Assessment of oral changes resulting from the use of electronic cigarettes: Literature review
Avaliação das alterações bucais decorrentes do uso dos cigarros eletrônicos: Revisão da literatura
Evaluación de cambios orales resultantes del uso de cigarrillos electrónicos: Revisión de la literatura

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Abstract
The aim of the present study is to carry out a bibliographic review regarding oral changes resulting from the use of electronic cigarettes. The PubMed database was used, with the following terms and operators: “electronic cigarette” OR “e-cigarette” OR “electronic cigarettes” AND “oral health” from January 2015 to January 2020. Books were also used and epidemiological data. The inclusion criteria were articles published in Portuguese and / or English, which met the theme proposed by the study. Non-systematic literature reviews and articles that did not strictly refer to the topic were excluded. The final result was a sample of 14 articles, in which 7.14% (n = 1) was classified as a systematic review, 78.57% (n = 11) as an “in vivo” study; 7.14% (n = 1) as a case control study and 7.14% (n = 1) as a pilot study. There was a low level of published evidence on the results of oral health. The effects mentioned referred to periodontal changes and cellular damage to oral tissues. In addition, the use of electronic cigarettes has not demonstrated health security in its entirety.
Keywords: Electronic Nicotine delivery systems; Oral health; Mouth abnormalities.

Resumo
O objetivo do presente estudo é realizar uma revisão bibliográfica a respeito das alterações orais decorrentes do uso dos cigarros eletrônicos. Foi utilizada à base de dados PubMed, com os seguintes termos e operadores: “electronic cigarette” OR “e-cigarette” OR “electronic cigarettes” AND “oral health” no período de janeiro de 2015 a janeiro de 2020. Livros foram também utilizados e dados epidemiológicos. Os critérios de inclusão foram os artigos publicados
1. Introduction

Tobacco consumption in the last 12 years has been reduced, results presented by the Surveillance System for Risk and Protection Factors for Chronic Diseases by Telephone Survey (Vigitel) showed that the reduction in consumption was 40% and that the national trend observed is that this harmful habit is reduced even more year after year (Dar-Odeh, 2010).

Despite the various implementations of public policies for smoking control, electronic cigarettes are becoming more popular and have been spreading under the misconception that the new form of consumption may be less harmful when compared to conventional cigarettes (Dar-Odeh, 2010). In electronic cigarettes, nicotine is released in the form of aerosol / liquid vapors, along with several other substances, including propylene glycol, vegetable glycerin, with a multitude of aromas and flavors. The idea is that nicotine and other chemicals present in the tobacco leaf undergo a process of removing impurities, theoretically resulting in a cleaner form for consumption (Farsalinos & Stimson, 2014).

Electronic cigarettes are marketed both for recreational use and for smokers who want to minimize or stop smoking. However, the current scenario concerns health entities, considering that the use of electronic cigarettes has increased among young adults, among people who have never smoked and ex-smokers (Sutfin, 2013). The harmful effects of conventional cigarettes and the problems caused by impaired innate immunity are known and frequently appear as the object of research. For the oral cavity, inflammatory gingival and periodontal changes are known, as well as predisposing factors for oral cancer (Huilgol, 2018).

On the other hand, for electronic cigarettes, few studies have been reported on the effects of electronic cigarettes on the oral cavity. Therefore, given the lack of proven scientific data, the commercialization, import and advertising of all types of smoking devices are currently prohibited in Brazil. Studies that assessed the content of emissions, that measure health impacts and that seek to describe the risks associated with these products still have inconsistencies regarding the use and risks attributed to these devices (Sutfin, 2013).

In countries where marketing is permitted, such as the USA, electronic cigarettes are already considered a public health problem (Sutfin, 2013). Therefore, the present study aims to carry out a bibliographic review regarding oral changes resulting from the use of electronic cigarettes, as well as to discuss the role of related aspects for the correct diagnosis of these changes and treatment mechanisms involved, expanding the understanding on the subject.
2. Methodology

To carry out this literature review, the PubMed database was used, using the following terms and operators: “electronic cigarette” OR "e-cigarette" OR "electronic cigarettes" AND “oral health”; and the publication date filter: articles published between January 2015 and January 2020. Books and epidemiological data were also used to construct this work.

Free articles, in Portuguese and / or English, and those in its full version, which apply to the theme proposed by the research, were included in the research in order to respond to the problems and objectives of the research. As exclusion criteria, non-systematic literature reviews and articles that did not strictly refer to the topic were excluded.

After the initial screening, the articles were downloaded and sequentially numbered and read in full, to verify the adequacy to the theme. Then, the information extracted from the selected articles were defined; categorization (case reports, clinical trial, case-control studies, pilot study, systematic reviews and meta-analyses); interpretation and analysis of results; and the presentation of the results obtained.

At the end, after the application of the election criteria for the selection of manuscripts for the development of the study, articles that did not meet the pre-established criteria were excluded and the selected articles of greater relevance on the topic were read in their entirety for obtaining information relevant to the object of this study.

