MINIREVIEWS

6974 Applications of time series analysis in epidemiology: Literature review and our experience during COVID-19 pandemic
Tomov L, Chervenkov L, Miteva DG, Batselova H, Velikova T

ORIGINAL ARTICLE

6984 Acute cholangitis: Does malignant biliary obstruction vs choledocholithiasis etiology change the clinical presentation and outcomes?
Tsou YK, Su YT, Lin CH, Liu NJ

6995 Usefulness of analyzing endoscopic features in identifying the colorectal serrated sessile lesions with and without dysplasia

7004 Roles of biochemistry data, lifestyle, and inflammation in identifying abnormal renal function in old Chinese
Chen CH, Wang CK, Wang CY, Chang CF, Chu TW

7017 Clinical efficacy and safety of Guipi decoction combined with escitalopram oxalate tablets in patients with depression
Yu J, Xu FQ

7026 Artificial intelligence technology and ultrasound-guided nerve block for analgesia in total knee arthroplasty
Tong SX, Li RS, Wang D, Xie XM, Ruan Y, Huang L

7034 Axenfeld-Reiger syndrome: A search for the missing links
Morya AK, Ramesh PV, Sinha S, Nishant P, Nain N, Ramavath RN, Gone C, Prasad R

7043 Self-management of osteoarthritis while waiting for total knee arthroplasty during the COVID-19 pandemic among older Malaysians
Mahdzir ANK, Mat S, Seow SR, Abdul Rani R, Che Hasan MK, Mohamad Yahaya NH

7053 “In situ bone flap” combined with vascular pedicled mucous flap to reconstruction of skull base defect

7061 Reference values of gait parameters in healthy Chinese university students: A cross-sectional observational study
<table>
<thead>
<tr>
<th>Page</th>
<th>Title</th>
<th>Authors</th>
</tr>
</thead>
<tbody>
<tr>
<td>7075</td>
<td>Effect of T-regulatory cells and interleukin-35, interleukin-10, and transforming growth factor-beta on diffuse large B-cell lymphoma</td>
<td>Wu H, Sun HC, Ouyang GF</td>
</tr>
<tr>
<td>7091</td>
<td>Meta-analysis of the efficacy and safety of daratumumab in the treatment of multiple myeloma</td>
<td>Wang P, Jin SY</td>
</tr>
<tr>
<td>7101</td>
<td>Varicella-zoster virus meningitis with hypoglycorrhachia: A case report</td>
<td>Cao LJ, Zheng YM, Li F, Hao HJ, Gao F</td>
</tr>
<tr>
<td>7107</td>
<td>Unusual presentation of penile giant condyloma acuminatum with spontaneous prepuce perforation: A case report</td>
<td>Hsu FC, Yu DS, Pu TW, Wu MJ, Meng E</td>
</tr>
<tr>
<td>7113</td>
<td>Primary renal lymphoma presenting as renal failure: A case report and review of literature from 1989</td>
<td>Lee SB, Yoon YM, Hong R</td>
</tr>
<tr>
<td>7136</td>
<td>Mucoepidermoid carcinoma of the lung with hemoptysis as initial symptom: A case report</td>
<td>Xie WX, Liu R, Li Z, Zhou PL, Duan LN, Fu DD</td>
</tr>
<tr>
<td>7144</td>
<td>Co-infection of <em>Chlamydia psittaci</em> and <em>Tropheryma whippelii</em>: A case report</td>
<td>Du ZM, Chen P</td>
</tr>
<tr>
<td>7150</td>
<td>Surgical treatment of severe anterior capsular organized hard core cataract: A case report</td>
<td>Wang LW, Fang SF</td>
</tr>
<tr>
<td>7156</td>
<td>First platelet transfusion refractoriness in a patient with acute myelocytic leukemia: A case report</td>
<td>Tu SK, Fan HJ, Shi ZW, Li XL, Li M, Song K</td>
</tr>
<tr>
<td>7162</td>
<td>Rare finding of primary aortoduodenal fistula on single-photon emission computed tomography/computed tomography of gastrointestinal bleeding: A case report</td>
<td>Kuo CL, Chen CF, Su WK, Yang RH, Chang YH</td>
</tr>
<tr>
<td>7170</td>
<td>Rituximab combined with Bruton tyrosine kinase inhibitor to treat elderly diffuse large B-cell lymphoma patients: Two case reports</td>
<td>Zhang CJ, Zhao ML</td>
</tr>
</tbody>
</table>
## Contents

