World Cancer Research Journal wcrj 2021; 8: e2039

# **CORRELATION BETWEEN CHRONIC** MUCOSAL TRAUMA AND ORAL CANCER. A CASE REPORT AND REVIEW OF THE LITERATURE

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**Abstract – Objective:** Squamous cell carcinoma is the most common malignancy in the oral cavity representing the 90% of all oral malignancies that affect oral cavity. Primary risk factors associated with oral cancer are alcohol and tobacco consumption. However, other emerging risk factors have been proposed in Literature, including chronic trauma of the oral mucosa related to dental prostheses and incongruous restorations. Therefore, the aim of this study was to review the literature on the correlation between oral carcinoma and chronic irritative trauma, with the help of a case report and with focus on clinical features and differential diagnosis.

Materials and Methods: A literature search on MEDLINE, PubMed, ScienceDirect, Cochrane Database of Systematic Reviews and Wiley InterScience was performed, using the following words: Oral Cancer, chronic Trauma and Denture Trauma in various combinations.

Results: The review produced 1356 results, but only 8 articles met the inclusion criteria and were involved in the review. No association has been proven between chronic mucosal trauma and oral cancer. Carcinoma has been shown to develop more frequently in trauma regions, particularly in the tongue. As for the data on the gender and age of affected patients, it emerges that most of the greatest incidence occurs around the sixth and seventh decade of life and men are more affected than women.

**Conclusions:** Due to the heterogeneity of the different studies and the strong presence of other factors such as smoking and alcohol, the role of trauma in carcinogenesis is unclear. Further molecular, epidemiological and case-control studies are needed to define this correlation. It is advisable to carry out more restricted checks in patients at risk with dental prostheses.

**KEYWORDS:** Case report, Chronic mucosal trauma, Denture trauma, Literature review, Risk factors, Squamous cell carcinoma.

## INTRODUCTION

Squamous Cell Carcinoma (SCC) is the most common malignancy in the oral cavity representing the 90% of all oral cancers that affect oral cavity<sup>1</sup>. In one of the most recent epidemiological study, SCC affects more than 300.000 people worldwide causing more than 150.000 deaths<sup>2</sup>. Main risk factors linked with the development of SCC are tobacco smoking (smokeless tobacco and some tobacco-specific N-nitrosamines WHO) and alcohol abuse (a review of the relationship between alcohol and oral cancer, 2011). Other factors related are human papillomavirus infections (mainly genotypes 16 and 18)3,4, ultraviolet radiation exposure5, nutritional factors<sup>6</sup> and chronic trauma<sup>7</sup>. Particularly, although

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not clear, some studies described a possible relationship between chronic irritative trauma and SCC development<sup>8-11</sup>. Chronic mechanical irritation caused by dental factors, incongruous prostheses or parafunctional habits can cause the onset of various lesions <sup>12</sup>. Basically, chronic trauma can:

- stimulate hyperproliferation of the epithelium (frictional hyperkeratosis<sup>13</sup>, *morsicatio bucca-rum*<sup>14</sup>);
- cause hyperplastic tissue growth (fibrous prosthesis hyperplasia, traumatic fibroma<sup>15,16</sup>);
- cause damage to the mucosa (chronic traumatic ulcer<sup>9</sup>).

Oral traumatic lesions are very common in the population<sup>12</sup>; for example traumatic fibroma, according to the epidemiological study by Santana *et al*<sup>17</sup> represent the 15% of diagnosed lesions. *Morsicatio buccarum* prevalence is 750 out of every million individuals<sup>18</sup>, with a greater predisposition of women, especially during pregnancy<sup>19</sup>. Brantes *et al*<sup>20</sup> in a recent cross sectional study show that the lesions most frequently found in patients with prostheses are: denture stomatitis (63%), inflammatory fibrous hyperplasia (19%) and traumatic ulceration (11%). In detail, chronic traumatic ulcer (i.e. lasting more

than two weeks) is relatively common<sup>21</sup>. In fact, in the literature, some SCC cases arising on chronic ulcers are described<sup>22,23</sup>. In addition, a recent literature study has proposed chronic traumatic ulcer on the lingual edge as a potentially malignant disorder<sup>24</sup>. Therefore, the aim of this work is to report a case of oral SCC arising from a previous chronic traumatic lesion and to carry out a review on the correlation between chronic irritative trauma and SCC, with a focus on clinical features.

