UltraPulse Carbon Dioxide Laser Plus Methyl Aminolevulinate-Photodynamic Therapy for the Treatment of Penile Cancer

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Keywords
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Abstract
The treatment of early-stage penile carcinoma is usually represented by wide excision or partial penectomy with or without inguinal lymph node dissection. However, laser ablation of the tumor may have a prominent role as an organ-sparing approach. In this regard, the combination of UltraPulse CO2 laser and photodynamic therapy (PDT) may be a valid option, especially when surgery is not feasible or refused. UltraPulse CO2 laser allows for the formation of gentle cutaneous abrasion that destroys the malignant tissue and, at the same time, improving the uptake of methyl aminolevulinate and amplifying the photochemical reaction of PDT in the tumor and surrounding tissue.

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Introduction

Penile cancer is a rare malignancy, with an incidence of about 1/100,000 men [1]. Usually, the treatment is represented by wide excision or partial penectomy with or without inguinal lymph node dissection [2]. However, laser ablation of the tumor may have a prominent role as an organ-sparing approach [2].

Case Report/Case Presentation

We report on the case of a 73-year-old Caucasian man, who presented to our institute with a 2-year-old history of multiple papulo-erythematous lesions on the glans (shown in Fig. 1a). These lesions were painless and fixed on palpation. The patient's past medical history included atrial fibrillation and circumcision for phimosis. A biopsy of one lesion was performed, showing pT1, G2 keratinizing squamous cell carcinoma infiltrating the sub-epithelial connective tissue. Therefore, the appropriateness of a wide surgical excision of the lesion was discussed. The patient refused any surgical procedure. Accordingly, an alternative treatment characterized by ablation with high-energy UltraPulse carbon dioxide (CO2) laser + photodynamic therapy (PDT) was discussed and deemed acceptable by the patient. We performed this combined treatment with the aim of allowing both the destruction of the tumor tissue and a greater penetration of the pharmacological treatment.

After a local disinfection and local anesthesia with mepivacaine hydrochloride 3%, the cutaneous lesions have been removed with 2 passages of UltraPulse CO2 laser at 80 mJ-5W using the handful Total-FXTM (Lumenis®, CA, USA), which created a spot of 0.8 mm in size. The ablation of the lesions was carried out with a minimum of 4 mm margin clearance (shown in Fig. 1b). Subsequently, methyl aminolevulinate (MAL) crème (Metvix 16%; Galderma,
France) was applied on the treated area, which was covered with an occlusive dressing for 3 h. After 3 h, the residual MAL creme was removed and the affected area was illuminated with 37 J/cm² red LED light, at 630 nm peak, and 70 mW/cm² for 7 min. After treatment, the patient experienced only a mild local burning sensation. Surgical eschars disappeared in 10 days using an emollient cream. The patient performed one session every 30 days for a total of 3 sessions. After 11 months of follow-up, resolution of the cutaneous lesions was evident on clinical observation (shown in Fig. 1c). There were no signs of local recurrence. A total body PET/CT was obtained and did not show any local or systemic recurrence.

**Discussion/Conclusion**

The treatment with CO₂ laser allows an effective vaporizing of affected areas, followed by the removal of the heat-separated tissue [1]. The UltraPulse system uses high fluencies that are delivered in short pulses to minimize nonspecific thermal damage to the adjacent tissues, permitting a precise treatment with minimal destruction of normal surrounding tissue, decreasing postoperative pain, decreasing recurrence, shortened healing time, bloodless field, and better cosmesis [3]. To ensure a faster and greater therapeutic response, we arbitrarily added MAL-PDT treatment in the same sessions [1, 2]. PDT relies on a light-induced photochemical reaction after the application of ester MAL, which metabolizes to the endogenous photosensitizer proto-porphyrin IX (PpIX) [4]. This reaction results in a release of singlet oxygen and other reactive oxygen species, leading to cell necrosis, allowing for a greater treated area, and inducing re-epithelization.

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Author Andrea Necchi was not available to confirm co-authorship, but the corresponding author Matteo Riccardo Di Nicola affirms that author Andrea Necchi contributed to the paper and had the opportunity to review the final version to be published and guarantees author Andrea Necchi co-authorship status and the accuracy of the author contribution and conflict of interest statements.

**Statement of Ethics**

Written informed consent was obtained from the participant for publication of the details of their medical case and any accompanying images. Study approval statement was not required for this study in accordance with local/national guidelines.

**Conflict of Interest Statement**

The authors have no conflicts of interest to declare.

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Author Contributions

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Data Availability Statement

All data generated or analyzed during this study are included in this article. Further inquiries can be directed to the corresponding author.

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