**World Journal of Clinical Cases**

**Thrice Monthly Volume 11 Number 29 October 16, 2023**

<table>
<thead>
<tr>
<th>Page</th>
<th>Title</th>
<th>Authors</th>
</tr>
</thead>
<tbody>
<tr>
<td>7179</td>
<td>Use of Ilizarov technique for bilateral knees flexion contracture in Juvenile-onset ankylosing spondylitis: A case report</td>
<td>Xia LW, Xu C, Huang JH</td>
</tr>
<tr>
<td>7187</td>
<td>Case of takotsubo cardiomyopathy after surgical treatment of liver hydatid cyst: A case report</td>
<td>Altas Y, Abdullayeva U</td>
</tr>
<tr>
<td>7193</td>
<td>Laparoscopic choledocholithotomy and transductal T-tube insertion with indocyanine green fluorescence imaging and laparoscopic ultrasound: A case report</td>
<td>Yoo D</td>
</tr>
<tr>
<td>7200</td>
<td>Hematopoietic stem cell transplantation of aplastic anemia by relative with mutations and normal telomere length: A case report</td>
<td>Yan J, Jin T, Wang L</td>
</tr>
<tr>
<td>7207</td>
<td>Emphysematous thrombophlebitis caused by a misplaced central venous catheter: A case report</td>
<td>Chen N, Chen HJ, Chen T, Zhang W, Fu XY, Xing ZX</td>
</tr>
<tr>
<td>7214</td>
<td>Aggressive angiomyxoma of the epididymis: A case report</td>
<td>Liu XJ, Su JH, Fu QZ, Liu Y</td>
</tr>
<tr>
<td>7221</td>
<td>Gastric and intestinal ectopic pancreas: Two case reports</td>
<td>Zhang H, Zhao HY, Zhang FH, Liang W</td>
</tr>
<tr>
<td>7234</td>
<td>Imaging misdiagnosis and clinical analysis of significant hepatic atrophy after bilioenteric anastomosis: A case report</td>
<td>Liang SY, Lu JG, Wang ZD</td>
</tr>
<tr>
<td>7248</td>
<td>Simultaneous thyroglossal duct cyst with parathyroid cyst: A case report</td>
<td>Chen GY, Li T</td>
</tr>
<tr>
<td>7253</td>
<td>Submandibular solid-cystic mass as the first and sole manifestation of occult thyroid papillary carcinoma: A case report</td>
<td>Chen GY, Li T</td>
</tr>
<tr>
<td>7258</td>
<td>Artificial intelligence and machine learning in motor recovery: A rehabilitation medicine perspective</td>
<td>Swarnakar R, Yadav SL</td>
</tr>
</tbody>
</table>

**LETTER TO THE EDITOR**

<table>
<thead>
<tr>
<th>Page</th>
<th>Title</th>
<th>Authors</th>
</tr>
</thead>
<tbody>
<tr>
<td>7258</td>
<td>Artificial intelligence and machine learning in motor recovery: A rehabilitation medicine perspective</td>
<td>Swarnakar R, Yadav SL</td>
</tr>
</tbody>
</table>
ABOUT COVER
Editorial Board Member of World Journal of Clinical Cases, Zeid J Khitan, FACP, FASN, MBBS, MD, Academic Research, Director, Full Professor, Department of Medicine, Marshall University, Huntington, WV 25701, United States. zkhitan@marshall.edu

AIMS AND SCOPE
The primary aim of World Journal of Clinical Cases (WJCC, World J Clin Cases) is to provide scholars and readers from various fields of clinical medicine with a platform to publish high-quality clinical research articles and communicate their research findings online.