#### **MATERIALS AND METHODS**

A review of the literature was made using three databases, PubMed, Web of Science and Scopus. The keywords used were: Oral Cancer AND chronic Trauma; Oral Cancer AND Denture Trauma; Oral Cancer AND Dental Trauma. We included article in English and studies in which the development of SCC is linked to oral trauma. Studies in animals, studies not in English, case reports and reviews were excluded. The review produced 1356 results, but only 8 articles met the inclusion criteria and were involved in the review (Figure 1).

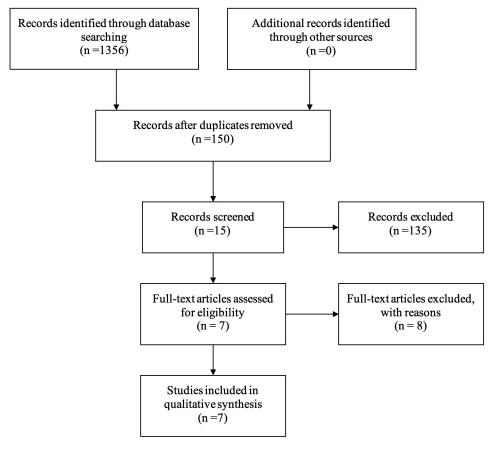


Fig. 1. The review processes.



**Fig. 2.** The lesion as appeared after the incisional biopsy.

#### **CASE REPORT**

A 54-years old woman came to visit for a suspected lesion in the left cheek. The patient had a history of periodontitis, slipped disc and she is a tobacco smoker (10 cigarettes/day). Her father was treated for a laryngeal carcinoma. The lesion present in her cheek was 3.5 x 1.5 cm, appeared erythematous with an irregular surface in correspondence of an area with sign of *morsication buccarum*. First, the lesion was treated by removing possible causal factor, as sharp tooth but the lesion did not resolve. Then, the lesion was biopsied (Figure 2), and a diagnosis of SCC was made.

#### Localization

In literature, tumours appear frequently in traumatized regions like tongue, cheek mucosa and less frequently in palate, gingiva, retromolar region. In the study of Piemonte et al<sup>8</sup>, in the 51% of the cases the tongue was the most involved location, followed by gums and alveolar crest (15%), buccal mucosa (13%), oral floor (9.5%), hard and soft palate (19%) and labial mucosa (2%). Another study, by comparing smoker patient and non-smoker patient, stated that lateral board of the tongue is the most involved site (66%) in the latter group, while for the former, the involvement is lesser (35%)<sup>25</sup>. Rotundo et al<sup>26</sup> showed that (22.5%) had the development of the tumour in the lateral edge of the tongue, (35%) in the oral floor, (22.5%) in the hard palate, (9%) in gingiva, (8.4%) in retromolar trigon and (1.4%) in cheek mucosa. About tongue localization of the tumour, another study affirmed that lateral edge of tongue was involved in the 77.5% followed by tongue venter 9.5%, dorsum 5.7% and the tip, 1.9%<sup>27</sup>.

## Age and Gender

As for the data on the gender and age of affected patients, the data vary in the articles but for the most, the greatest incidence occurs around the sixth and seventh decade of life and men are most affected than women<sup>8,25,27,28</sup>. Conversely, in the study of Perry et al<sup>25</sup>, incidence of oral cancer was higher for men than women in smoker and drinker group (65% *vs.* 35%); however, incidence was the opposite in non-smoker and non-drinker group (39% *vs.* 61%). Perry et al<sup>25</sup> stated that women who are neither smokers nor drinkers have a 4 times higher risk of developing oral cancer<sup>25</sup>. The data about gender and sex are collected in Table 1.

# Causal factors

The studies involved in the review process presented heterogeneity for number of patients, data, and causal factors studied, so controversies still exist for the role of trauma in tumour formation.

Particularly, a higher consumption of cigarettes per day increases the risk of developing oral cancer. The Odds Ratio (OR) for a consumption of 6-20 cigarettes / day is 3.1 and 7.96 for more than 20 cigarettes/day<sup>29</sup>.



