the trial by Howley et al[3] the pulp chamber was unroofed using
- Pulpotomy: in another (the other) two trials by Nguyen et al[ref-4] and Casas et al[ref-2].
- Final restoration : Resin restorations was (were) performed in three..... Full coverage crowns whether stainless steel crown (SCC) or SCC with white esthetic veneer (were used) in Howley et al(trial)3.
- Results of studies;
- nor pulpectomy in this (the) 23 month(s) trial by Howley et al[3]. The clinical failure rate was 13.1% for pulpotomy and 4.4% (for pulpectomy) at

Page 7:
- radiological failure: was in (remove "in") 41% in the pulpotomy group and 18% in pulpectomy group at 2 years follow up trial by Casas et al[ref-2].
- Pain: No pain was reported in either group(s) in Howley et al[3], trial9.
- No pain was reported in either group(s) in the Casas et al[2]. trial21.
- Soft tissue pathology: No soft tissue pathology was reported in either group(s) in Howley et al[3], (trial)9.
- Pathological mobility: Pathologic mobility was not reported for any tooth in three trials ()
- Pathological radiolucency: At 2 years follow up, 5 teeth (11.11%) showed
- follow up, 7 teeth (58%) showed periapical radiolucency in the
- pathological root resorption: while for internal resorption, one tooth (3%) showed perforating
- At 2 years follow up, pathologic external or internal root resorption occurred in 6 teeth (13.3%) (in) the pulpotomy group and in 2 teeth (4.34%) (in) the pulpectomy group in the
- Aminabadi et al. trial20
- At 2 years follow up, pathologic external root resorption occurred

Pathological radiolucency:

Pulp canal obliteration:
At 23 months, pulp canal obliteration was seen in 18 teeth (60%) in the pulpotomy group in Howley et al.'s (trial). At 2 year follow up, no teeth showed pulp canal obliteration.

Please add reference number to authors names in the titles of tables 4 to 7.

Page 11, discussion section:
- 2nd paragraph: After exclusion of one trial due to its high risk of bias, only three trials were left to be included in the meta-analysis.
- Overall failure was reported for two pulpotomized incisors in two trials. (on the other hand,) tooth survival, although it is an important outcome, but it is not commonly reported.
- The overall risk of bias of primary studies was low for three trials, except for the unclear risk for blinding of clinical assessment which was not effective. We did not have access to all the trial protocols to assess the selective reporting bias except (for) only one trial.

References

Are the rationale for, and objectives of, the Systematic Review clearly stated?
Yes

Are sufficient details of the methods and analysis provided to allow replication by others?
Yes

Is the statistical analysis and its interpretation appropriate?
Yes

Are the conclusions drawn adequately supported by the results presented in the review?
Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: pediatric dentistry

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.
Thank you Dr. Mariem very much for reviewing our study. As for the protocol registration, yes it was registered in the Evidence based Committee of Cairo University as the preliminary step of my Ph.D. in 2014.

**Competing Interests:** No competing interests were disclosed.