

Title	Human infection with a novel avian influenza A(H5N6) virus
Author(s)	Yang, ZF; Mok, KP; Peiris, JSM; Zhong, NS
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## **U.S.-Citizen International Medical Graduates**

TO THE EDITOR: The Perspective article on U.S.citizen international medical graduates (IMGs) by Eckhert and van Zanten (April 30 issue)1 used data from the National Resident Matching Program (NRMP)<sup>2</sup> to assert that "the forecasts that there would soon be no room in U.S. residency programs for IMGs were not accurate: since the number of positions for postgraduate-year-1 [PGY-1] residents increased by 26% between 2004 and 2014." Before the 2013 "all-in" policy of the NRMP, data for the Residency Match did not accurately reflect the number of PGY-1 residents who actually entered training, because programs could fill positions (e.g., with IMGs) outside the Match. The Accreditation Council for Graduate Medical Education (ACGME) collects actual counts of residents occupying positions in accredited programs.3

Before "all-in," the 2011 Match placed 23,420 PGY-1 residents, and the ACGME counted 26,737 entry-level residents in 2011-2012 (in "pipeline" programs leading to initial board certification). After "all-in," the 2014 Match placed 26,678 PGY-1 residents, and the ACGME counted 27,534 firstyear pipeline residents in 2014-2015, which is consistent with closer compliance with the "allin" policy. According to the ACMGE counts, the number of positions actually occupied by entrylevel pipeline residents increased from 24,069 in 2004-2005 to 27,534 in 2014-2015, a 14.4% increase.

Paul H. Rockey, M.D., M.P.H. Rebecca S. Miller, M.S. Thomas J. Nasca, M.D.

Accreditation Council for Graduate Medical Education Chicago, IL prockey@acgme.org

No potential conflict of interest relevant to this letter was reported.

1. Eckhert NL, van Zanten M. U.S.-citizen international medical graduates — a boon for the workforce? N Engl J Med 2015; 372:1686-7.

2. National Resident Matching Program (NRMP). Results and data: 2014 main residency match. Washington, DC: NRMP, April 2014 (http://www.nrmp.org/wp-content/uploads/2014/04/ Main-Match-Results-and-Data-2014.pdf).

3. Accreditation Council for Graduate Medical Education (ACGME). Data resource book, academic year 2013-2014 (https:// www.acgme.org/acgmeweb/tabid/259/Publications/Graduate MedicalEducationDataResourceBook.aspx).

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THE AUTHORS REPLY: We appreciate the clarification by Rockey et al. with regard to data from the ACGME. Although the magnitude of the increase in graduate medical education positions during the past decade is smaller than we indicated, our statement concerning the continued supply of residency positions for IMGs remains valid. Data from the 2015 Match indicate that there were 1.51 positions per senior at U.S. allopathic medical schools, with IMGs (both U.S.-citizen and non-U.S.-citizen) filling 6301 of these excess positions,<sup>1</sup> despite the increasing pool of seniors at U.S. allopathic medical schools and students and graduates of U.S. osteopathic medical schools.

N. Lynn Eckhert, M.D., Dr.P.H.

Partners HealthCare International Boston, MA

Marta van Zanten, Ph.D.

Foundation for Advancement of International Medical Education and Research

Philadelphia, PA

Since publication of their article, the authors report no further potential conflict of interest.

1. National Resident Matching Program (NRMP). Advance data tables: 2015 main residency match. Washington, DC: NRMP, March 2015 (http://www.nrmp.org/wp-content/uploads/2015/03/ ADT2015\_final.pdf).

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## Human Infection with a Novel Avian Influenza A(H5N6) Virus

novel, highly pathogenic avian influenza A(H5) viruses in clade 2.3.4.4 have recently emerged and spread in Asia, Europe, and North America,

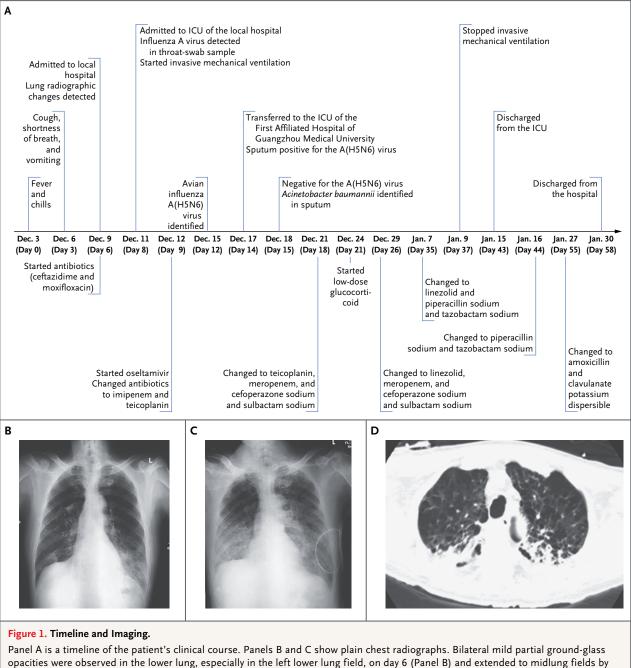
TO THE EDITOR: Multiple reassortant strains of giving rise to multiple subtypes (H5N2, H5N5, H5N6, and H5N8).1-3 We report on a 59-year-old man in Guangzhou, China, with influenza A(H5N6) infection and associated illness. He had

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day 13 (Panel C). Panel D is a computed tomographic scan of the chest (lung window), showing bilateral ground-glass opacity and consolidation on day 14. ICU denotes intensive care unit.

