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Condyloma acuminatum (CA) is a sexually transmitted disease caused by human papilloma virus (HPV). Infection with HPV is a major risk factor for development of anal squamous cell carcinoma. In clinical practice, it is frequently limited to the perianal skin or anal canal and it is rarely described a rectal mucosal extension. Several therapeutic options are developed for CA, including chemical or physical destruction, immunological therapy, or a surgical excision. However, these treatments still have some degree of limitations and important side effects compromising patient compliance and reducing treatment efficacy. CO$_2$-laser emits a continuous beam, absorbed by biologic tissues, that vaporizing intracellular water, destroys target lesions. Argon plasma coagulator (APC) is a non-contact method of endoscopically delivered high-frequency thermal coagulation allowing well-controlled superficial tissue destruction. We present a case of a young female affected by anorectal condiloma extended to rectum, that was successfully treated with combined ginecological (carbon dioxide laser) and endoscopic approach (Argon Plasma Coagulator), with no side effects during the follow up and complete remission after two applications.

KEY WORDS: Argon Plasma, Carbon dioxide laser, Condyloma, HPV.

Introduction

Condyloma acuminatum (CA) is a sexually transmitted disease, caused by human papillomavirus (HPV) infection that has been increasing during the last decade. Infection with HPV is a major risk factor for development of anal or cervix squamous cell carcinoma. Despite over 100 genotypes of the virus, HPV 16 and 18 are considered pathogenic as they are seen in the majority of cervical and anal cancers. Epidemiologic studies identify high risk groups in: adolescents, young adults, persons with multiple sexual partners and human immunodeficiency virus (HIV) affected patients. There is not an agreement on the influence of sex, racial or demographics factors. Most HPV infections are cleared by the immune system and do not result in clinical manifestations. HPV viral subtype, immune status of the patient and environmental co-carcinogens determine the clinical presentation of infection. Anogenital lesions can be flat, dome-shaped, keratotic, pedunculated and cauliflower-shaped; they may occur singularly, in clusters or as plaques. Symptoms are mostly typical (anal itch/burning and external anal tags) but can be underestimated by patients that often delay treatment. Gold treatment for anogenital warts is still debated and not all methods have been completely tested. Therapeutic options for anogenital warts may be topical (podophyllin...
resin, imiquimod, trichloroacetic acid and podophyllotoxin) or surgical (carbon dioxide and Nd:YAG-laser vaporisation, surgical excision, loop excision, cryotherapy and electrodessication) treatments. However, recurrences are common and there is no single treatment that is superior to others. The Argon Plasma Coagulator (APC) is a device intended for thermal coagulation of tissue. Having been introduced in open and laparoscopic surgery, APC was adapted for use in flexible endoscopy in 1991 and has many potential uses. Endoscopic applications of APC continue to expand in the management of various gastrointestinal procedures that requiring: hemostasis, ablation of tissue or resolution of other technical problems (Tab. I). We present the case of a young female affected by an anorectal condiloma extended to rectum that was successfully treated by combined approach: carbon dioxide laser (CO2 Laser) for perineal lesions and APC for rectal localizations. This minimal-invasive treatment did not show side effects during the follow-up and reached a complete remission after two applications.

**Case presentation**

This the case of a 30 year-old female who was visited by gynecologist for the appearance of anal itching and burning and external anal painless tags, about 6 months before. The gynecologist, based on the anamnesis and clinical presentation diagnosed the presence of perineal condylomatous multiple warts in anogenital area and recommended serological test and surgical removal. Serological tests for syphilis and HIV gave negative results. Therefore, perineal warts were vaporized by CO2 laser, setting in pulse mode, after local anesthesia (2% lidocaine). Each condyloma was vaporized along with 3 mm border of normal-appearing epithelium without penetrating into the underlying subcutaneous tissue. A smoke evacuation system was used. Bleeding was coagulated with an electrosurgical system. Lidocaine spray for perineum was used 10 minutes later the procedure. The procedure was performed in the outpatient setting, no significant side effects were recorded. A follow-up visit was scheduled 30 days after the first treatment. Upon examination, the patient showed recurrence of perianal warts. So, the specialist recommended a further CO2

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<td>Shortening metallic stents</td>
<td>(*) Gastric antral vascular ectasia.</td>
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Fig. 1: (A) Rectal mucosa wart and first APC session: anorectal lesions in endoscopic retro-vision; (B) water-immersion vision; (C) elimination of intra-anal warts; (D) post-treatment result.
laser session to eradicate lesions together with a proctoscopy to rule out rectal involvment.

