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Multicentric castleman disease in a HIV patient

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Abstract

A 63 year old newly diagnosed HIV positive male patient had presented with a history of fever and weight loss for 3 months, as well as generalized lymphadenopathy. Inguinal lymph node biopsy revealed features of Castleman disease. Immunohistochemistry revealed +ve staining for EBV and -ve staining for HHV-8. The patient was diagnosed to have idiopathic Multicentric Castleman disease, and treatment with antiretroviral therapy led to complete resolution of lymphadenopathy.

Keywords

HIV; Fever; Disease.

Introduction

A patient afflicted with HIV is at a risk of numerous opportunistic diseases. Generalized lymphadenopathy, a manifestation of HIV itself, has a host of differential diagnoses. Multicentric Castleman disease is a rare lymphoproliferative disorder associated with HIV as reported in this case.

Case Presentation

A 63 year old male patient who had recently been diagnosed with HIV, presented with a history of fever and weight loss of 8 Kgs over 3 months. Examination revealed lymphadenopathy in cervical, axillary and inguinal regions. The baseline HIV viral load was 29,50,000 copies/ml and CD4 was 132 cells/microl (7.85%). VDRL and TPHA were positive. PET CT revealed multiple enlarged lymph nodes (Figure 1).

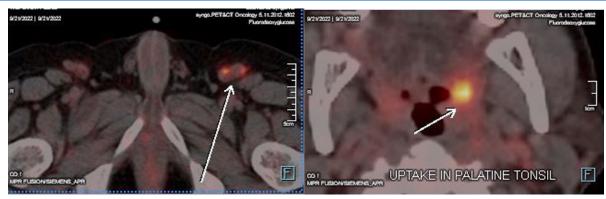


Figure 1: PET CT: FDG avid left inguinal node with an SUV Max of 11.06 (left) and FDG avid asymmetric uptake in left palatine tonsil with an SUV Max of 12.87 (right).

Several differential diagnoses were considered at this point:

- 1. Tuberculosis
- 2. HIV related lymphadenopathy
- 3. Non-tuberculous mycobacterial infection
- 4. Secondary syphilis
- 5. Lymphoproliferative disorder

The patient received Inj Ceftriaxone 2 g IV BD for suspected late latent syphilis. A biopsy was performed from the left inguinal lymph node and relevant investigations were sent for the above differential diagnoses. Histopathological examination revealed interfollicular plasmacytosis, atretic lymphoid follicles with penetrating vessels, interfollicular zones with prominent proliferation of high endothelial venules and focal endothelial necrosis/apoptosis. All these features were suggestive of Castleman Disease. An immunohistochemistry was performed, which was negative for HHV-8 and positive for EBV. Figures 2, 3 and 4 describe the histopathological features in detail.

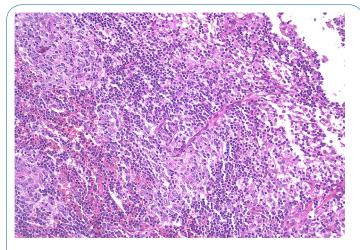


Figure 2: Atretic lymphoid follicles with penetrating vessels (Low power magnification).

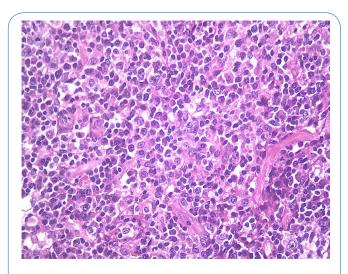


Figure 3: Increased number of plasma cells (High power magnification).

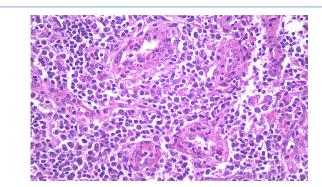


Figure 4: Interfollicular zone with prominent proliferation of high endothelial venules and focal endothelial necrosis/ apoptosis (High power magnification).

Hence, he was diagnosed to have EBV +ve, HHV8 -ve, idiopathic Multicentric Castleman disease. The patient was started on antiretroviral therapy (Tenofovir alafenamide fumarate 25 mg + Emtricitabine 200 mg + Dolutegravir 50 mg) after diagnosis. No chemotherapy was used. After 6 months, he had complete resolution of symptoms, reduction in HIV viral load to 408 copies/ml, improvement in CD4 to 204 cells/microl (10%) and PET CT showed a complete radiologic and metabolic resolution of the lesions (Figure 5).

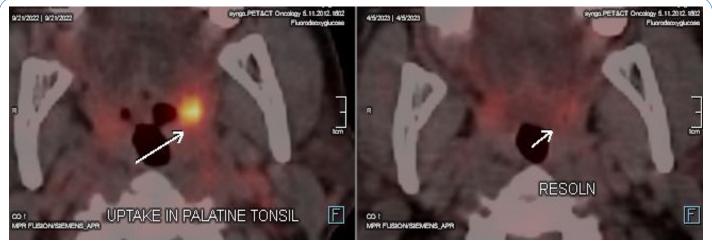


Figure 5: Complete resolution of enlarged left inguinal lymph node (top) and palatine tonsil (bottom). Left side of the image shows the old PET CET and right side shows the new PET CT, after 6 months.

Discussion

Castleman Disease (CD) includes a heterogenous group of lymphoproliferative disorders that share common histopathological features. Multicentric Castleman Disease (MCD) can be HHV-8 or KSV (Kaposi Sarcoma Virus) +ve, which is the most common MCD associated with HIV, or HHV-8 –ve / idiopathic MCD (iMCD). Disease in Indian patients may be HHV-8 –ve, but +ve for EBV [1], as seen in this patient and may have a role in causation. The primary treatment for MCD is chemotherapy. HHV-8 +ve patients are treated with Rituximab, while HHV-8 –ve patients are treated with Siltuximab as the first line chemotherapy agents [2]. Considering the high risk of infectious complications following cytotoxic therapy in the AIDS population, patients with AIDS-related MCD may be first managed with antiretroviral therapy, leaving more aggressive therapeutic approaches for patients if relapse occurs [3]. There are multiple case reports of HIV associated MCD being treated with just antiretroviral therapy, without chemotherapy [4,5], as we did in this patient, and led to complete resolution of signs and symptoms.

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