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**TABLE 1.** Data regarding age and gender of the patients affected by oral carcinoma.

| Authors                           | Gender (M/F, %) | Average age (%) |
|-----------------------------------|-----------------|-----------------|
| Albuquerque et al <sup>27</sup>   | 71% M/ 29%F     | 63.7 yrs        |
| Bektas-Kayhan et al <sup>28</sup> | 64% M/36% F     | 52-53 yrs       |
| Perry et al <sup>25</sup>         | 72% M/28% F     | 61.7 yrs        |
| Piemonte et al <sup>8</sup>       | 54% M/46% F     | 63 yrs          |

Furthermore, another study found that lower cigarette consumption per day revealed a direct effect, whereby the strength of the pack-year association increased with the increase in cigarettes/day, indicating that for the same packs per day 'year, smoking more cigarettes a day for a shorter duration was more deleterious than fewer cigarettes a day for a longer duration<sup>30</sup>.

In the study of Albuquerque et al<sup>27</sup>, 44% of men drink and smoke and do not use a prosthesis vs. 51% of women. In this case, the women who have a tumour on the tongue and do not drink and smoke are 77%. Of this 77%, one third of patients wear prosthesis but not significantly association was found although these data show a higher incidence in women wearing prosthesis. Another study examined the relation between the use of a prosthesis and the formation of an oral tumour. One-hundred-thirty-four patients out of 211 patients affected by SCC wear different prostheses, i.e. crowns, bridges and dentures and of these 34% wear dentures, 22.3% have dentures both above and below; the rest of the patients (64%) have crowns or bridges. However, again, no statistical analysis showed a significant association between the prosthesis and tumour location and onset<sup>31</sup>. In the study of Bektas-Kayhan et al<sup>28</sup>, 31 out of 47 patients (66%) had a fixed or removable prosthesis and the 44% of patients with tongue cancer had signs of mechanical trauma. However, the high percentage of smokers did not make simple to understand the role of the sole mechanical trauma in carcinogenesis. In another study, patients were divided into non-smokers and smokers/ former smokers. Smoking appears to be an important predisposing factor in males compared to females (72% vs. 28%) but, by eliminating the most important factors such as smoking and alcohol, women are more affected than men. In smokers, the localization of the tumour is prevalent in the edge of the tongue, in the floor. In non-smokers, on the other hand, the site mainly affected is that of the lingual border, and trauma was a probable causal factor<sup>25</sup>. In the study of Piemonte et al<sup>8</sup> patients were divided into smokers, non-smokers, drinkers, and non-drinkers and by different sources of trauma such as prostheses, dentures and teeth. The article highlights that the multivariate statistical analysis presents an association between trauma, age, and alcohol use while it does not present it for sex and the use of smoking. Rotundo<sup>26</sup> showed that smoking and alcohol are associated with tumour formation. The use of prostheses would not represent a risk (OR 0.67); this risk occurs if the prosthesis should cause the formation of ulcers and decubitus (OR 3.98). Out of 71 patients, 33.8% had a removable prosthesis. In the statistical analysis carried out, it was found that the presence of recurrent oral lesions caused by prostheses can increase the risk of developing oral carcinoma. However, in another study, Vaccareza *et al*<sup>11</sup> observed that the use of a prosthesis has an impact on tumour development. In multivariate analysis, however, the use of the prosthesis appears less incisive than other factors such as alcohol and tobacco.

# **CONCLUSIONS**

Due to the heterogeneity of the different studies and the strong presence of others factors as smoke and alcohol, the role of traumatism in carcinogenesis is not clear. Chronic trauma and its causal role in oral carcinogenesis remains a dilemma. Further molecular, epidemiological, and case-control studies are needed. Finally, especially in patients at risk, the dentist plays a fundamental role in an early diagnosis of SCC. In addition, patients at risk (drinkers and smokers or with a positive family history) with prostheses should be subjected to repeated checks over a short period of time in order to intervene early in the case of development of traumatic injury and consequent hypothetical development of a possible neoplastic lesion.

#### ACKNOWLEDGMENTS

One of the authors, CC performed his/her activity in the framework of the International PhD in Innovation Sciences and Technologies at the University of Cagliari, Italy.

#### **CONFLICT OF INTEREST:**

The Authors declare that there is no conflict of interest.

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