Diagnosis and treatment of epididymal tuberculosis: A review of 47 cases

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Abstract:

Objective: To analyze the clinical manifestations, diagnosis, and treatment outcomes in a series of patients with epididymal tuberculosis.

Methods: This study is a retrospective data analysis of 47 cases of histologically-confirmed epididymal tuberculosis in patients treated at our hospital from November 2014 to December 2018.

Results:The average age of the patients was approximately 41.98 years.The epididymal lesion location was left-sided in 15 patients(31.9%),right-sided in 22 patients (46.8%),and bilateral in 10 patients (21.3%).The main symptoms were painless swelling of the scrotum in 21 cases (44.7%) and scrotal drop pain in 21 cases (44.7%). Scrotal physical examination revealed epididymal beaded enlargement in 12 patients (25.5%), testicular mass in 1 patient (2.1%), scrotal tenderness alone in 7 patients (14.9%), ill-defined epididymal-testicular border in 21 patients (44.7%), and sinus formation in 6 patients (12.8%). After 2 to 4 weeks of anti-tuberculosis chemotherapy, the patients underwent a surgical resection procedure. We found that 10 (83.3%) of the 12 patients whose main symptom was epididymal beaded enlargement underwent simple epididymal resection .Of the 21 patients whose main clinical manifestation was ill-defined testis-epididymis demarcation, 16 (72.2%) underwent epididymis-testicular resection.All patients underwent postoperative chemotherapy for 3 to 6 months. Postoperative follow-up showed good response to treatment.

Conclusion:It is difficult to diagnose early-stage epididymal tuberculosis. Epididymal tuberculosis is likely to have invaded surrounding tissues when signs such as epididymal beaded changes and ill-defined epididymis-testis border are present. Surgical resection combined with preoperative and postoperative chemotherapy is an effective approach to treating this condition.

Keywords:epididymal tuberculosis, clinical characteristics, chemotherapy

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Introduction: Epididymal tuberculosis is a rare extrapulmonary form of tuberculosis that occurs in young adults(Surati et al. 2012). Patients with this disease may have no obvious clinical symptoms or only mild symptoms. The disease typically develops slowly and early diagnosis is difficult; delayed diagnosis and misdiagnosis are common. Recently, due to the emergence of multi-drug resistant bacteria (MDR), anti-tuberculosis drug resistance, and the widespread use of glucocorticoids, the incidence of male genitourinary tuberculosis. This term is incorrect. Please use "Male genital TB" or "Urogenitak TB", including epididymal tuberculosis, has been increasing worldwide. Epididymal tuberculosis remains an important health problem in many developing countries, including China(Kulchavenya et al. 2012). The pathological features of epididymal tuberculosis are extensive tissue destruction and fibrosis, eventually leading to destruction of the epididymis and surrounding genital tissues and organs and complications such as infertility and other serious effects on male reproductive system function.

At present, early diagnosis and regular chemotherapy are the keys to curing male reproductive system tuberculosis and to avoid surgical treatment. However, due to its late onset and nonspecific clinical signs and symptoms, and the lack of rapid, sensitive, and specific diagnostic methods, the disease is often misdiagnosed or the diagnosis is delayed. Therefore, surgical treatment combined with chemotherapy has been the preferred treatment approach to this disease. This study, a retrospective analysis of a series of cases of epididymal tuberculosis, is intended to more fully characterize the clinical features and outcomes of surgical treatment of this disease.

Material and Methods: We collected the medical records of 47 patients with histologically-diagnosed epididymal tuberculosis from November 2012 to October 2018 at the Second Hospital of Lanzhou University. Age, clinical signs and symptoms, diagnostic methods, and treatment effects were collected from the patients' electronic medical records, and the final results were compared and analyzed. The following information was supplied relating to ethical approvals (i.e., approving body and any reference numbers): Ethics Committee of Lanzhou University Second Hospital Number:2019A-079. Written consent was obtained from all patients.

Results:_The mean age of the patients was 41.98 years (range, 19-72 years); 15 patients (31.9%) had tuberculosis in the left epididymis, 22 patients (46.8%) had tuberculosis in the right epididymis, and 10 patients (21.3%) had bilateral epididymal tuberculosis. Of the 47 patients, 8 (17.0%) had a history of tuberculosis, including 3 cases of pulmonary tuberculosis, 4 cases of renal tuberculosis, and 1 case of prostate tuberculosis (Table 1). What about active TB alongside with actual disese?

