A Case of Concurrent Acute Viral Myocarditis and Intussusceptions in a 3-Year-Old Child

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Chickenpox is an acute disease caused by the varicella-zoster virus (VZV), a herpesvirus that causes human infection worldwide. Primary VZV infection routinely occurs during childhood and is usually a self-limiting illness in immunocompetent children. However, chickenpox can be a severe disease in adolescents, adults, and immunosuppressed or immunocompromised patients. Although vaccination substantially attenuates disease manifestations, significant complications such as secondary soft tissue infection, encephalitis, and pneumonia can occur. We present a rare autopsy case of concurrent acute myocarditis and intussusceptions in a 3-year-old female child who presented with chickenpox followed by abdominal pain and sudden death. The present case emphasizes the potential for fatal complications of viral infections, which should be considered in cases of sudden unexpected infectious death in children.

Key Words: Chickenpox; Intussusception; Myocarditis; Sudden death; Viral disease

Introduction

Varicella-zoster virus (VZV) is a herpesvirus that causes two clinically distinct forms of disease, varicella (chickenpox) and herpes zoster (shingles). Primary VZV infection results in characteristic diffuse vesicular rash of chickenpox. Although vaccination substantially attenuates disease manifestations, significant complications such as secondary soft tissue infection, encephalitis, and pneumonia can occur. The specific types of complications among patients with chickenpox also tend to vary by age. However, the most frequent chickenpox-related complications in children...
include secondary bacterial infections and pneumonia.

Intussusception is the most common cause of intestinal obstruction in young children, and the majority of cases occur in those less than 24 months of age [1,2]. The exact pathogenesis and underlying causes of intussusception in children remain unknown, except for several pathogens, including adenovirus. A previous study suggested an association between viral infection and development of intussusceptions in young children [3]. In addition, myocarditis and consequent damage due to inflammation of the heart may result in myocardial dysfunction and heart failure. Myocarditis causes are diverse, including infection, toxin, and autoimmunity. The most common cause of the infectious etiology is viral infection. We report an autopsy case of a 3-year-old female child with sudden unexpected death who presented with intussusceptions and acute myocarditis.

**Case Report**

The 3-year-old female child weighed 14 kg and measured 91 cm. She was not diagnosed with serious disease, malnutrition, or anatomical malformations before her death. Five days before her death, she presented with mild fever, itching sensation, and characteristic diffuse vesicle formations on her arms, legs, and face. She visited the pediatric clinic, was diagnosed with chickenpox, and was prescribed anti-viral drugs and antihistamines. Three days later, the dermatologic manifestation had subsided, but her fever persisted; a cough also developed. A physical exam revealed no pathologic signs in her respiratory tract. The next day, she presented with newly developed symptoms of abdominal pain and dehydration due to vomiting. Although the dehydration was corrected with intravenous administration of fluids, her abdominal pain remained intermittent. Twenty-four hours later, she unexpectedly collapsed at her home and was dead on arriving at the hospital.

At autopsy, external examination revealed no structural abnormalities, anatomical malformation, or physical damage. Internal examination revealed multiple areas of gray-whitish lesions in serial sections of the myocardium (Fig. 1). Microscopic findings of the myocardium showed inflammatory infiltration with interstitial edema, degeneration of adjacent cardiomyocytes, and focal necrosis. Most of the infiltrates were mononuclear lymphocytes, but occasional neutrophils were also observed (Fig. 2A). Immunohistochemical staining showed most of the mononuclear lymphocytes to be CD3-positive T lymphocytes (Fig. 2B, C). Areas of intussusception were found at three distinct

![Fig. 1. Gross autopsy findings of the heart (A) with multiple areas of gray-whitish lesions on the cut surface after formalin fixation (B).](http://www.kjlm.or.kr)
Fig. 2. (A) Microscopic findings of inflammatory infiltration of the myocardium. The infiltrates are mostly lymphocytes (H&E, ×400). Immunohistochemical staining of inflammatory cells with CD3 (B) and CD20 (C) (×400).

