CASE REPORT

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Inflammatory perforated cecal tumor due to intestinal histoplasmosis in an immunocompetent patient

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ABSTRACT

In this case report, the case of a patient who presents with abdominal pain and acute appendicitis signs and symptoms is highlighted. This patient underwent surgery and a cecal tumor with perforations was observed intraoperatively. An oncological surgical approach was decided due to macroscopic evidence. Furthermore, a later histopathological result of intestinal histoplasmosis which is a rare pathology that frequently occurs in immunosuppressed patients was detected. However, in this rare case, the middle-aged male patient does not present immunosuppression problems.

Keywords: Cecal tumor, Immunocompetent patient, Intestinal histoplasmosis

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INTRODUCTION

Histoplasmosis is an infection caused by a fungus called *Histoplasma capsulatum* [1]. Its dissemination to the gastrointestinal tract is infrequent as it occurs in 1 of 5000 cases of histoplasmosis. Even rarer, its presence in patients with preserved immunity reaches 0.05% [2]. Indeed, the most frequent symptoms are diarrhea, abdominal pain, and weight loss [3]. Additionally, the gastrointestinal location is pathophysiologically caused by the large amount of lymphoid tissue in this system. The clinical treatment is Amphotericin B or ketoconazole. However, despite being a specific treatment, relapses have been detected when applied. Noteworthy, surgical treatment is reserved for complications that manifest an acute abdomen due to intestinal perforation.

CASE REPORT

A 24-year-old male patient residing in Sacha (Orellana, eastern Ecuador), without significant pathological history, came to our hospital due to the presence of moderate intensity colic-type abdominal pain that had lasted 14 hours. It began in the mesogastrium and it was later located in the right iliac fossa. In addition, the patient had vomited four times without apparent cause.

Upon admission, vital signs as 130/70 mm Hg blood pressure, 95 beats per minute heart rate and 37 °C temperature were measured. In fact, an abdomen with positive appendicular signs (McBurney, Blumberg, Dunphy) drew attention in the physical examination. Therefore, laboratory examinations were carried out showing 18,200 leukocytosis with neutrophilia (85.5%) and elevated C-reactive protein (CRP) of 6.

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Emergent surgical resolution (appendectomy) was decided. An opened approach was performed with a McBurney-type incision. Afterward, inflammatory fluid of approximately 100 mL was observed. Additionally, the presence of a cecum mass of approximately 4×3 cm was detected. This mass deforms the cecum and retracts it. Indeed, partial necrosis with perforations showing evidence of leakage, gas, and purulent liquid was found (Figure 1).

Given the suspicion of a malignant neoplasm, the applied surgery was a laparotomy where multiple pericolic, epiploic nodes, in the root of the mesentery, the largest up to 2 cm in diameter, were found with no evidence of tumor implants in the liver or peritoneum. Due to the aforementioned cecum mass, a right hemicolectomy and lymph node emptying with low vascular pedicle ligation (right colic) and latero-lateral ileo-transverse anastomosis were performed (Figures 2 and 3). Then, the patient was managed with a postsurgical diagnosis of perforated cecal tumor.

Further exams such as serology tests [enzyme-linked immunosorbent assay (ELISA)], tumor markers cancer embryonic antigen (CEA), and carbohydrate antigen (CA19-9) were negative, without pathological findings in computed tomography of the chest, abdomen, and pelvis.

The patient evolved adequately, and he was discharged on the fourth postoperative day with adequate intestinal transit and good tolerance to the enteral diet.

The pathology report macroscopically revealed a tumor with necrotic areas and microscopically showed intestinal



Figure 1: Necrotic tumor mass in the cecum (shown by arrow).



Figure 2: Surgical piece. Right colon.



Figure 3: Surgical piece. Right colon and cecal tumor (shown by arrow).

mycosis compatible with histoplasmosis, negative for malignancy (Figure 4). An evaluation from the infectious disease service was requested. Furthermore, serology and viral load with western blot for suspected acquired immunodeficiency disease were performed and these analyses were negative. Treatment with Amphotericin B started at a dose of 1 mg per kg of weight per day for one week, continued later with itraconazole 200 mg each day for 12 months. Being the treatment of choice, the patient evolved clinically and surgically in an appropriate way. The patient is currently being followed up by Internal Medicine.

DISCUSSION

Histoplasmosis is a granulomatous disease caused by an intracellular dimorphic fungus called *H. capsulatum*,

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Figure 4: Histopathology: intestinal histoplasmosis: (A) Staining (hematoxylin & eosin, $10 \times$), (B) (hematoxylin & eosin, $40 \times$), and (C) (Giemsa, $40 \times$).

for which soil is the natural habitat [1, 4].