3. Results

A total of 33 articles were evaluated. Of these, 57, 57% (n = 19) were excluded: 52, 63% (n = 10) for not referring to the proposed theme and 47, 37% (n = 9) for being classified as a literature review. The final result was a sample of 14 articles, in which 7,14% (n = 1) was classified as a systematic review, 78,57% (n = 11) as an “in vivo” study; 7,14% (n = 1) as a case control study and 12.5% (n = 1) as a pilot study.

Table 1. Classification of articles regarding the objectives and main results found.

<table>
<thead>
<tr>
<th>Author (year)</th>
<th>Country</th>
<th>Purpose</th>
<th>Results</th>
</tr>
</thead>
<tbody>
<tr>
<td>Tatullo (2016)</td>
<td>Italy</td>
<td>To assess variations in periodontal health and general health in a population of smokers who used electronic cigarettes for a period of 120 days.</td>
<td>A progressive improvement was observed both in the periodontal indexes and in the perception of the general health status. In addition, patients reported a reduced need for smoking.</td>
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<td>Pilot study</td>
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<td>Sundar (2016)</td>
<td>U.S</td>
<td>To determine mechanisms of gingival epithelial inflammation and premature senescence caused by aerosols of electronic cigarettes in epithelial cells and fibroblasts of the periodontal ligament.</td>
<td>They highlighted the pathological role of FB (with or without nicotine) for cells and tissues in the oral cavity. The results showed that CE caused an increase in oxidative / carbonyl stress and inflammatory processes with cytokine release in periodontal ligament fibroblasts, progenitors of the human gingival epithelium (HGEpp) and EpiGingival 3D epithelium. They found an association between the levels of prostaglandin-E2 and cyclooxygenase-2, and the positive regulation of the receptor for end products of advanced glycation (RAGE) by carbonyl stress in the epithelium / gingival tissue. DNA damage was also observed along with a reduction in the histone deacetylase 2 (HDAC2) enzyme through RAGE-dependent mechanisms in the gingival epithelium.</td>
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<tr>
<td>In-vivo study</td>
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<tr>
<td>Author(s)</td>
<td>Year</td>
<td>Study Type</td>
<td>Country</td>
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<tr>
<td>Cho (2017)</td>
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<td>In vivo</td>
<td>South Korea</td>
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<tr>
<td>Huigol (2018)</td>
<td></td>
<td>In vivo</td>
<td>U.S</td>
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<tr>
<td>Alharthi (2018)</td>
<td></td>
<td>In vivo</td>
<td>Saudi Arabia</td>
</tr>
<tr>
<td>Akinkugbe (2018)</td>
<td></td>
<td>In vivo</td>
<td>U.S</td>
</tr>
<tr>
<td>Alqahtani (2019)</td>
<td></td>
<td>In vivo</td>
<td>Saudi Arabia</td>
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</table>
| Binshabaib (2019)|       | In vivo    | Saudi Arabia | The aim of this study was to compare the periodontal clinical status and cytokine profile of gingival fluid (CFG) between conventional cigarette smokers (G1), electronic cigarette users (G2) and non-smokers (G3). | The mean scores of plaque index, probing depth and level of clinical insertion were significantly higher among individuals in G1 than in G3. Probing bleeding was most frequently manifested among patients in G3. Compared to G3, marginal bone loss was significantly greater in groups G1 and G2. The fluid volume of the marginal cracks was significantly higher in G1 compared to groups 2 and 3. The
Concentrations of IL-1β, IL-6, IFN-γ, TNF-α and MMP-8 were significantly higher in the samples of marginal cleft fluids in G1 (P <0.05) than in G2 and 3.

