Cytomegalovirus Colitis Mimicking Rectal Tumour in an Undiagnosed HIV Patient

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Keywords
CMV colitis · Rectal tumour · HIV infection · Coinfection

Abstract
Introduction: Cytomegalovirus (CMV) is the most common opportunistic agent in HIV-infected patients. It can affect the entire gastrointestinal tract, but frequently involves the oesophagus and the colon. Case Report: We report the case of a 70-year-old female, ultimately diagnosed with HIV infection, whose inaugural clinical manifestation was CMV colitis with endoscopic findings resembling a rectal tumour in which initial histological evaluation was not able to provide a proper diagnosis. Discussion/Conclusion: Since clinical presentation is variable and histopathological yield without immunohistochemical analysis is poor, recognizing CMV infection in the absence of known risk factors may be difficult. It is crucial to consider this entity with suspicious lesions or when initial evaluation, either clinical or histopathological, is inconclusive, thus avoiding potentially debilitating and superfluous treatment or life-threatening complications.

Palavras Chave
Colite CMV · Tumor rectal · Infecção VIH · Coinfecção

Resumo
Introdução: O citomegalovírus (CMV) é o agente oportunista mais comum nos doentes com infecção pelo vírus da imunodeficiência humana (VIH). Pode afetar todo o trato gastrointestinal, mas frequentemente envolve o esófago e o cólon. Caso Clínico: Os autores apresentam o caso de uma doente do sexo feminino com 70 anos, com diagnóstico inicial de câncer de recto por VIH, cuja manifestação clínica inaugural foi colite a CMV, com achados endoscopicos iniciais mimetizando um tumor rectal. Discussão/Conclusão: Uma vez que a apresentação clínica é variável e a rentabilidade da análise histopatológica sem imunohistoquímica é baixa, o reconhecimento da infecção por CMV na ausência de fatores de risco pode ser difícil. É crucial considerar esta entidade perante lesões suspeitas ou quando a avaliação inicial, clínica ou histopatológica, for inconclusiva, de forma a evitar potenciais terapêuticas debilitantes e/ou superfluas bem como complicações graves.
Introduction

Cytomegalovirus (CMV) is the most common opportunistic agent in HIV (human immunodeficiency virus)-infected patients and in up to 25% of cases it provides the basis for the initial clinical diagnosis of acquired immunodeficiency syndrome [1]. CMV can affect the entire gastrointestinal tract, but frequently involves the oesophagus and the colon [2]. In fact, colitis is the main extraocular manifestation of CMV disease in HIV-infected patients [3]. It is always important to be aware of this entity and diagnose it promptly, as life-threatening complications can occur, such as colonic perforation or gastrointestinal tract bleeding [4]. However, CMV infection diagnosis can be difficult, mainly if no risk factors are suspected (e.g., HIV infection, prolonged immunosuppression) or if its clinical suspicion was not raised for pathologists.

Case Report

Here we present the case of a 70-year-old, melanodermic, female patient who is a native of Cape Verde but has been resident in Portugal since 1997. She had previous history of arterial hypertension, medicated with lisinopril and amlodipine. The patient was referred from the Surgery Department because of an ulcerated rectal tumour with inconclusive histopathology on biopsies obtained from a previous colonoscopy. She also complained of anorexia and weight loss that was not quantified. The physical examination was unremarkable. The blood tests showed a microcytic and hypochromic anaemia (Hb 8.5 g/dL, MCV 72.6 fl, MCH 24.3 pg) and elevation of C-reactive protein (39.6 mg/L). The colonoscopy was repeated, which revealed irregular and ulcerated mucosa, with necrotic areas, between 10 and 18 cm from the anal verge (Fig. 1). On hepatic flexure, we found another ulcerated area involving one third of the luminal circumference (Fig. 2). Based on these findings (two ulcerated segments in the colon, blood test changes, and previous inconclusive biopsies), we requested CMV immunohistochemistry (IHC) on colonic samples, which were positive for its presence (Fig. 3). The patient was prescribed valganciclovir (1 g, 3 times daily) and was tested for HIV infection, which was also positive. When questioned at this stage, the patient confessed that her husband was diagnosed with HIV infection a few months earlier, information that she had not previously disclosed. There was clinical and endoscopic improvement after the valganciclovir therapy (21 days). The patient was referred to an immunodeficiency consultant and started a highly active antiretroviral therapy with a good response.

Discussion

This is a case of HIV infection with an initial clinical presentation of CMV colitis. In fact, the index colonoscopy showed only an ulcerated lesion in the rectum that resembled a malignant tumour. Several cases of CMV colorectal pseudotumors are reported in the literature [1, 5], and it is important to be aware of this condition, mainly in patients with risk factors associated with immunosuppression.

To identify CMV colitis, colonoscopy is preferred over flexible sigmoidoscopy, as more than one third of affected patients have gross disease restricted to proximal regions of the colon. Furthermore, it is also important to
take multiple biopsy specimens as viral inclusions are most often present in the endothelial cells of the deeper layers of the bowel wall [2]. Histopathological examination of colorectal mucosal biopsies, stained with haematoxylin and eosin (HE), is a specific method (92–100%) but often insensitive (10–87%). IHC has been found to be more sensitive than HE staining (78–93%) but is not routinely used – only if there is a clinical suspicion raised by the clinician [6]. Another diagnostic tool is tissue polymerase chain reaction (PCR), which allows for rapid results and compared to IHC is more sensitive, with a high negative predictive value [7]. In fact, the European Crohn’s and Colitis Organisation recommend tissue PCR or IHC as the methods of choice for diagnosing CMV colitis [8].

Concerning treatment, all patients with CMV gastrointestinal disease should receive induction therapy with an antiviral agent for 3–6 weeks or until signs and symptoms have resolved. The options are ganciclovir, foscarnet (both intravenous), or oral valganciclovir (if the patient can absorb and tolerate oral therapy and/or in mild disease). Antiretroviral therapy should be initiated as soon as CMV retinitis has been excluded and the patient can tolerate oral medication. If the patient is already on antiretroviral therapy, this should be continued. Chronic maintenance therapy is not recommended for CMV gastrointestinal disease unless there is concurrent retinitis or recurrent gastrointestinal disease after induction therapy has been discontinued [9].

Thus, this case illustrates the importance of considering CMV infection when we face colorectal lesions that are suspicious or when the initial diagnosis (clinical or pathological) is not conclusive.

Statement of Ethics
The authors declare that the subject gave her written informed consent to publish this case.

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Author Contributions
All authors contributed to the literature review. Manuel Rocha prepared the manuscript. Verónica Borges, Guilherme Simões, Sara Santos, Verónica Gamelas, and Carlos Bernardes revised the manuscript.
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