Fig. 3. Gross autopsy findings of three areas of intestinal intussusceptions in the small intestine without visible peritonitis (A), and one of the opened intussusceptions (lesions are indicated by arrows) (B).
sites about 123 cm, 150 cm, and 195 cm from the pyloric sphincter (Fig. 3). The intussusceptions measured 1 cm, 2.5 cm, and 1.5 cm. Gross autopsy and microscopic examination revealed no evidence of peritonitis. No poisonous agent, drug, or chemical was detected in the blood and gastric contents. Bacterial culture of post-mortem blood was negative.

**Discussion**

Our patient’s illness began five days before her death as typical chickenpox with symptoms of fever and myalgia that developed into abdominal pain, vomiting, and sudden death. The intussusceptions could have been a constellation of the death process. Proof of death due to intussusceptions requires (1) confirmation of vomitus in the airway and (2) compatible events and symptoms prior to death [4]. However, on autopsy, her airway was clear of vomitus and the patient only had a history of dehydration. Therefore, additional investigation was required for better assessment of the cause of death other than the observed intussusceptions.

The other possibility was acute myocarditis. The clinical presentation of myocarditis is usually variable. Affected patients typically present with a broad spectrum of signs and symptoms ranging from subclinical to cardiogenic shock, arrhythmia, and even sudden death. Patients may have prodromal symptoms of fever, myalgia, and malaise prior to onset of heart dysfunction. However, nonspecific signs and symptoms such as respiratory distress, anorexia, abdominal pain, and vomiting may be the most prominent presenting features, and might lead to an incorrect initial diagnosis in many children [5–7]. In this context of potentially affected patients presenting with serious complications such as cardiogenic shock and sudden death, it is possible that acute myocarditis caused death in this case, based on the clinical history of prodromal symptoms as well as the histologic findings.

Our unique autopsy case demonstrates synchronous intestinal intussusceptions and acute myocarditis in a female child after chickenpox. Intussusceptions are the most common cause of intestinal obstruction in young children; the majority occur in children less than 24 months of age [1,2]. The exact pathogenesis and underlying causes of intussusception in children remain unknown, except for several known viral infections, including enteric adenoviruses. Previous studies suggest a strong association between viral infection and development of intussusceptions in young children [3]. However, myocarditis results from inflammation of the heart muscle; the consequent damage results in myocardial dysfunction and may ultimately lead to heart failure. Its causes are diverse, including infection, toxin, and autoimmunity. The most common cause of the infectious etiology is viral infection, including Coxsackie B virus, adenovirus, parvovirus B19, Epstein-Barr virus, cytomegalovirus, and human herpes virus. In accordance with the results of previous studies, Morentin et al. [8] reported that the main cause of sudden unexpected death from infectious disease was myocarditis (35.7%) followed by bronchopneumonia (32.1%). Therefore, the concurrent viral etiology of intussusceptions and myocarditis in this child might explain the presence of two distinct diseases in autopsy.

The present case emphasizes the possibility of fatal complications, which should be considered especially in cases of sudden unexpected infectious death of a child. Although our case is limited by the absence of serological or virological data, chickenpox is usually a clinical diagnosis. No further diagnostic testing is necessary in cases with typical clinical presentation. In addition, virus serology tests are not recommended for diagnosis of myocardial infection in patients with suspected myocarditis. Microbiological confirmation is sought only in cases with atypical features. Furthermore, development of skin lesions typical of chickenpox makes an alternative diagnosis highly unlikely.

In conclusion, we present a case of a 3-year-old female child with intussusceptions and acute myocarditis possibly precipitated by chickenpox.
Although sudden unexpected death in children and young people has been scarcely studied, a significant proportion (16%) of all sudden deaths and a considerable percentage (5%) of mortality from infectious diseases affect children and young people [8]. Traditionally, microbiological post-mortem investigation has been underestimated. Most recently, however, genomics and proteomics have been proposed for investigation of the potential role of bacterial toxins in some cases of sudden death [9]. The only limitation of the present case is that, due to the routine autopsy process and protocol, microbiological analyses could not be performed except for microbiological culture. Therefore, this article emphasizes the need for a specific forensic procedure if there is suspicion of sudden unexpected death from infectious disease, including detailed rules on sample collection and autopsy protocols including polymerase chain reaction amplification and genetic sequencing, particularly in the context of negative microbiological cultures [8,10].

Conflicts of Interest
No potential conflict of interest relevant to this article was reported.

References