WJCC mainly publishes articles reporting research results and findings obtained in the field of clinical medicine and covering a wide range of topics, including case control studies, retrospective cohort studies, retrospective studies, clinical trials studies, observational studies, prospective studies, randomized controlled trials, randomized clinical trials, systematic reviews, meta-analysis, and case reports.

INDEXING/ABSTRACTING
The WJCC is now abstracted and indexed in Science Citation Index Expanded (SCIE, also known as SciSearch®), Journal Citation Reports/Science Edition, Current Contents®/Clinical Medicine, PubMed, PubMed Central, Reference Citation Analysis, China National Knowledge Infrastructure, China Science and Technology Journal Database, and Superstars Journals Database. The 2023 Edition of Journal Citation Reports® cites the 2022 impact factor (IF) for WJCC as 1.1; IF without journal self cites: 1.1; 5-year IF: 1.3; Journal Citation Indicator: 0.26; Ranking: 133 among 167 journals in medicine, general and internal; and Quartile category: Q4.

RESPONSIBLE EDITORS FOR THIS ISSUE
Production Editor: Hua-Ge Yu; Production Department Director: Xiang Li; Editorial Office Director: Jin-Lei Wang.
Unusual presentation of penile giant condyloma acuminatum with spontaneous prepuce perforation: A case report

Fu-Chieh Hsu, Dah-Shyong Yu, Ta-Wei Pu, Min-Jui Wu, En Meng

Specialty type: Medicine, research and experimental
Provenance and peer review: Unsolicited article; Externally peer reviewed.
Peer-review model: Single blind
Peer-review report's scientific quality classification
Grade A (Excellent): 0
Grade B (Very good): 0
Grade C (Good): C, C
Grade D (Fair): 0
Grade E (Poor): 0
P-Reviewer: Ali N, Finland; Sarier M, Turkey
Received: June 6, 2023
Peer-review started: June 6, 2023
First decision: August 30, 2023
Revised: September 8, 2023
Accepted: September 22, 2023
Article in press: September 22, 2023
Published online: October 16, 2023

Abstract

BACKGROUND
Giant condyloma acuminatum (GCA) is an uncommon condition affecting the anogenital area.

CASE SUMMARY
Here, we report an 88-year-old male patient who presented with a 5-year history of a progressive appearance of multiple cauliflower-like warts over his penile shaft, pubis region, and groin as well as urine leakage along the distal penile shaft. Physical examination revealed an ulcerative skin perforation with pus discharge under the distal prepuce base, which was initially suspected to be a urethral fistula. However, during surgery, it was discovered that the perforation was caused by a giant condyloma lesion that had obliterated the prepuce opening, with infection and high pressure causing subsequent skin perforation. He underwent circumcision and wide excision with electrocauterization of the warts. He was discharged after the surgery, and the residual lesion was treated with imiquimod and low-dose oral tegafur-uracil.

CONCLUSION
Penile GCA can cause prepuce perforation and can be postoperatively treated with imiquimod and low-dose oral tegafur-uracil.

Key Words: Giant condyloma acuminatum; Prepuce perforation; Buschke-Löwenstein tumor; Fistula; Foreskin; Case report
Core Tip: This case report highlights a rare presentation of a giant condyloma acuminatum (GCA) in an elderly male patient. The patient presented with progressive, cauliflower-like warts on the penile shaft, pubis region, and groin, accompanied by urine leakage and an ulcerative skin perforation with pus discharge under the prepuce. Surgery revealed that the perforation was caused by an obliteratorative giant condyloma lesion, mimicking urethral rupture. The patient underwent circumcision, wide excision of warts, and tailored subsequent treatment with favorable outcomes. This case emphasizes the importance of individualized treatment for GCA and the successful management of spontaneous prepuce perforation.