Anoscopy performed after deep sedation in an outpatient operating room, one week after CO2 session, revealed the presence of circumferentially rectal mucosa warts at about 1 cm above the dentate line (Fig. 1). Rectal biopsies were taken for histological confirmation. After the biopsy sample and polygeline-adrenaline solution infiltration (1:10.000), rectal warts were destroyed by APC in endoscopic retro-vision. Smoke evacuation system was used during the procedure to minimize airborne spread of the papillomavirus. All APC applications were performed by using the Olympus ENDOPLASMA® (Olympus Europe GmbH, Hamburg, Germany) combined with electrosurgical unit PSD-60. Energy transmission on the mucosa surface was guaranteed by a 2,3 mm diameter, 220 cm lengths, APC flexible probe (MAJ-1012, Olympus), with nozzle orientation at the tip of the catheter that was inserted through the working channel of a standard flexible endoscope. None early or late side effects (as pain or bleeding) were reported by the patient. Histological finding was consistent with condyloma acuminatum. An anoscopy follow-up was scheduled 3 months after the APC treatment. A the next endoscopic session, patient showed clearance of external disease and markedly decreasing of rectal lesions that was treated with a new APC session without adverse events (Fig. 2). Rectal biopsies were taken again for histological confirmation. Repeated flexible sigmoidoscopy was also done six months after in order to verify the successful eradication. No clinical or subclinical disease was shown at the anoscopy.

Discussion

There is an important link between the prevalence of squamous-cell cancer of the anal and HPV infection that is present in approximately 1% of the sexually active population. HPV infections may cause squamous intraepithelial neoplasia, which may progress from low grade to high grade and may be found in areas adjacent to squamous cell carcinoma. Previous reports demonstrating that HPV DNA was found in 75-90% of anal squamous cell cancer and in more than 95% of cervical carcinoma. In most cases, the CA diagnosis is based on anamnesis and clinical observation. Differential diagnosis must be made with the flat warts of syphilis (serological testing), molluscum contagiosum lesions, granuloma inguinale and Bowenoid papulosis. All patients with external anogenital warts should have proctoscopy and histological examination of all intra-anal lesions. There are many treatment options and patients are retreated at regular intervals until successful eradication. Topical treatment include podophyllotoxin, imiquimod cream and trichloroacetic acid. Surgical therapies include cryotherapy, CO2, and Nd:YAG-laser vaporisation, APC and surgical excision, that is highly effective but is generally reserved for large or bulky warts that cannot be treated otherwise. Patients with multiple sites of HPV infection could be less likely to respond to topical treatment. Indeed, combination of APC with intra-anal application of imiquimod cream could be effectively used in multiple CA localizations. However, Viazis N et al reported that the effectiveness of treatment with APC alone and APC plus imiquimod is not statistically significant. In our study, APC therapy has been used for the treatment of intra-anal warts proving to be easy, safe, and feasible. APC is a non-contact method of delivering high-frequency monopolar current in a controlled manner. The ionized argon gas contacts the tissue closest to the probe, which allows for either en face or tangential application, generally resulting in a 1-3 mm deep coagulation zone. Complication rates of APC are low, especially when compared with other thermal methods of treatment, and is thought to be secondary to the limited depth of coagulation. Moreover, using a submucosal injection of normal saline solution, the incidence of deep tissue injury decreases significantly. Considering that, a polygeline-adrenaline solution infiltration was adopted to even more decrease depth of tissue injury and to limit post-biopsy bleeding.

Conclusion

All patients with external anogenital warts should be submitted to proctoscopy and histological examination of intra-anal lesions. Elimination of these lesions is also important to reducing the risk of possible sexual transmission of HPV infection and to form a basis for prevention of anal cancer. Our experience, although limited to a case report, shows that patient affect by CA with perineal and intra-anal localizations, had symptomatic and endoscopic improvement after combined surgical treatment. This combined approach for multiple site condylomata between different surgical techniques, CO2-
laser and APC, should be considered as a safe and effective treatment. However, larger series of patients need to be studied to further assess the efficacy, complications rating and rate of recurrence.

Riassunto

Il condiloma acuminato è una malattia sessualmente trasmesa provocata dall’infezione da Papilloma Virus Umano e rappresenta un importante fattore di rischio per lo sviluppo del carcinoma anale a cellule squamose. Frequentemente si riscontra a livello della cute peri-anale e più raramente viene descritto a livello della mucosa rettale. Sono stati adottati diversi trattamenti per l’escissione delle lesioni condilomatose. Tra questi vi sono terapie mediche e chirurgiche che tuttavia presentano alcune limitazioni e spesso effetti collaterali che ne limitano l’efficacia e l’accettabilità da parte del paziente. Il laser a CO2 emette un fascio continuo che, assorbito dai tessuti biologici, elimina le lesioni trattate vaporizzando l’acqua all’interno delle cellule. L’Argon Plasma Coagulator è un metodica di escissione mediante coagulazione termica controllata, senza contatto, con la quale si ottiene una completa distruzione del tessuto superficiale. In questo studio riportiamo il caso di una giovane donna affetta da condilomatosi anorettale trattata con successo in maniera combinata, con approccio ginecologico (laser ad anidride carbonica) ed endoscopico (argon plasma coagulator), senza comparsa di effetti collaterali nel follow-up e con guarigione completa dopo due trattamenti.

References