<table>
<thead>
<tr>
<th>Researcher (2019)</th>
<th>Geographical Location</th>
<th>Study Type</th>
<th>Details</th>
</tr>
</thead>
<tbody>
<tr>
<td>Holliday (2019)</td>
<td>UK</td>
<td>Systematic review</td>
<td>A systematic review was carried out to evaluate the in vitro effects of nicotine on human gingiva, periodontal ligament and oral epithelial cells, specifically: cell viability, cell fixation, cell proliferation and production of inflammatory mediators, in tobacco users. The nicotine levels found in tobacco smokers, nicotine users on replacement therapy and users of electronic cigarettes, are unlikely to be cytotoxic to human periodontium cells, although saliva levels in smokeless tobacco users may be high or enough to achieve cytotoxicity.</td>
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<tr>
<td>Atuegwu (2019)</td>
<td>U.S</td>
<td>Case-control study</td>
<td>The aim of this study was to evaluate the association between the use of nicotine in electronic devices and periodontal disease. The results showed that users of nicotine and electronic devices increased the chances of being diagnosed with gum disease (OR 1.76, 1.12–2.76) and bone loss around the teeth (OR 1.67, 1.06–2.63). These odds were higher among participants with a history of marijuana and a history of illicit drug use.</td>
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<tr>
<td>Cichonska (2019)</td>
<td>Poland</td>
<td>In vivo study</td>
<td>The aim of this study was to estimate changes in the physicochemical properties in the saliva collected from users of electronic cigarettes. Among electronic cigarette users, statistically significant differences were observed in the values of lysozyme and lactoferrin; however, no statistically significant differences were found for the IgA value. In the group of traditional cigarette smokers, statistically significant differences were observed between all parameters tested in relation to the group of non-smokers. In relation to IgA, statistically significant differences were found between users of electronic cigarettes and smokers of traditional cigarettes, with a disadvantage for the latter.</td>
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<td>Ji (2019)</td>
<td>U.S</td>
<td>In vivo study</td>
<td>The aim of this study was to assess whether aerosol exposure from electronic cigarettes impacted the genetic pathways of normal human oral keratinocytes, in particular the response path to unfolded protein. The results indicated that aerosols in electronic cigarettes positively regulate the genes of the response pathway to unfolded proteins in normal human oral keratinocytes.</td>
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<tr>
<td>Jeong (2019)</td>
<td>South Korea</td>
<td>In vivo study</td>
<td>The aim of this study was to examine the association between tobacco use through conventional cigarettes and electronic cigarettes with periodontal disease. Periodontal disease was more prevalent among users of electronic cigarettes (OR 2.34, 1.52-3.59) than users of conventional cigarettes (OR = 2.17, 1.76-2.68). In addition, electronic cigarette users had a significant relationship with tooth decay, toothache and dental damage.</td>
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<tr>
<td>Vora; Benjamin</td>
<td>U.S</td>
<td>In vivo study</td>
<td>This study aimed to evaluate self-reported gum disease among users of electronic cigarettes and users of other tobacco products. The groups with the highest relative chances for diagnosing periodontal disease were pipe users (OR 3.1, 1.5-6.4), users of electronic cigarettes (OR 2.6, 1.6-4.3), users of several tobacco products (OR 2.8, 2.3-3.5) and ex-smokers (&lt;12 months) (OR 2.8, 2.1-3.7).</td>
</tr>
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Source: Authors.
4. Discussion

Few studies are published on the potential health consequences of electronic cigarettes and on its effectiveness as a strategic method in helping smoking cessation. Studies carried out with samples of the chemical vapors and liquid substances used in these devices, warned about the harmfulness of toxic and carcinogenic substances, similar to those found in conventional cigarette smoke (Besaratinia, 2017). The concentrations of most carcinogenic compounds present in the aerosols of electronic cigarettes are lower than those present in conventional cigarette smoke, however there is no “safe” level of exposure given the potential of carcinogens (Vora, 2019).

The literature shows that reports of electronic systems associated with nicotine started in 1963, developed by tobacco companies. Research carried out at the University of California, San Francisco, found that in 1990 Philip Morris began to develop a device that aerolized a nicotine solution, using technology for pharmaceutical applications, as it was not known how such products would be designed by Food and Drug Administration. The widespread intention was to solve health problems and reduce the number of smokers, in a kind of nicotinic therapy by changing the form of consumption (Farsalinos & Stimson, 2014).

Research found that some models of electronic cigarettes marketed “without nicotine”, contained low levels of nicotine. The results also showed that half of the samples contained four tobacco-specific nitrosamines (NNN, NNK, NAB and NAT), the first two being classified as carcinogenic to humans (Sutfin, 2013).

The acute use of electronic cigarettes can trigger oxidative stress and increase resistance to airflow, and cause changes in the secretion of proteins from the immune system in human airways. Pulmonary repercussions are important factors in observing the consequences caused by smoking (Besaratinia, 2017).

Ganapathy (2017), examined the effects of aerosol exposure from electronic cigarettes on possible DNA damage in oral and lung epithelial cells. The results presented showed that exposure to aerosols from electronic cigarettes can cause significant levels of mutagenic DNA damage. Observation regarding the mechanisms that modulate DNA damage, identified an increase in reactive oxygen species, a decrease in total antioxidant capacity and a decrease in the expression of proteins essential for the repair of DNA damage; potentially increasing DNA damage and the risk of cancer. Yang, Sandeep & Rodriguez (2020) in a systematic review gathered evidence from available research on the impact of electronic cigarette use on oral health. Reports of exposure to electronic cigarettes were associated with an increased risk of deteriorating periodontal, dental and gum health, as well as changes in the oral microbiota. It has also been reported that vapors generated by electronic cigarettes have cytotoxic, genotoxic and carcinogenic properties.

5. Conclusion

The study presented a current review of the implications that electronic cigarettes can have in the oral cavity. There was a low level of published evidence on the results of oral health. The effects cited referred to periodontal changes and damage at the cellular level of oral tissues. It is worth mentioning that the use of electronic cigarettes is not completely safe for health.

References