INTRODUCTION

Giant condyloma acuminatum (GCA), also known as Buschke-Lowenstein tumor, was first reported by Abraham Buschke in 1896 and later by Loewenstein in 1925[1]. It typically appears as a slowly growing mass resembling a cauliflower in the genital or anorectal region. The tumor tends to infiltrate deeper tissues at a relatively slow rate[2]. Here, we present a case of penile GCA where the initial symptom mimicked a urethral fistula resulting from the obliteration of the preputial opening by the giant condyloma lesion and subsequent perforation of the distal preputial skin.

CASE PRESENTATION

Chief complaints
An 88-year-old male patient complained of multiple warts over the pubis region and exogenitalia for 5 years and urine leakage from the distal penis noted for 2 wk.

History of present illness
He denied multiple sexual partners. The patient was not immunocompromised and was negative for human immunodeficiency virus and syphilis.

History of past illness
The patient denied any prior systemic diseases.

Personal and family history
The family history revealed nothing noteworthy.

Physical examination
Physical examination revealed an ulcerative distal preputial hole, measuring 1 cm in diameter, with pus discharge under the ventral penile shaft so that the initial impression was a urethral fistula (Figure 1A). In addition, multiple condyloma lesions and plaques, presenting as cauliflower-like warts, were noted over the exogenitalia, penile shaft, pubis, and groin region (Figure 1B). The warts were clustered, gray-to-black in color, hard, and not painful. Some warts were spread sporadically over the scrotum, and multiple verrucous plaques were also found in the perianal region on rectal examination.

Laboratory examinations
A serum test revealed leukocytosis with elevated white blood cell levels of 12.13 (10^3/μL). The urine sediment examination revealed pyuria, characterized by an elevated urine white blood cell count of 10-20 per high-power field. All lab results were found to be normal during the follow-up interval.

Imaging examinations
Distant metastasis was excluded through chest and abdominal computed tomography scans.
Figure 1 Images of physical examination. A: An ulcerative hole (indicated by a orange arrow) measuring 1 cm in diameter, with pus discharge, was located under the base of the prepuce; B: Multiple condyloma lesions were present over the exogenitalia, penile shaft, pubis, and groin region, appearing as cauliflower-like warts and plaques.

**FINAL DIAGNOSIS**

The pathology of the circumcised redundant prepuce (Figure 2) confirmed the diagnosis of giant condyloma acuminata with moderate dysplasia and destructive local invasion without metastasizing. The human papillomavirus (HPV) genotyping results for the redundant prepuce specimen indicated a positive presence of HPV-11.

**TREATMENT**

During the operation, we found that the prepuce opening was completely obliterated by a giant condyloma lesion with a distal preputial perforation, which mimicked a urethral fistula. Urine drained from a spontaneous perforation that was present directly below the glans, nearly at the level of the coronal sulcus. Finally, circumcision along the hole and wide excision with electrocauterization of the residual warts were performed.

**OUTCOME AND FOLLOW-UP**

The postoperative course was uneventful, and the wounds healed gradually with adjuvant topical 5% imiquimod cream thrice a week and oral tegafur-uracil (UFUR), containing 100 mg of tegafur and 224 mg of uracil, two pills per day, for 6 mo (Figure 3).

**DISCUSSION**

CA lesions, which grow extensively and display invasive local growth patterns without spreading to distant sites, are referred to as Buschke-Löwenstein tumors. These tumors are also known as verrucous carcinoma or giant condylomas. Buschke-Lowenstein tumors were first reported by Abraham Buschke in 1896 and later by Loewenstein in 1925[1]. GCA is an uncommon condition that is characterized by a cauliflower-like growth with exophytic extension and tends to invade nearby tissues. This condition has a high rate of recurrence and carries the potential for malignant transformation [3]. GCA is associated with various types of HPV, immunosuppression, and other risk factors that require further investigation. GCA is associated with several factors, such as having multiple sexual partners, chronic genital infections, poor hygiene, and immunodeficiencies[4].

