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Photodynamic Therapy for the Treatment of Oral HPV Lesion in a Patient with Oral Lichen Planus. A Case Report

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Abstract

Human papilloma viruses (HPVs) are a group of double-stranded DNA, that cause different type of lesions, also on oral cavity. Lichen planus (LP) is a chronic inflammatory mucocutaneous disease affecting the oral cavity and skin. Photodynamic therapy (PDT) is a treatment used in all fields of medicine, based on the interaction between a photosensitizer, the appropriate wavelength, and oxygen. The reaction generates ROS in cells that take up the photosensitizer, causing cell death by necrosis or apoptosis. Different parameters are described in the literature for PDT. We would like to report a case of a patient with diagnosis of Oral lichen planus that developed, and HPV lesion treated with a particular type of PDT. I have not found other cases of Photodynamic Therapy for the treatment of oral HPV lesions, in the scientific literature.

Keywords: HPV lesion; Oral lichen planus; Photodynamic Therapy; Toluidine Blu; Diode light

Introduction

Human papilloma viruses (HPVs) are a group of double-stranded DNA [1]. Human Papillomavirus (HPV) infection is considered the most common sexually transmitted infection. About 6 million people are diagnosed each year and approximately 9.0- 13.0% of the world population is already infected with the disease [2]. There are a lot of subtypes of HPV (more than 120), and some of these HPV plays a role in the pathogenesis of Head and Neck Squamous Cell Carcinomas and Oropharyngeal Squamous Cell Carcinoma [2,3]. Different types of HPV lesions could occur in oral cavity. Oral squamous papilloma is benign exophytic lesions presenting as papillary or verruciform proliferation. That is softened/flaccid in 66.7% of cases and pedunculated in 75% of the lesions. The most commonly associated with HPV 6, 11, and 16 [4]. Verruca vulgaris is a benign epidermal proliferation that affect overall the skin and sometimes the oral mucosa and the clinical presentations vary according to the viral type and the anatomical site infected. Verruca vulgaris is most commonly induced by HPV-2, HPV-4 or HPV-40 and it rarely occurs on the tongue [5]. Condyloma acuminate appear most frequently as soft, pink cauliflower like growths in moist areas, such as the genitalia, mouth and other places [6]. Focal

epithelial hyperplasia (FEH), or Heck's disease, is a rare disease of the oral mucosa, with multiple soft lesions associated with infection of subtypes 13 or 32 [7].

Lichen planus (LP) is a chronic inflammatory mucocutaneous disease affecting the skin, mucous membranes or both. It is recurrent and of unknown etiology. The typical clinical findings related to oral lichen planus (OLP), especially the presence of a reticular, bilateral and symmetrical pattern, are normally deemed sufficient for the clinical diagnosis of the disease, but a biopsy is recommended to confirm clinical diagnosis [8]. Various topically treatments are described for the OLP including: Corticosteroids, immunosuppressants such as cyclosporin, tacrolimus, and retinoids. The most commonly used drugs are Corticosteroids, but also azathioprine, calcineurin inhibitors, mycophenolate mofetil, dapsone, retinoids, and hydroxychloroquine can be used in recalcitrant cases [9].

Photodynamic therapy (PDT) is a particular type of treatment used in all fields of medicine, based on the interaction between a photosensitizer, the appropriate wavelength, and oxygen. The reaction generates ROS in cells that take up the photosensitizer,

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causing cell death by necrosis or apoptosis [10]. Different types of photosensitizer are used such as Methylen Blue, Toluidine Blue, Indocianine green and so on, to perform PDT. Different types of lights are described in the literature, lasers or LEDs, and there are a lot of different parameters for each type of disease or lesions [10]. I would like to describe a case of HPV lesion, in a patient with OLP, treated with a single session of PDT.

Case Presentation

A 58 years old female patient, with a previous diagnosis of oral lichen planus went to my private practice for a follow up visit. The medical anamnesis was positive for hysterectomy about 30 years ago, arthrosis, mild hypertension, gastro-esophageal reflux. A new asymptomatic lesion between the gingival papilla of the element 24 and 25 was present. It is an exophytic pink-white lesion, of irregular shape, of about 0,5 cm, near gingival lesion already present due to the oral lichen planus. The patient did not want to perform a histological examination in that day because she had an important event the day after. I have decided to treat the region between the gum around the element 24 and 25 with a session of photodynamic therapy (PDT). PDT was performed with a 630 nm LED red light (FotoSan 630, CMD Dental, Cophenagen, Denmark) in combination with Toluidine Blue (0,1 mg/ml). The lights applied have very mild affinity with mammalian cells.



Figure 1: HPV lesion in patient with a diagnosis of OLP.



Figure 2: Reduced HPV lesion after PDT.

For this there are no adverse effects during the treatments. The intensity of the light that emitted diodes is between 2000 and 4000MW/cm². This device works with three different modalities that correspond to different time cycles of application: 10, 20, and 30 seconds, respectively. The dye was applied on the entire surface of the lesion beyond the margins and even encroaching on healthy

tissue. The light diode was then turned on with a wavelength of 630 nm with 5 cycles of 30 seconds. a long-pipped tip was used and circular movements of about 0.5 cm above it were performed. At the end of the 5 applications, the dye was completely removed with a gauze and the patient performed a final rinsing with water. One week later, the exophytic lesion was reduced, of about 0,3 mm and hade a regular shape. The gingival tissue around the element 24 returned pink (Figures 1 & 2). I have decided to remove the exophytic lesion with a biopsy for an histological examination, that confirmed the diagnosis of squamous papilloma from HPV infection. After one week, at the follow up visit the tissues healed (Figure 3). I have informed the patient about the risk connected with the development of oral cancer in patient with oral lichen planus and HPV lesions.



Figure 3: Healing of the lesion.

Discussion

For the treatment of oral HPV the gold standard is considered surgical excision with different instruments. The use of Imiquimod has been documented in the literature, especially for small lesions [11]. Researchers also experimented the use of lasers, but I found no work on using PDT for HPV lesions [12,13]. Different types of photodynamic therapy have been used for the treatment of OLP, with very variable parameters. The alpha aminolaevulinic acid, methylene blue, toluidine blue are the most documented photosensitizers [14,15]. In some studies, it was seen that PDT for the treatment of OLP gave the same or better results to conventional therapy with corticosteroids or immunosuppressants [16]. The same device that was used in this work was tested for PDT on OLP erosive lesions, with excellent results [17]. Photodynamic therapy has also been used successfully for the treatment of other viral infections of the oral cavity, such as for herpes simplex virus [18,19]. Photodynamic therapy sessions could be a valid alternative to surgical excision in the treatment of squamous papilloma and could also be proposed to reduce the viral load of HPV, thus reducing the risk of possible malignant transformation. Further clinical studies must be performed to establish the most appropriate protocol, with the most effective parameters.

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