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Main symptoms at the time of onset included: painless swelling of the scrotum in 21 patients (44.7%); scrotal drop pain in 21 patients (44.7%); urinary tract irritation such as urinary frequency, dysuria, and hematuria in 4 patients (8.5%); and scrotal skin ulceration in 1 patient (2.1%). Systemic symptoms such as low-grade fever, fatigue, and night sweats occurred in 6 patients (12.8%). All patients underwent scrotal physical examination, which revealed epididymal beaded enlargement in 12 patients (25.5%), testicular mass in 1 patient (2.1%), scrotal tenderness alone in 7 patients (14.9%), ill-defined epididymal testicular border in 21 patients (44.7%), and sinus formation in 6 patients (12.8%)(Table 1).

Urinalysis and chest imaging were performed in all patients._Twenty patients (42.6%) were positive for white blood cells in urine and 8 patients (17.0%) were positive for red blood cells in urine. Did kidney MSKT performed for them? Chest imaging was positive in 8 patients (17.0%).Please explain The preoperative diagnoses were epididymal tuberculosis in 41 patients (87.2%), epididymal tumor in 3 patients (6.4%), and epididymal mass in 3 patients (6.4%). All patients had evidence of surgical resection of the lesion. How did you explain pathology in urinalyses?

Among the 47 cases, 41 (87.2%) were initially identified as epididymal tuberculosis and treated pre-operatively with rifampicin, isoniazid, pyrazinamide, and ethambutol for 2 to 4 weeks. Twenty-six patients (55.3%) had epididymectomy and

21 patients (44.7%) had epididymectomy combined with orchiectomy— and all recovered well after surgery.

We found that 10 (83.3%) of the 12 patients whose main symptom was epididymal beaded enlargement underwent simple epididymal resection and 2 (16.7%) underwent epididymis-testicular resection. Of the 21 patients whose main clinical manifestation was ill-defined testis-epididymis demarcation, 16 (72.2%) underwent epididymis-testicular resection and 5 epididymal resection (Table 2).

Macroscopically, all specimens contained solitary or confluent pale, grayish caseous necrotic nodules. Some lesions invaded the entire epididymis, some showed involvement of the testis with adherence to the scrotum to form a cold abscess, and some showed the formation of ulcerated sinus tracts in the skin. Microscopically, the centers of the specimens were red-stained with amorphous granular, irregularly sized foci of caseous necrosis surrounded by tuberculous granulation tissue (epithelioid cells, Langhans giant cells, and lymphocytes) (Figure 1,2).

All patients received anti-tuberculosis treatment with rifampicin, isoniazid, pyrazinamide, or ethambutol after surgery for 3 to 6 months. No recurrence was reported.

Discussion:

The number of TB patients in China accounted for 10%-12% of the total number of TB patients in the world, ranking second in the world(Glaziou et al. 2018; Zhang et al. 2016). People in Gansu Province of China have a high incidence of tuberculosis and are relatively delayed in seeking medical care(Zhang et al. 2016). The presentation of tuberculosis is increasingly atypical. Unfortunately, the prevalence of drug-resistant strains of TB is increasing(Lee et al. 2015). The main treatment of epididymal tuberculosis is early anti-tuberculosis treatment. However, since most of the patients with epididymal tuberculosis in Gansu are in the terminal stage, surgical treatment combined with chemotherapy has been the best treatment modality for this type of patients.

Reproductive system tuberculosis can occur in any age, mainly in men 30 to 50 years old. Due to the long incubation period, it is not common in children. The most commonly involved organ is the epididymis, followed by the seminal vesicle, prostate, testis, and vas deferens(Yadav et al. 2017). Isolated epididymal tuberculosis is very rare(Gueye et al. 1998). One new study indicates that isolated epididymal tuberculosis may be the first or only manifestation of early genitourinary tuberculosis(Viswaroop et al. 2005). Similarly, in our patient cohort, there were 39 cases (82.9%) of isolated epididymal tuberculosis.

The pathogenesis of epididymal tuberculosis includes blood-borne transmission and transurethral reflux of *Mycobacterium tuberculosis* caused by factors such as trauma, alcohol abuse and excessive sexual activity(Tzvetkov & Tzvetkova 2006). Epididymal tuberculosis lesions first appear in the tail of the epididymis, owing to its rich blood supply and to retrograde infection from the vas deferens. The lesions of epididymal tuberculosis gradually invade the body to the head, finally affecting the entire epididymis. In severe cases, the testis can be involved(Chung et al. 1997). In this group, 21 patients (44.7%) with epididymal tuberculosis invading the testis underwent radical resection Resection can not be radical. It always is partial. of the testicle. In our study, the main cause of the patient's visit was painless mass of the epididymis, which is consistent with previous studies(Kho & Chan 2012). However, the proportion of patient with scrotal pain is higher, mainly because the patients included in this study all had advanced epididymal tuberculosis or involvement of the testis or scrotum.

