Normocomplementemic Urticarial Vasculitis Associated with A/H1N1 in a Child. Case Report

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Abstract
Urticarial vasculitis is an entity characterized by urticarial rashes that are typically pruritic and painful, associated with angioedema and purpura. Influenza viruses are common respiratory pathogens that can cause seasonal infections and global human morbidity/mortality. A 4.5-year-old girl presented with fever and painful urticarial skin lesions leaving post-inflammatory hyperpigmentation. The nasal specimens showed a positive for influenza A/H1N1. On the basis of these findings, a diagnosis of urticarial vasculitis associated with A/H1N1 was concluded. Taking the results together, we suggest that urticarial vasculitic lesions can be considered to the list of cutaneous manifestations during the seasonal flu.

Keywords Child · Vasculitis · Urticarial vasculitis · Influenza infection

Introduction

Generally, influenza infection is an acute, self-limited, and uncomplicated disease that is caused by influenza A and B, and occurs every winter season [1]. Abrupt onset of fever, headache, myalgia, cough, sore throat, and rhinitis is mentioned as frequent clinical manifestations of uncomplicated influenza infections [1, 2]. Cutaneous involvement with both influenza infection and vaccination has been reported as a macular/maculopapular rash or vasculitic/purpuric eruption [3]. Urticarial vasculitis (UV) is a type of leukocytoclastic vasculitis that may present with a generalized urticarial-appearing eruption, fever, and arthralgias. This is a clinicopathologic entity thought to be mediated by a Type-III hypersensitivity reaction mediated by antigen-antibody complexes deposited on vascular endothelium creating inflammation and vasculitis. Urticarial lesions are typically fixed for more than 24 h and may be associated with petechiae, purpura, or post-inflammatory hyperpigmentation. The prevalence is unknown and occurs in middle-aged women being rare in children [3, 4]. Leukocytoclastic vasculitis following seasonal influenza vaccination has been reported in several cases [5]. However, urticarial vasculitis associated with A/H1N1 infection has not been reported previously. This case describes a 4.5-year-old girl who presented with fever and painful urticarial skin lesions, and diagnosed with A/H1N1 associated with urticarial vasculitis.

Case Report

A previously healthy 4.5-year-old girl was admitted to our outpatient department with a 6-day history of nonproductive cough, nasal congestion, myalgia, and fever with a temperature of 39 °C twice a day during the period of seasonal flu. After a 3-day history of flu symptoms, urticarial lesions were noticed on various parts of the body mostly the face and upper extremities, accompanied by arthralgia, edema, and pain on hands and wrist. It was found out that the patient was diagnosed with “urticaria” and was treated with antihistaminic treatment in another hospital 3 days before. In the last 2 days,
on resolution left a violaceous-brownish purpuric lesion. Some urticarial rashes subsided, leaving post-inflammatory hyperpigmentation. After 2 weeks, the entity was totally resolved without scarring or residual hyperpigmentation. In the following 2 months, no new active skin lesions were detected.

Discussion

This is an unusual presentation of H1N1 influenza A infection presenting with findings of urticarial vasculitis. The clinical reports published to date have not demonstrated this pattern of cutaneous manifestation; however, it was likely secondary to an influenza vaccine. Dermatological manifestations due to influenza infections and influenza vaccines include a variety of conditions such as a rash (macular-maculopapular), purpura, and petechiae. There have been reported that petechial eruptions were seen in 3–10% of children infected with pandemic H1N1 influenza [6]. However, vasculitic-purpuric syndromes are an extremely rare manifestation in influenza infection. Silva ME et al. [5] described a case of a child with a febrile petechial rash associated with influenza A virus infection. Rocco Urso et al. [6] have reported a 23-year-old Caucasian woman was hospitalized for fever, severe abdominal pain, hematuria, arthritis, and purpuric manifestations associated with pandemic 2009 (H1N1) virus by reverse transcriptase PCR assay. A report conducted in the USA showed a case with vaso-occlusive retinal vasculitis following H1N1 infections [7]. Regarding the systematic review of current literature for influenza-associated vasculitis in children, we searched for articles published up to 2010 in the following databases: PubMed, Medline, Embase, Cochrane Library, and CINAHL. We used search terms including “influenza,” “childhood,” “vasculitis,” and “urticarial vasculitis” to identify reports that presented data on influenza-associated vasculitis. To our knowledge, there are no reports concerning cases of urticarial vasculitis associated with the A/H1N1 infection.

Urticarial vasculitis, first described by McDuffie in 1973, is a clinicopathologic entity characterized by urticarial rashes that are typically pruritic and painful, associated with angioedema and purpura [8]. The urticarial plaques of UV persist for more than 24 h, can have a central dark red or brown macule, and may resolve with minimal residual hyperpigmentation. Typical UV criteria are urticarial lesions lasting more than 24 h and 2 of 3 of the following: pain or tenderness, purpura, or resolution with hyperpigmentation. The histopathological spectrum of urticarial vasculitis shows minimal vascular damage to severe leukocytoclastic vasculitis caused by endothelial cell damage, fibrin deposition, and acellular infiltrate of neutrophils with or without leukocytoclastic and lymphocytes. The pathophysiology of UV is thought to be a formation of immune complexes in the blood that then deposits in vessel walls due to third-type hypersensitivity reaction against any of antigens [9].
For patients evaluated because of the novel 2009 H1N1 influenza infection, the incidence of rash is about 2% typically macular or maculopapular in children [10]. Ryan-Poirier reported that macular, maculopapular, and petechial rashes appear in a few counts of children with influenza virus infections. The classic presentations of influenza A virus infection consist the sudden onset of fever, chills, myalgia, cough, sore throat, rhinorrhea, croup, bronchitis, bronchiolitis, and pneumonitis [11]. Despite the documentation of UV following influenza vaccination, no report has described an association of UV with influenza infection.

In conclusion, we have reported the first case of UV associated with influenza A virus infection diagnosed by clinical features and proven by skin biopsy, treated with oseltamivir, antihistaminic drugs, and prednisolone. Influenza virus can also sometimes be associated with dermatologic findings, and therefore, such associations should be kept in mind during seasonal influenza infection.

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Compliance with Ethical Standards

Conflicts of Interest The authors declare that they have no conflict of interest

Human and Animal Rights and Informed Consent This article does not contain any studies with human or animal subjects performed by any of the authors. Informed consent was taken from parents.

Consent to Participate Informed consent was taken from parents.

Consent for Publication The manuscript has not been submitted or accepted elsewhere. If accepted, the paper will not be published elsewhere in the same form, in English or in any other language, without written consent of the copyright holder.

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