

Leukoerythroblastic reaction in a patient with COVID-19 infection

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A 46-year-old previously healthy female developed flu-like symptoms several days prior to presenting to a local hospital. The patient did not note a recent travel history, specifically denying travel to China. Sick contact exposure was not known. Chest X-ray and CT of the chest showed findings concerning for lobar pneumonia. Subsequent radiology studies showed worsening lung findings along with worsening respiratory symptoms necessitating intubation and ventilation. She was transferred to a university hospital due to the need for an elevated level of care.

Upon arrival to the university hospital, it was confirmed that the patient was positive for the novel Corona virus disease 2019 (COVID-19). Her initial complete blood count (CBC) showed a normal leukocyte count (7.3 x 103/uL) with lymphopenia (200/uL), normocytic anemia (hemoglobin = 10.4 g/dL), and a normal platelet count (213 x 103/uL). Three days after admission to the university hospital (approximately one week after initial hospital admission), the patient developed leukocytosis (14.1 x 103/uL) with a left-shifted population of neutrophilic cells, prompting peripheral blood smear evaluation. Mild monocytosis (900/uL) along with the persistent lymphopenia was also noted. The peripheral blood smear was consistent with a leukoerythroblastic picture with normocytic anemia with occasional nucleated red blood cells (Figure 1 panel A), mild anisocytosis, and rare dacrocytes. Schistocytes were absent. Neutrophilia with left shifted myeloid cells, including occasional myelocytes and rare promyelocytes were noted (Figure 1 panels B, C, and D). Lymphopenia was confirmed with many smudge cells seen. Platelets were adequate with some large platelets seen.

Figure 1

Review of the peripheral smear at 100x magnification shows a nucleated erythroid (Panel A), rare blast, with prominent nucleoli and immature chromatin pattern (Panel B), a left shifted myeloid series with immature promyelocytes and metamyelocyte (Panel C), and occasional monocytes noted (Panel D).

Leukoerythroblastic reactions, defined as immature erythroid and immature myeloid cells circulating in the peripheral blood, are uncommon. Leukoerythroblastic blood findings are typically seen in disorders

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associated with bone marrow fibrosis including myelofibrosis and other myeloproliferative disorders, and cancers with metastatic disease to the bone marrow. Leukoerythroblastosis can rarely be seen in viral infections such as parvovirus [1,2].

Corona virus disease 2019 (COVID-19) is an infectious viral disease caused by a pathogen named severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) [3]. COVID-19 has created a public health emergency in many countries, including the United States, and is expected to continue to spread. Few strains of corona virus are known to cause flu-like upper respiratory illnesses in humans, but the current strain, SARS-CoV-2, was not known to cause any human illnesses prior to this outbreak. Thus, very little information about COVID-19's pathogenic role in humans is known. Other coronaviruses have been previously found responsible for infection in both animals and humans, including the viruses responsible for the Middle East Respiratory Syndrome (MERS-CoV) [4] and severe acute respiratory syndrome (SARS-CoV) [5]. Most of the patient morbidity and mortality attributable to COVID-19 has been reported in Wuhan, China, [6]. Other patients with symptoms of severe infection have now been reported in South Korea, Italy, German, and Iran among others. Following the recent availability of COVID-19 testing, it is likely the number of diagnosed cases will significantly increase.

Our understanding of this novel virus continues to evolve. The spectrum of disease, clinical manifestations, and pathophysiology underpinning this infection will be further elucidated over the coming months. Although we are only reporting the findings of a single patient with COVID-19, our aim is to describe the unusual finding of leukoerythroblastosis in this viral infection. We cannot definitively conclude that our findings are secondary to infection with COVID-19. However, following clinical improvement, the neutrophilia has resolved with the other blood findings slowing improving. With our description of these findings, it is our hope to avoid additional work-up and diagnostic testing in other patients with COVID-19, as well as contribute to our understanding of this novel infection.

References:

1. Anand A, Gray ES, Brown T, Clewley JP, Cohen BJ. Human parvovirus infection in pregnancy and hydrops fetalis. N Engl J Med. 1987 Jan 22;316(4):183-6.

 Ozdemir N, Akı H, Hakyemez HT, Cokuğraş FÇ, Apak H. Parvovirus B19 infection mimicking juvenile myelomonocytic leukemia. Int J Infect Dis. 2010 Sep;14 Suppl 3:e379-80. doi: 10.1016/j.ijid.2010.04.004. Epub 2010 Jul 6.

3. Fauci AS, Lane HC, Redfield RR. Covid-19 - Navigating the Uncharted. N Engl J Med. 2020 Feb 28. doi: 10.1056/NEJMe2002387.

4. Gostin LO, Lucey D. Middle East Respiratory Syndrome: A Global Health Challenge. JAMA. 2015 Aug 25;314(8):771-2. doi: 10.1001/jama.2015.7646

5. https://www.cdc.gov/coronavirus/types.html

6. Wu Z, McGoogan JM. Characteristics of and Important Lessons From the Coronavirus Disease 2019 (COVID-19) Outbreak in China: Summary of a Report of 72 314 Cases From the Chinese Center for Disease Control and Prevention. JAMA. 2020 Feb 24. doi: 10.1001/jama.2020.2648.

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