Spontaneous resolution of persistent lymphadenitis: a case of Kikuchi–Fujimoto disease
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Cervical lymphadenopathy is common in all age groups. Persistently enlarged cervical lymph nodes often pose a diagnostic challenge and necessitate focused clinical evaluation with targeted investigations. Pathological examination of excised node yields conclusive answer in the vast majority of cases with unsettled diagnosis. We present a case of a young man with persistent posterior cervical lymphadenopathy which on excision biopsy turned out to be Kikuchi–Fujimoto disease. With watchful follow-up, he had a self-limiting clinical course in the next few months.

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Introduction
Persistent and enlarged cervical lymph nodes more than 6 weeks can be owing to infective, malignant, and immunologic etiologies. A proper and detailed systemic examination and laboratory and histopathological evaluation may be necessary to establish a correct diagnosis in a case of persistent adenopathy. Kikuchi–Fujimoto disease (KFD), an entity of unknown etiology, is an uncommon cause of cervical lymphadenopathy in young adults, which usually has a self-limiting clinical course. The histopathological changes in the surgically resected nodes are characteristic enough to yield a conclusive diagnosis. Herein, we aim to provide a brief presentation of a patient with persistent cervical lymphadenopathy who turned out to be having KFD on histopathological analysis.

Case summary
A 24-year-old male student, resident of Ernakulam urban area, with atopic background presented with a history of fever, cough, and swelling in the neck for 6 weeks. His history was remarkable for intermittent sneezing, rhinorrhea, and wheeze associated with respiratory infections. There was history of loss of weight of 4 kg in 6 months without any alteration of bowel and bladder habits. He had no addictions. He was a chemistry student and had history of exposure to chemical fumes in the laboratory. He denied any history of contact with patients with tuberculosis or sexual exposure. Physical examination revealed multiple firm left-sided posterior cervical lymph nodes of varying size, the largest being 2×2 cm in size. The nodes were nontender, discrete, and mobile. There was no inflammation of the overlying skin.

Respiratory system examination finding was normal. Abdomen examination revealed no clinically appreciable enlargement of liver or spleen. No lymph node enlargement at any other site was appreciated. Skin and joint examination was essentially unremarkable.

Blood examination revealed a normal hemoglobin and leukocyte count (hemoglobin 13.5 g/dl, total cholesterol 7600 mm3). Erythrocyte sedimentation rate at the end of first hour was 58 mm. An ELISA test result for HIV infection was negative. Mantoux test was nonreactive with no induration. Serum low-density lipoprotein level was within normal range. Chest radiograph (Fig. 1) and ultrasonography of the abdomen were unremarkable. With the clinical and investigation reports, the diagnosis remained elusive. The differentials entertained were chronic suppurative lymphadenitis, tuberculous lymphadenitis, lymphoma, autoimmune diseases, etc.

As the diagnosis was not settled with the available investigations, an excision biopsy of the left cervical lymph node was done and subjected to histopathological examination. Microbial tests including bacterial and fungal culture and gene Xpert for mycobacterium tuberculosis were also sent. Biopsy sample for histopathology examination revealed histiocytic infiltration of the node with confluent necrosis, karyorrhexic nuclear debris, and pyknosis.

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characteristic of KFD (Fig. 2). Microbial tests returned negative. Virology tests to exclude EBV and cytomegalovirus involvement were not done owing to nonaffordability. HIV card test was nonreactive. Tests for autoimmune diseases were also asked for (ANA screening by immunofluorescence, anti-ds DNA), which were negative. He was initiated on symptomatic measures (antipyretics and supportive measures) and was kept on close follow-up. He became afbrile in 2 weeks with total resolution of lymph nodes in 5 months.

Discussion
KFD is also known as ‘histiocytic necrotizing lymphadenitis.’ It was first described in Japan in 1972 by Kikuchi and Fujimoto independently [1]. It is relatively common among people of Asian origin. The etiology is largely unknown, although viral infections and immunological factors has been postulated. HHV-6, HHV-8, HTLV-1, EBV, and cytomegalovirus are postulated to be associated with KFD [2].

Affected patients are typically young, with a mean age of diagnosis of 21 years. Female predisposition has been noted with a female to male ratio of 4 : 1. Lymphadenopathy usually involves one or more of posterior cervical lymph node group [3]. The disease is often associated with fever, myalgias, arthralgias, and weight loss. Diarrhea, chills, and sweating are uncommon, and rarely, hepatosplenomegaly may be seen [4].

Systemic lupus is a mimicker of KFD, and both of them share similar histologic features, which warrants detailed evaluation to differentiate between them [5]. KFD is essentially a histological diagnosis characterized by paracortical necrosis, histiocytes (crescentic nuclei), and karyorrhexis. The clinical and immunological features required for diagnosis of systemic lupus erythematosus (SLE) are well documented and specific. Lupus lymphadenitis has been reported in between 12% and 59% of patients with SLE, but in contrast to KFD, is rarely the presenting feature. ANA is usually negative in KFD.

Other differential diagnosis include lymphoid malignancies like non-Hodgkin lymphomas and reactive lymphadenopathy owing to infectious etiologies, such as EBV, herpes simplex virus, and toxoplasmosis. As their management protocols differ from KFD, these entities need to be excluded before a final diagnosis of KFD is committed. Patients with KFD should be followed up regularly for years to detect possible evolution of SLE [6] and lymphoma [7].

Although the natural history can occasionally be unpredictable regarding severity and complications, most cases are self-limiting, improving within 6 months. Lymph node enlargement usually subsides in few weeks to maximum 6 months. Treatment is mainly symptomatic and supportive. NSAIDs and corticosteroids can be used to reduce inflammation depending on severity. Hydroxychloroquine has also shown some benefit in treating the disease, and antibiotics are not found to be beneficial [8]. There is a 3% risk of recurrence. Recurrences usually occur within a few weeks of the first episode [7]. Death from Kikuchi disease is extremely rare (2.1%) and usually
occurs owing to liver, pulmonary hemorrhage, DIC, fatal hemophagocytic syndrome, or heart failure [4].

**Conclusion**

To conclude, KFD should be considered in the differential diagnosis of any patient presenting with unexplained lymphadenopathy with or without constitutional features. Reaching an appropriate diagnosis is especially important as it can be mistaken for complicated diagnosis like SLE, tuberculosis, or even lymphoma. Even though self-limiting illness, follow-up should be observed to detect recurrence or associated conditions such as SLE and lymphoma.

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**Conflicts of interest**

There are no conflicts of interest.

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