> a 30-year history of smoking and had stopped set, he regularly purchased and handled live smoking in January 2014. He had undergone sur- poultry in live poultry markets. gery and chemotherapy for colon cancer, the last

Fever and chills developed on December 3, chemotherapy dose having been administered in 2014. In the next 3 days, his body temperature April 2014. During the weeks before illness on- rose to 40°C, and the fever was not relieved by

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acetaminophen; vomiting, cough, and shortness of breath also developed. He was hospitalized on day 6 of his illness and began receiving antibiotic treatment (Fig. 1A). Chest radiography revealed mild bilateral ground-glass opacities in his lower lung (Fig. 1B). Two days later, pulse oximetry showed that the arterial blood oxygen saturation levels had declined (to approximately 50 to 70%) in spite of the administration of highflow oxygen (5 to 6 liters per minute), and invasive mechanical ventilation was commenced. A throat-swab sample was positive for influenza A virus, as assessed by means of real-time reversetranscriptase-polymerase chain reaction (PCR), and treatment with oseltamivir (150 mg twice a day for 13 days, administered through a nasogastric tube) was initiated on day 9. On day 12 of his illness, an influenza A virus isolate, designated A/Guangzhou/39715/2014 (H5N6), was obtained from a throat-swab specimen (see the Supplementary Appendix, available with the full text of this letter at NEJM.org, for details). Influenza A(H5N6) was identified by means of subtypespecific PCR and sequencing of the virus isolate (GenBank accession numbers, KP765785 through KP765792).

On day 14, the patient was transferred to the isolation ward of the intensive care unit in the First Affiliated Hospital of Guangzhou Medical University. A chest radiograph and a computed tomographic scan showed progression of lung consolidation (Fig. 1C and 1D). Influenza A RNA was detectable by means of real-time PCR in sputum samples but not from throat-swab, blood, urine, or stool samples. The next day, virus was undetectable in throat-swab, sputum, and bronchoalveolar-fluid specimens. Multidrug-resistant Acinetobacter baumannii was detected in sputum but not in blood cultures. There was evidence of neutrophilia and lymphocytopenia (Table S1 in the Supplementary Appendix). Serum levels of aspartate aminotransferase, creatinine, lactate dehydrogenase, creatine kinase, and myoglobulin were within the normal range throughout his illness. Antibiotic therapy was changed to teicoplanin, meropenem, and cefoperazone-sulbactam on day 18. Low-dose methylprednisolone treatment was started on day 21 and was administered until day 42. Imaging showed resolution of the bilateral lung infiltrations. Fever recurred on day 25 in association with eosinophilia. Invasive mechanical ventilation was stopped on day 37,

and the patient was later discharged (day 58). This case shows that a novel H5 clade 2.3.4.4 influenza virus can cause human infection, similar to other influenza A(H5N1) viruses.<sup>4</sup>

Zi-Feng Yang, M.D., Ph.D.

State Key Laboratory of Respiratory Disease Guangzhou, China

Chris K.P. Mok, Ph.D. Joseph S.M. Peiris, Ph.D.

University of Hong Kong Hong Kong, China

Nan-Shan Zhong, M.D.

State Key Laboratory of Respiratory Disease Guangzhou, China nanshan@vip.163.com

Drs. Yang and Mok contributed equally to this letter.

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Disclosure forms provided by the authors are available with the full text of this letter at NEJM.org.

1. World Health Organization. Antigenic and genetic characteristics of zoonotic influenza viruses and development of candidate vaccine viruses for pandemic preparedness. February 2015 (http://www.who.int/influenza/vaccines/virus/201502\_zoonotic \_vaccinevirusupdate.pdf?ua=1).

2. European Food Safety Authority. Highly pathogenic avian influenza A subtype H5N8. Parma, Italy: EFSA, 2014 (http://www.efsa.europa.eu/en/efsajournal/doc/3941.pdf).

**3.** Outbreaks of avian influenza A (H5N2), (H5N8), and (H5N1) among birds — United States, December 2014–January 2015. MMWR Morb Mortal Wkly Rep 2015;64:111.

**4.** Writing Committee of the Second World Health Organization Consultation on Clinical Aspects of Human Infection with Avian Influenza A (H5N1) Virus. Update on avian influenza A (H5N1) virus infection in humans. N Engl J Med 2008;358:261-73.

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