An estimated prevalence of 0.1% has been reported in the general population, with the condition usually presenting between the 4th and 6th decades of life[3,5]. In the latest case series, the male-to-female ratio was reported as 2.7:1, with patients ranging in age from 24 to 77 years old and a mean age of 43.9 years. Additionally, there was a tendency for the condition to present at younger ages[6].

Cases of GCA have been reported in the pediatric age group, and a limited number of cases also have been reported in pregnant women[7,8]. The types of HPV causing condyloma acuminatum warts of the anal and genital regions were 6, 11, 42, 44, and 54[9]. Persistent HPV infection was observed in 10% of patients with GCA, and the progression to invasive cancer in such cases typically extends over a 15 to 20-year timeframe[10]. Intraepithelial neoplasia commonly arises during the subclinical phase, while the clinical phase is characterized by the presence of visible lesions and symptoms such as condyloma acuminatum or invasive cancer[11]. Papillomatous lesions are frequently observed in the male genital
area, including the penis, scrotum, urethral opening, and perianal region, and in the female genital area, including the vaginal opening, vulva, perineum, and perianal region.

Because of the absence of a dependable serological test to distinguish between past and current HPV infections and the inability to isolate the virus, the definitive diagnosis relies on the detection of HPV-DNA using molecular tests. Nucleic acid amplification tests, such as polymerase chain reaction (PCR), are presently employed as the established benchmark for HPV genotyping. At present, there is no universally agreed-upon standard for the most effective sampling technique for HPV DNA PCR testing in circumcised heterosexual men. Sarier et al. designed a study involving six different anatomical sampling sites for HPV DNA PCR testing in circumcised heterosexual men with genital warts. Their findings revealed that the detection rate approaches 99% when wart biopsy is combined with swab sampling of the penile shaft, coronal sulcus, scrotum, and external urethral meatus.

GCA is primarily characterized by the presence of a perineal mass, but patients may also experience symptoms such as constipation, hemorrhoids, difficulty defecating and urinating, dysuria, abdominal distension, and fatigue. Interestingly, based on initial examination, our case presented as a urethral fistula resulting from an obstruction in the urethra, caused by the growth of GCA up to the meatus. It was not until further surgical examination that we discovered that the preputial opening had been obstructed by GCA, causing the perforated prepuce to present in a manner mimicking a urethral fistula. This event has not been reported in GCA-related case studies. Theoretically, the slow growth pattern of GCA led to chronic inflammation and infection within the prepuce, resulting in the accumulation of pus and eventual spontaneous perforation of the prepuce. The resulting perforated hole allowed for urine diversion and successful decompression of preputial obstruction.

The malignancy of GCA can be considered in two distinct ways: Through the confirmation of neoplastic histology in malignant transformation and through the infiltration of adjacent deep tissues, regardless of whether histological confirmation of malignancy is present, in the case of malignant behavior. Trombetta and Place reported that in their series of 51 patients with GCA, 58% of the patients had neoplasia present at the time of diagnosis. Chu et al. observed a neoplastic transformation rate of 56% in their series of 42 patients with GCA, as 56% of the patients had histologically
confirmed malignancy during follow-up. In cases where squamous cell carcinoma develops in the presence of GCA, treatment must be tailored to the patient’s individual circumstances, and in some cases, only palliative care may be provided to prevent the development of further comorbidities[14].

