COVID Toes: Digital Vascular Changes in Patients with a COVID-19 Infection

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Received: 12 May 2020; Accepted: 28 May 2020

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ABSTRACT

COVID-19 is a highly infectious disease caused by the SARS-CoV2 virus and has resulted in a global pandemic. Primarily understood as a respiratory disease, ongoing studies have observed the effects of COVID-19 on other body systems. COVID toes, also known as pseudo-chilblains, is an increasingly common dermatological condition in which pruritic, erythematous rashes appear on the toes. Generally benign in adults, COVID toes have also been observed in Multisystem Inflammatory Syndrome in Children (MIS-C) associated with COVID-19. Here we present a case of a 9-year-old Hispanic asymptomatic COVID-19 male and a 53-year-old African American female with symptomatic, confirmed COVID-19, with a comparison of the prognostic differences between children and adults of COVID toes and a review of the current literature.

Keywords: COVID Toes, COVID-19, Pseudo-Chilblains, Pandemic, Pernio

Introduction

COVID-19 is a highly infectious viral illness caused by the severe acute respiratory syndrome coronavirus 2 or SARS-CoV2 virus (Ahn *et al.*, 2020). As a coronavirus, COVID-19 attacks alveolar cells in the respiratory system causing pneumonitis (Hamming *et al.*, 2004). However, this is a tremendous oversimplification, as COVID-19 in select individuals appears to trigger a variety of immune phenomena resulting in multi-system damage (Huang *et al.*, 2020). Colloquially known as the coronavirus, COVID-19 has evolved from a local outbreak to a global pandemic, resulting in over 4 million infections and 300 thousand deaths worldwide (Lai *et al.*, 2020). Due to its high pathogenicity and transmissibility, largescale changes to social interactions have altered the fabric of human interactions through social distancing measures, widespread shutdown of economies, closure of schools and non-essential businesses, and the curtailment of travel and large gatherings (Matrajt and Leung, 2020; Prem *et al.*,

2020). Current knowledge of COVID-19's clinical presentation and treatment options have been rapidly evolving, with ongoing research revealing new discoveries on a daily basis.

Recent studies have highlighted new dermatological findings related to COVID-19 from assorted constellations of benign rashes in adults to a more deadly Multisystem Inflammatory Syndrome in Children (MIS-C) associated with COVID-19 (Andina *et al.*, 2020; Belhadjer *et al.*, 2020; Galván Casas *et al.*, 2020; Nirenberg and Del Mar Ruiz Herrera, 2020). COVID toes, a type of benign pseudo-chilblains, were one of the first rashes ascribed to COVID-19, however its significance as a diagnostic tool or prognosticator for recovery are still largely being investigated (Cordoro *et al.*, 2020; Zhang *et al.*, 2020). With a spectrum of physical manifestations, consistent diagnosis of COVID toes may pose a challenge for physicians and patients alike, especially in the context of asymptomatic COVID-19 (Galván Casas *et al.*, 2020).

Here we present a case of a 9-year-old Hispanic asymptomatic COVID-19 male and a 53-year-old African American female with symptomatic, confirmed COVID-19, with a comparison of the prognostic differences between children and adults with COVID toes and a review of the current literature.

Case Presentations

Case 1

A 53-year-old African American physician female presented to an allergic clinic for recent onset skin lesions of the bilateral ten toes. Her past medical history included hypertension, hyperlipidemia, allergic rhinitis, and allergies to latex and betadine. Her social history was remarkable for recent exposure to COVID-19 through her husband who was hospitalized 1-month prior for COVID-19 positive pneumonia. At the time of exposure, she rapidly developed flu-like symptoms of myalgia, fatigue, non-productive cough, and subjective low-grade fever within a week. She was seen at an urgent care facility where her vitals revealed 98% oxygen saturation on room air, blood pressure of 130/60 mmHg, and pulse of 70 beats per minute. Due to the evolving local, state, and federal guidelines, she did not meet COVID-19 diagnostic testing criteria at that time and was given a course of Azithromycin. Her cough improved, however she developed tachypnea and dyspnea on exertion, with persistence of her malaise and fever. Due to her lack of improvement, a chest x-ray was ordered which revealed a left lower lobe pneumonia. The patient was managed on outpatient cefdinir and albuterol. Nasopharyngeal COVID-19 swab and PCR analysis were performed and returned positive, confirming diagnosis of COVID-19 infection. Four-weeks after the onset of her respiratory symptoms, the patient noticed bilateral blue/grey discoloration of her proximal and distal foot phalanges, worse on the first and second digits, but no pain,

pruritis, or loss of sensation (Fig. 1 and Fig. 2). 10 weeks after her initial COVID-19 diagnosis, her toe discoloration has resolved without any topical or oral treatment. (Fig. 3)



Figure 1: Picture of toes 7 weeks post-onset of COVID-19 symptoms.



Figure 2: Picture of toes 9 weeks post-onset of COVID-19 symptoms.



Figure 3: Picture of toes 10 weeks post-onset of COVID-19 symptoms, with subjective resolution of discoloration.

Case 2

A 9-year-old half Hispanic, half Asian male whose parents were both emergency medical technician first responders (both having tested COVID-19 antibody positive), presented to telemedicine dermatology clinic with a 3-week history of tender nodules of the toes. Past medical history was significant for attention deficit disorder, managed on methylphenidate, and asthma, well-managed on Beclomethasone

and albuterol inhalers. Social history is significant for daily walks in cold weather during quarantine orders from March to April 2020. He was feeling well, afebrile, no malaise, asthenia or myalgias. For two weeks his right second toe, one week the 4th toe on the right, and then the left foot second toe developed warm, tender erythematous nodules (Fig. 4). Initially severely tender and painful, the lesions caused difficulty with walking and weight bearing. He was treated with oral cephalexin (40 mg/kg/day) for 10 days and the patient has had slow and steady improvement. The child was well managed with topical mometasone, local warming, and acetaminophen for pain. The patient was fully resolved without sequelae 4 weeks later.