The gold standard for diagnosing tuberculosis is the isolation and culture of *Mycobacterium tuberculosis*. In cases of suspected genitourinary tuberculosis incorrect term, we usually look for *M. tuberculosis* in the urine or tissue. Traditionally,

the appearance of so-called sterile pyuria on microscopic urinalysis is considered to be a typical manifestation of urogenital involvement. In our patient cohort, leukocytes in urine were present microscopically (50%) or grossly in a majority of cases (50% and 10%, respectively). Again, what was a reason? Kidney TB? Hematuria and acidic urine have been associated with urinary tuberculosis, but they are nonspecific findings. Is it your date? Color Doppler ultrasound is the first choice for imaging analysis of epididymal tuberculosis(Viswaroop et al. 2005). CT and MR have little value in diagnosing epididymal tuberculosis; they are mainly used to diagnose tuberculosis in lung and kidney and provide support for the diagnosis of epididymal tuberculosis. Epididymal fine needle biopsy is a good method for the diagnosis of epididymal tuberculosis.

Tuberculous epididymitis can be the only manifestation of genitourinary tuberculosis. Therefore, even in the absence of clinical and laboratory markers of renal and urologic tuberculosis, all men with identifed epididymal lesions should undergo a fine needle aspiration biopsy. Kim(Kim et al. 1993) suggests that epididymal tuberculosis can often be diagnosed by B-ultrasound biopsy, which supports the above conclusions. Polymerase chain reaction has been an important method for diagnosme. Combined with pathological biopsy, it can improve the diagnosis rating epididymal tuberculosis in recent years(Chawla et al. 2012). It has the characteristics of high sensitivity, high specificity, and short turnaround tie. However, at present, the diagnosis of simple epididymal tuberculosis is difficult, and there is no preoperative diagnostic method with high sensitivity and specificity. The clinical diagnosis of scrotal abscess and ulceration is not difficult. It is often possible to make a clear diagnosis by examining for acid-fast bacilli in samples of the ruptured tissue through pus or secretion smears.

The differential diagnosis of epididymal tuberculosis includes bacterial epididymitis, epididymal sperm granuloma, epididymal tumor, and other diseases. Patients with bacterial epididymitis often have testicular-epididymis pain, and scrotal swelling and heat. Features include a shorter course of illness, and symptoms are generally relieved after antibiotic treatment. Carl(Carl & Stark 1997) reported that epididymal tuberculosis should be highly suspected when patients have persistent or repeated epididymitis episodes and symptoms cannot be controlled after adequate antibiotic treatment.

Epididymal sperm granuloma mainly occurs in the head of the epididymis, characterized by a smooth solid mass, and antibiotics and anti-tuberculosis treatment are ineffective. Epididymal tumors are rare, accounting for about 0.9% of male reproductive system tumors. Most tumors occur during periods of sexual activity in

men between 20 and 40 years of age. The disease is characterized by slow growth, large volume, and no tenderness. CT helps to confirm the diagnosis.

Epididymal tuberculosis, like other tuberculosis diseases, requires early, regular, full-course, moderate, combined anti-tuberculosis treatment. The drug treatment method uses 3 to 4 anti-tuberculosis drugs for 6 to 9 months(Peloquin 2014). Surgical treatment is necessary if there is no response to drug treatment or in cases of cold abscess not only cold formation. Because the early symptoms of epididymal tuberculosis are not obvious, abscesses or involvement of surrounding tissues such as the testicles have often developed at the time of treatment, so most patients need surgery. In 28 cases in our patient cohort (59.6%) presenting as a testicular painless mass, the epididymal-testis border was unclear, or sinus formation was found, suggesting that the surrounding tissues were invaded. Although surgical resection is effective, patients still need regular anti-tuberculosis chemotherapy for 3 to 6 months after surgery and close follow-up.

Conclusion: epididymal tuberculosis is easy to diagnose in patients with a history of tuberculosis. However, in isolated epididymal tuberculosis, early symptoms are not obvious and cases are typically advanced at the time of diagnosis. There are obvious symptoms and signs such as epididymal enlargement, falling pain and bead-like changes. Epididymal tuberculosis has invaded surrounding tissue by the time it is discovered. Surgical resection combined with preoperative and postoperative chemotherapy is an effective treatment approach. When epididymal tuberculosis manifests bead-like changes, testicular involvement, or sinus formation, good results can be achieved with definitive epididymal-testicular resection. This experiment involved a relatively small number of cases, and the patients did not undergo needle biopsy. Therefore, further research is needed to further characterize this disease and refine the treatment.

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