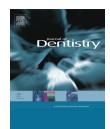
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Review

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Evaluation of the efficacy of potassium nitrate and sodium fluoride as desensitizing agents during tooth bleaching treatment—A systematic review and meta-analysis

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ABSTRACT

Objectives: This meta-analysis was performed to evaluate the efficacy of potassium nitrate and sodium fluoride as desensitizing agents during tooth bleaching treatment.

Data, sources and study selection: An electronic systematic literature search was conducted in Cochrane Center Register of Controlled Trials, MEDLINE (PubMed) and EmBase in April, 2014 in English and without time restrictions. Study information extraction and methodological quality assessments were accomplished by two reviewers independently. Methodological quality was assessed by using the "Criteria for judging risk of bias in the 'Risk of bias' assessment tool". Dichotomous data was summarized by odds ratio (OR) with 95% confidence interval (CI) and continuous data was summarized by mean difference (MD) or standardised mean difference (SMD) with 95% confidence interval (CI). Statistical analyses were carried out by using Review Manager 5.2.

For evaluation of percent of patients experiencing tooth sensitivity (POTS), the pooled OR of desensitizers vs. placebo was 0.45 (95% CI: 0.28–0.73, P = 0.29). The pooled SMD of desensitizers vs. placebo was -0.47 (95% CI: -0.77 to -0.18, P = 0.13) in evaluation of level of tooth sensitivity (LOTS). The results of shade evaluation remained inconsistent by evaluating subjective shade guide unit difference (Δ SGU or SGU) and objective colour difference (Δ E).

Conclusions: This meta-analysis was performed to evaluate the efficacy of desensitizing agents, potassium nitrate and sodium fluoride, for tooth bleaching treatments. Potassium Q2 nitrate and sodium fluoride reduce tooth sensitivity while no consistent conclusion of tooth colour change was found.

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$\frac{19}{1}$ **1.** Introduction

18 Q3 Following improvement in the economy and in people's living 19 standards, an increasing number of people became self-aware 20 that teeth play an important role in their appearance. These patients with discoloured teeth demand whiter teeth and a 21 22 more aesthetically pleasing smile. According to a previous 23 review, teeth whitening may be improved by physical or 24 chemical approaches.¹ When comparing whitening tooth-25 pastes, professional cleaning, microabrasion of enamel with 26 abrasives and acid, resin-bonded composites, porcelain 27 veneers and crowns, tooth bleaching was considered a more 28 accepted and conservative approach to improving the colour of teeth and a pleasant smile.¹⁻⁶ The main chemical 29 approaches of tooth bleaching include at-home treatment, 30 in-office procedure and bleaching with over-the-counter (OTC) 31 bleaching products.^{7,8} For at-home tooth bleaching treatment, 32 33 10% carbamide peroxide (CP) delivered in a custom-fitting 34 mouth tray was introduced by Klusmier in late 1960s.⁹ This 35 successful technique was published by Haywood and Hey-36 mann in 1989 and has become the gold-standard treatment in tooth whitening.⁹⁻¹² Later the in-office procedure, of applying 37 a high concentration of hydrogen peroxide (HP) to tooth 38 39 bleaching with (or without) exposure to various kinds of heat or curing lights to enhance the effects became popular.^{8,11–15} 40 Typically a 30–35% concentration of HP is used in vital and 41 42 nonvital tooth bleaching during in-office procedures.¹¹ Unlike at-home bleaching techniques, in-office bleaching is super-43 44 vised by dentists. OTC products have increased in popularity 45 with patients and are self-administered. Different OTC products have become available in markets, including 46 47 whitening strips and gels, whitening rinses, paint-on gels with brushes, toothpastes, etc.^{1,8,16} The safety and efficacy of 48 OTC products remain questionable.^{1,8,16,17} Evidence has 49 50 shown that bleaching products based on CP and/or HP are 51 relatively safe and effective when following manufacturer's instructions.1,7,18 52

53 Nevertheless adverse effects are another concern often 54 expressed with bleaching.^{19–21} Many publications have voiced 55 concerns about oral health and potential tooth structure change after bleaching. Difficulty in dental hygiene, an 56 57 unpleasant sensation in the mouth, gingival irritation, during and postoperative tooth sensitivity, structural integrity of 58 59 dental hard tissue and restoration are common risks reported with tooth bleaching.^{20,22–24} Tooth sensitivity and/or gingival 60 irritation may be typical side effects associated with tooth 61 bleaching techniques.²⁵ In previous reports, up to 66% of 62 patients experienced side effects (tooth sensitivity and/or 63 gingival irritation) after overnight vital bleaching.²⁶ This kind 64 of discomfort is sometimes responsible for patients' hesitancy 65 in tooth bleaching. 66

Modifications and improvements have been made to 67 68 bleaching products. As compared with decreasing the con-69 centration of peroxide products and administration of analgesic, the application of a desensitizing agent seemed 70 71 to be an effective option to reduce tooth sensitivity.^{27–29} 72 Potassium nitrate and sodium fluoride as desensitizers are used widely to treat tooth sensitivity. These agents may be 73 74 contained in bleaching gel and delivered by using a custom

tray during treatment. Other delivery systems can be used independently by placing them into a subject's mouth for a short time before bleaching is introduced.^{4,29-32} The mechanism of action of potassium nitrate remains unknown. Several randomized clinical trials have been published assessing the safety and efficacy of tooth bleaching treatments. Some of them conclude that desensitizers based on potassium nitrate and sodium fluoride reduce tooth sensitivity.^{29,31-34} Others question the efficacy of tooth bleaching when a desensitizing agent is used.^{4,30} The results of these studies are sometimes even conflicting, possibly because of the small number of patients. Therefore the aim of this meta-analysis is to provide a more accurate estimate of the efficacy of two desensitizing agents, potassium nitrate and sodium fluoride, during bleaching treatment. 75

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2. Methods

All the work was performed by two reviewers, independently. A discussion ensued if any disagreement occurred and unresolved issues were solved by consulting a third reviewer.

2.1. Inclusion criteria

Clinical studies were included in this meta-analysis that satisfied the following inclusion criteria. Participants: patients with discoloured teeth (intrinsic discolouration and/or extrinsic discolouration) undergoing tooth bleaching treatment had to be randomized to test or control groups. The main approaches of tooth bleaching had to include at-home treatments and/or in-office procedures. Interventions: desensitizer agents had to be based on potassium nitrate and/or sodium fluoride intervention products. Comparisons: control products had to be placebo or other desensitizing agents. Outcomes: outcomes had to be arranged into two categories: tooth sensitivity evaluation and shade evaluation. Studies design: studies had to be designed as randomized controlled trials (RCT) or controlled clinical trials (CCT).

2.2. Exclusion criteria

Clinical studies were excluded in this meta-analysis that contained the following criteria: (1) those estimating the efficacy of over-the-counter (OTC) products which contained desensitizing components, such as dentifrice, whitening strips; (2) duplicate studies; (3) irrelevant studies; (4) in vitro studies.

2.3. Search strategy and study selection

An electronic systematic literature search was conducted in 117 Cochrane Center Register of Controlled Trials, MEDLINE (PubMed) 118 and EmBase in April, 2014 in English and without time 119 restrictions. Additional studies were identified by searching 120 reference lists of included studies and contacting experts. The 121 medical subject headings (MeSH) and text words "hydrogen 122 peroxide", "carbamide peroxide", "whitening", "tooth whiten-123 ing", "bleaching", "tooth bleaching", "brightening", "aesthetics", 124

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