Figure 4: Picture of toes at start of symptoms, taken by patient's parent via telemedicine.

Discussion

thousand deaths in the United States alone (Aggarwal *et al.*, 2020). Age over 60, male gender, and racial minorities such as African Americans and Latino/Hispanic have been disproportionately affected by COVID-19, with higher rates of infection, hospitalization, and mortality (Department of H, Mental Hygiene C-RT, 2020; Yuan *et al.*, 2020; CDC Covid Response Team, 2020). During its early stages, skin manifestations of COVID-19 were not widely recognized due to the focus on the more life-threatening disease processes of acute respiratory distress and pneumonia. Rapidly developing knowledge regarding asymptomatic carriers, comorbidities, and treatment have fueled varied COVID-19 approaches to containment and preventative measures (Adhikari *et al.*, 2020). From the pandemic onset, new COVID-19 associated symptoms have been documented across various organ systems such as anosmia, ageusia, delirium, and diarrhea (Jin *et al.*, 2020; Vaira *et al.*, 2020). Dermatologically, COVID-19 mimics other viral exanthems, with reports demonstrating a spectrum that include, but are not limited to pseudo-chilblains, vesicular (varicella-like) eruptions, maculopapular rashes, urticarial lesions, and skin necrosis (Huang *et al.*, 2020; Galván Casas *et al.*, 2020; Cordoro *et al.*, 2020). Due to the low sensitivity and scarcity of diagnostic testing during the early COVID-19 pandemic, various studies have accepted clinical diagnoses

of COVID-19, further complicating attempts to correlate laboratory results with COVID skin manifestations (Zainol Rashid *et al.*, 2020). In a recent study, 41% of patients with COVID toes had a confirmed COVID-19 infection and serology, however patients with convincing physical exam and history of exposure have also tested negative for SARS-CoV2 RNA, which further obscures the role of testing in supporting diagnosis and quarantine recommendations (Galván Casas *et al.*, 2020).

Despite these barriers, COVID toes or pseudo-chilblains, have recently come into focus in which patients have detailed painful, pernio-like acral lesions that are raised and discolored in patients of all ages (Nirenberg and Del Mar Ruiz Herrera, 2020; Cordoro et al., 2020). Typically emerging within a week of other COVID-19 symptoms, the etiology and subsequent pathophysiology of COVID toes are unknown and unlike classical chilblains, in which lesions occur due to small vessel inflammation in response to cold temperature or vitamin D deficiency (Piccolo et al., 2020; Shahi et al., 2015). The frequency of COVID toes is underestimated as many of the children with these features do not have other features of COVID-19 (Andina et al., 2020; Molloy and Bearer, 2020). Unclear reporting guidelines and underdiagnosis have estimates between 2% to upwards of 20% in all COVID-19 patients for COVID toes (Galván Casas et al., 2020; Cordoro et al., 2020; CDC Covid Response Team, 2020). Preliminary studies highlighted severe presentations in which peripheral ischemia resulted in acral cyanosis, gangrene, ulceration, or necrosis (Zhang et al., 2020). Patients were found to be in a hypercoagulable state with ultimately poor prognosis for recovery despite anticoagulation attempts (Zhang et al., 2020). In contrast, recent studies in Spain, the Middle East, and the United States have described a more benign course in which patients were generally younger and often asymptomatic (Galván Casas et al., 2020; Piccolo et al., 2020; Alramthan and Aldaraji, 2020). Minimal images of COVID toes have been shared of African American patients in the aforementioned studies, such that healthcare providers may have difficulty in identification of subtler presentations in darker skin tones (Buster et al., 2020). As seen with the first case patient, COVID toes may have an atypical appearance as her lesions were not painful, overtly swollen or enlarged, and had only mild discoloration relative to the surrounding skin. In contrast the nodules in the lighter skinned Hispanic/ Asian child are easily identified.

Continued research and reporting are needed to better understand COVID toes and associations in the COVID-19 disease process. It appears at this time that COVID toes may be a good prognostic indicator for recovery. However, as our adult patient demonstrates, patients can develop significant symptoms. Furthermore, clinical lesions may be more difficult to detect in patients of color. Implications regarding long term immunity are unknown. Early demographics for COVID-19 related hospitalizations and deaths have shown a higher prevalence among older or immunocompromised patients, particularly those with pre-existing health conditions(16). COVID toes of adults and children, appear to be of similar morphology

and presentation as well as a good prognosis for recovery. Recent efforts by the American Academy of Dermatology's COVID-19 Dermatology Registry and large-scale studies are critical in deciphering how the virus affects skin differently among patients (Freeman *et al.*, 2020). Current treatment for COVID toes mirror that of classical chilblains in which the off-label utilization and combination of high potency corticosteroids, analgesics and topical calcium channel blockers have had mixed results (Galván Casas *et al.*, 2020). Usage of aspirin has been described in adults, however, risk of Reye's syndrome in children with aspirin usage limits prescribing (Prakash and Weisman, 2009).

Conclusion

COVID toes are a unique dermatological finding in COVID-19 patients, with a pseudo-chilblain appearance and course. As seen with the case patients, COVID toes can have a more classical morphology with painful, red lesions versus atypical ones with only discoloration and no pain. This spectrum of presentations coupled with the imprecision of COVID-19 testing complicates the ability for physicians to correlate COVID toes with a COVID-19 infection. Ongoing studies are continuously updating the body of knowledge in regard to COVID-19 and its extrapulmonary manifestations.

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