The infection is acquired by inhalation of the microconidia found in the feces of birds and bats. These microconidia reach the pulmonary alveoli where they are phagocytosed by alveolar macrophages and then microconidia are spread to the entire mononuclear phagocytic system. It is the most common endemic mycosis in individuals infected with human immunodeficiency virus [1, 5].

In South American countries, it is endemic to eastern areas [6]. This would be the main risk factor that can be associated with the patients. Histoplasmosis disseminated in the gastrointestinal tract occurs in 1 of 5000 cases. Histoplasmosis has been well documented in the literature within the immunocompromised population. Furthermore, the gastrointestinal condition is rare in a patient who does not have his immunological system compromised. The incidence of which represents 0.05% of the cases. In fact, symptoms are absent in the majority of cases. Therefore, diagnosis is complicated [2, 7].

The presence of disseminated histoplasmosis with gastrointestinal involvement is common in patients with CD4+ cell counts of <200 cells/mm³ [8].

As described, the main symptoms are diarrhea, weight loss, and abdominal pain [3]. Indeed, it is commonly presented as a bowel mass, ulcer, or polypoid lesion in the cecum and terminal ileum [5].

The localization at the gastrointestinal level is a consequence of the large amount of lymphoid tissue present there. Some patients' reports mention its presence as an acute abdomen setting confused with acute appendicitis [3].

Histopathologically, the differential diagnosis of histoplasmosis should include other bacterial and fungal infections. In low magnification, Hematoxylin and Eosin (H&E) preparations, *H. capsulatum* and *Leishmania* are very similar. Consequently, *Toxoplasma* may resemble *Histoplasma* superficially. However, it is smaller, and it is usually not seen within macrophages. Any of the special fungal stains will clarify the difference between *Histoplasma, Leishmania, and Toxoplasma* successfully [7].

At the West China Hospital, Sichuan University, an intestinal study was carried out from October 2005 to March 2015, where four immunocompetent patients were diagnosed endoscopically. These researchers concluded that histoplasmosis presence in immunocompetent patients may exist without being reported since most fungi tests are not routinely performed [8].

The manifestations mimic many other gastrointestinal diseases, such as Crohn's disease, tuberculosis, carcinomas, and lymphomas. Histoplasmosis is not often considered as a cause in the differential diagnosis, which leads to inappropriate therapies [8, 9].

The prognosis for disseminated histoplasmosis is poor: 84% died in a study prior to development of Amphotericin B. Although Amphotericin B kills the organism in vivo, a relapse may occur after therapy completion [8–10].

Ketoconazole is associated with fewer side effects. In 1981, Hawkins reported excellent responses in four patients with disseminated histoplasmosis. Mabey reported a successful treatment of one patient with histoplasmosis who was treated with 200 mg of ketoconazole daily. Slama found that 13 of 14 immunocompetent patients with progressive or disseminated cavitary histoplasmosis were cured with 200 mg of ketoconazole daily [8, 9].

A treatment with 200 mg of itraconazole is taken orally twice a day for three days, and then 1 or 2 times a day for three months are reasonable. However, when considering disseminated histoplasmosis, it is recommended to be applied for 12-24 months [10].

Medical management of gastrointestinal histoplasmosis is recommended and surgical resection in case of urgent cases is reserved for acute emergencies such as intestinal perforation or when required for diagnosis [8].

CONCLUSION

Histoplasmosis is a frequent infection in immunocompetent people that generally does not present symptoms. However, when symptoms appear, treatment is clinically performed administering antifungals. When a patient presents an acute abdomen with signs of intestinal perforation, his treatment is surgical and antifungal treatment is applied for a long time because the infection is widespread.

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Stalin R Ordoñez – Conception of the work, Interpretation of data, Drafting the work, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Bernabé E Aguilar – Design of the work, Revising the work critically for important intellectual content, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Jhony A Delgado – Acquisition of data, Revising the work critically for important intellectual content, Final

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Gabriela A Arboleda – Acquisition of data, Drafting the work, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Victor R Medina – Interpretation of data, Revising the work critically for important intellectual content, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

Andrea E Valladares – Acquisition of data, Analysis of data, Drafting the work, Final approval of the version to be published, Agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved

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Conflict of Interest

Authors declare no conflict of interest.

Data Availability

All relevant data are within the paper and its Supporting Information files.

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