Serum sickness classically refers to a systemic immunological reaction caused by injected serum and occurs due to deposition of an antigen-antibody-complement complex. It occurs when an antigen (i.e., serum) persists in the circulation for a long duration stimulating IgG or IgM antibody synthesis. These antibodies, along with complement, form complexes which are deposited in various organs. The manifestations develop according to the organ involved. Since heterologous serum is rarely used, such a systemic reaction due to serum is rare. Other causes like drugs are now more commonly implicated and the resultant reaction is called a serum sickness like reaction. Minocycline can cause such a reaction rarely.\(^{2,3}\)

Our patient had all the features of serum sickness like reaction, namely angioedema, lymphadenopathy, arthralgia and fever. It is interesting to note that like most previous reports our patient was also female. Although there may be hematological alterations like hypocomplementemia or signs of renal impairment, their absence does not exclude this diagnosis and a combination of clinical signs is sufficient for the diagnosis of SSLR. Such a reaction should be recognized at the earliest and the drug should be stopped. We hope this report helps to increase awareness about this rare adverse effect of minocycline, which is being used increasingly commonly for the treatment of acne in India.

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Nimesulide induced bullous fixed drug eruption of the labial mucosa

Sir,
A fixed drug eruption (FDE) is a distinct drug induced reaction pattern that characteristically recurs at the same skin or mucosal site. We report a case of bullous FDE confined to the labial mucosa following intake of nimesulide, a commonly used nonsteroidal anti-inflammatory drug in India.

A 23-year-old man presented with multiple oral blisters of one day’s duration associated with marked burning and soreness. He had developed the lesions within 5 to 6 hours of taking a single tablet of nimesulide (which he had taken by himself). He had also been taking roxithromycin for three days which had been prescribed for his sore throat by an otolaryngologist. In the past he had taken multiple courses of roxithromycin for his recurrent sore throat without any complaints, but this was the first time that he had taken nimesulide.

The patient was afebrile and there were no other associated systemic complaints. Examination revealed multiple, tense bullae filled with clear fluid, varying in size from 0.5-1.5 cm, distributed over the upper labial mucosa with slight surrounding erythema (Figure 1). The rest of the oral mucosa and other mucosal sites were free of lesions.

A biopsy from the bullae was taken. Histopathological

Figure 1: Bullous FDE lesions involving the labial mucosa
examination revealed necrotic keratinocytes and a marked inflammatory infiltrate in the dermis, findings consistent with a fixed drug eruption.

The patient was asked to stop both the drugs and to avoid nimesulide in the future. He was given a short course of steroids in a tapering dose for two weeks following which the lesions improved. The lesions healed with mucosal hyperpigmentation.

Brocq first introduced the term fixed drug eruption in 1894. The pathogenetic mechanism underlying FDE is still enigmatic. The most commonly accepted hypothesis is persistence of memory T cells in the affected skin. CD8+ cells phenotypically resembling effector memory T cells have been shown to be greatly enhanced in the lesions of FDE. The lesions of FDE usually start as an erythematous macule that subsequently evolves into a plaque. Vesicles and bullae develop at a later stage and are usually hemorrhagic. The lesions can occur on any part of the skin and mucous membranes. The sites of predilection are the limbs, sacral region, genitalia, palmar and plantar skin. The oral mucosa may be involved in association with skin lesions or alone.

The drugs that commonly cause mucosal FDE include co-trimoxazole, oxyphenbutazone, and tetracycline. Classically the transitional epithelium of the mucocutaneous junctions is involved but strict localization of the lesion to the labial mucosa, as was seen in our patient, is unusual. Systemic manifestations are uncommon and include fever, malaise, nausea, diarrhea, abdominal pain, urethritis, and conjunctivitis.

Nimesulide is a nonsteroidal anti-inflammatory agent with antipyretic and analgesic properties. It is being commonly prescribed in India. Some of the side effects reported with its use are pruritus, urticaria, purpura, maculopapular rash and localized toxic pustuloderma. Due to severe hepatotoxicity and hemolytic anemia associated with its use, nimesulide is likely to be withdrawn from the market in many countries.

To the best of our knowledge only 8 cases of FDE secondary to nimesulide have been reported and there is only one other report with primarily oral mucosal involvement. This report emphasizes an uncommon mucosal localization of bullous FDE due to nimesulide.

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