In the past, various therapeutic options have been used for the treatment of GCA, including topical chemotherapy, intraligual injection of 5-fluoracil (5-FU), interferon, cryotherapy, curettage, CO2 Laser vaporization, wide resection alone or with neoadjuvant or adjuvant chemotherapy, chemotherapy alone, radiation therapy, and isolated perfusion[2]. Nevertheless, a universally accepted treatment approach with optimal effectiveness for anogenital warts remains elusive. It is understood that even after the disappearance of visible lesions following treatment, the virus can still be detected in apparently normal epithelial tissue for several months. The use of podophyllin, which inhibits cell division during the metaphase stage and induces localized tissue necrosis, resulting in the eradication of warts, has yielded unsatisfactory outcomes for GCA, but it has demonstrated positive outcomes for cases of classic condyloma acuminatum[2]. In the highly uncommon instances of extragenital GCA where resection is restricted owing to potential functional impairment, the use of combined therapy involving topical podophyllotoxin and imiquimod, along with weekly cryotherapy, has been documented as effective. This approach offers advantages over surgical treatment alone, as it does not result in functional consequences[15]. Surgical excision is particularly favored for solitary and giant genital verrucous lesions because it allows for both therapeutic intervention and diagnostic pathological assessment to detect precancerous changes[11]. Performing surgical treatment with negative margins is currently considered the optimal approach and should be pursued whenever feasible[16]. In the field of urology, surgical excision is particularly effective when dealing with cases of anterior urethral carcinoma[17]. Various treatment combinations, in addition to surgery as both neoadjuvant and adjuvant therapy, have been reported, yielding varying outcomes[18]. However, there is no standardized selection process to determine the appropriate therapeutic approach. It is advisable to adopt an individualized therapeutic approach for each case[19]. There are multiple options for surgical treatment to remove GCA, including traditional surgery, as well as alternative methods such as electrosurgery, radiofrequency, and carbon laser surgery, which is used for intramural and common extragenital verrucous. Because of the high likelihood of recurrence, it is strongly advised to conduct long-term follow-up for individuals with GCA[20].

Although the estimated rate of malignant transformation is as high as 56%, there is no propensity for metastasis[2]. In addition, given the advanced age of the patient and the recommended first-line treatment protocol of 5-FU-based chemotherapy[6], we tailored a treatment plan for this patient that involved local wide excision with circumcision and electrocautery combined with adjuvant topical 5% imiquimod cream thrice a week and oral UFUR, containing 100 mg of tegafur and 224 mg of uracil, two pills per day, rather than a more aggressive reconstructive surgery. The patient continued to use the above medication for six months, and the wound remained in good condition without new local wart recurrence.

**CONCLUSION**

While it is rare to see penile GCA cases mimicking a urethral fistula, our case demonstrates that obliteration of the preputial opening in GCA can cause spontaneous prepuce perforation. The obliteration of the prepuce and glans by GCA caused chronic inflammation and infection, with eventual spontaneous perforation of the prepuce.

**FOOTNOTES**

**Author contributions:** Hsu FC contributed to data collection and manuscript writing; Wu MJ and Meng E contributed to conceptualization and supervision; Yu DS and Pu DW contributed to manuscript revision and study supervision; and all authors have read and approved the final manuscript.

**Informed consent statement:** Informed written consent was obtained from the patient for the publication of this report and any accompanying images.

**Conflict-of-interest statement:** All the authors report no relevant conflicts of interest for this article.

**CARE Checklist (2016) statement:** The authors have read the CARE Checklist (2016), and the manuscript was prepared and revised according to the CARE Checklist (2016).

**Open-Access:** This article is an open-access article that was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution NonCommercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: https://creativecommons.org/Licenses/by-nc/4.0/

**Country/Territory of origin:** Taiwan

**ORCID number:** Fu-Chieh Hsu 0000-0001-8071-1635; Dah-Shyong Yu 0000-0001-7594-0070; Ta-Wei Pu 0000-0002-0538-407X; Min-Jui Wu 0009-0008-5402-9374; En Meng 0000-0002-1262-5361.

**S-Editor:** Wang JJ

https://www.wjgnet.com
REFERENCES


