of healing of suppurative nodes and is the only evidence-based effective treatment (9). Surgical excision remains controversial because of potentially high rates of significant scarring (10). For nonsuppurative lymphadenitis, a watch-and-wait approach is recommended because most resolve rapidly (8).

Given our findings, the National TB Program in Georgia subsequently created a management protocol. This protocol recommends no intervention for nonsuppurative lymphadenitis and needle aspiration for suppurative local lymphadenitis.

In summary, we found an increasing rate of BCG-associated lymphadenitis after a shift to exclusive BCG SSI vaccine use in Georgia. Countries with a BCG vaccination policy should have a clear protocol on management of BCG vaccine–related adverse events to avoid inappropriate treatment in children.

References

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Fatal Accelerated Cirrhosis after Imported HEV Genotype 4 Infection


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To the Editor: Hepatitis E is a viral hepatitide that is endemic in many developing countries. In its classic form, it results from ingesting fecally contaminated water that carries hepatitis E virus (HEV), and it frequently resolves without treatment. When hepatitis E is imported to the United States, it originates mainly from persons who have acquired HEV genotype 1 infection from South Asia (1). We report imported HEV genotype 4 infection (Technical Appendix Figure, panel A) in a patient during which cirrhosis and fatal hepatic decompensation ensued.

The patient was a 68-year-old man of Chinese ethnicity who had been a California resident since 1985. He sought treatment for mild jaundice in April 2013 in Hong Kong, where he had been staying for 7 weeks. Sixteen years before, he had undergone orthotopic liver transplantation at Stanford University Medical Center (Palo Alto, California, USA) for hepatitis B cirrhosis. Since then, he had received entecavir and tacrolimus for maintenance and had been vaccinated against hepatitis A virus. Until his current illness, routine liver function tests had not indicated hepatic dysfunction (values in November 2012: alanine aminotransferase 69 IU/L, aspartate aminotransferase 2 IU/L, alkaline phosphatase 67 IU/L, total bilirubin 0.5 mg/dL).

When the patient returned to the United States, 3 weeks after onset of jaundice, the initial work-up showed the following values: alanine aminotransferase 149 IU/L, aspartate aminotransferase 24 IU/L, alkaline phosphatase 67 IU/L, total bilirubin 0.5 mg/dL).

When the patient returned to the United States, 3 weeks after onset of jaundice, the initial work-up showed the following values: alanine aminotransferase 149 IU/L, aspartate aminotransferase 59 IU/L, alkaline phosphatase 193 IU/L, total bilirubin 2.8 mg/dL (online Technical Appendix Figure, panel B, http://wwwnc.cdc.gov/EID/article/21/9/15-0300-Techapp1.pdf). Hepatitis B virus DNA and anti-nuclear antibodies were not detected, and the tacrolimus level was stable. Ultrasound revealed a normal transplanted liver. A liver biopsy specimen showed mild portal, biliary, and lobular inflammation and early biliary injury (Figure,
The prednisone dosage was escalated, and mycophenolate mofetil was added. Liver enzyme activity showed some improvement, but the bilirubin level continued to rise (online Technical Appendix Figure, panel B).

A biopsy specimen taken 3 months later showed grade 3 hepatitis with bile ductular reaction, bridging hepatocytic necrosis and fibrosis, and regenerative nodule formation (Figure, panels C, D). A blood sample taken about this time tested positive for HEV RNA. The patient was then given ribavirin (1,000 mg/d). Before hepatitis E was diagnosed, tacrolimus was given (1 mg 2×/d); when the diagnosis was confirmed, the tacrolimus dose was reduced to 0.5 mg every other day.

Eight months after onset of the patient’s condition, marked hepatic decompensation occurred (online Technical Appendix Figure), culminating in esophageal variceal hemorrhage. The patient was placed on a waiting list and then underwent liver transplantation, but he died during the operation from complications of hemorrhage. Biopsy of the liver explant revealed intense lobular inflammation with the hepatocellular reactive changes persisting and stage IV fibrosis (Figure, panels E, F).

The patient had lived and worked in Hong Kong before he became a resident of the United States. He had not visited Hong Kong in the 3 years preceding his most recent trip, nor had he traveled to Europe. Review of his medical records revealed no evidence of hepatic dysfunction after his previous travels. Considering that his most recent visit to Hong Kong coincided with the incubation period of hepatitis E (2), he most likely acquired HEV genotype 4 infection during that visit.

In China over the past decade, national notifications of HEV infection have risen, with 28,232 cases reported in 2013 (3). In Hong Kong, where a rising trend in hepatitis E notifications also has been observed (150 cases reported in 2012 [4]), HEV infections are almost all associated with HEV genotype 4 (5).

This patient’s HEV subgenomic sequence was closely related to human and porcine HEV genotype 4 sequences.
reported from mainland China and Hong Kong (online Technical Appendix Figure, panel A). Porcine liver has been implicated as a possible HEV transmission vehicle in that region (6); although we do not know whether the patient ate food that carried HEV, the possibility underscores the importance of avoiding eating inadequately cooked animal-derived food products during international travel (2).

Chronic hepatitis with accelerated cirrhosis has been reported in solid-organ transplant recipients infected with HEV genotype 3, but not with genotype 4 (7). Serial liver biopsy specimens from the patient showed persistent and worsening hepatitis and rapid onset of fibrosis that intensified (online Technical Appendix Figure, panel B).

Testing for HEV infection is recommended during initial assessments of posttransplant hepatic dysfunction because histologic appearances in liver biopsy specimens may not clearly distinguish between graft rejection and acute viral hepatitis (Figure, panels A, B). Early diagnosis of hepatitis E should lead to prompt administration of anti-viral therapy and appropriate adjustments to the immunosuppressant drug regimen, particularly because some drugs can exert opposing effects on HEV replication (8).

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References

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Measles Reemergence in Ceará, Northeast Brazil, 15 Years after Elimination

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To the Editor: Measles was endemic in Brazil before 2000 and caused large outbreaks every 2 or 3 years (1). Although measles was eliminated in Brazil in 2000, cases have continued to be imported (2,3). During 2001–2014, the median annual number of measles cases reported in Brazil was 50 (range 2–712). The median annual number of Brazilian states with reported cases was 2.5 (range 1–7). Since elimination, the highest numbers of cases reported in Brazil occurred in 2013 (220) and in 2014 (712) (3–5). According to the Pan American Health Organization, endemic transmission is reestablished when epidemiologic and laboratory evidence indicate that a chain of transmission of a virus strain has continued uninterrupted for ≥12 months in a defined geographic area (6).

From December 2, 2013, through December 31, 2014, in the state of Ceará, Brazil, 681 measles cases were reported. A measles case was considered confirmed when a patient exhibited fever, rash, and ≥1 of 3 symptoms and signs (i.e., cough, runny nose, conjunctivitis); was positive for IgM and negative for IgG against measles virus; and had not been vaccinated in the previous 21 days. D8 genotype, the same virus genotype that was circulating in Europe, was the only genotype identified, and how the virus was introduced into the region was not clear (4,5). From 2000 to 2013, vaccine coverage among children 12 months of age remained >95% in Ceará, although that coverage was not homogeneous for the whole state. In 14.7% (27/184) of municipalities, the vaccination coverage was much lower...
Fatal Accelerated Cirrhosis after Imported HEV Genotype 4 Infection

Technical Appendix

Technical Appendix Figure. A) Phylogenetic tree comparing a 258-nt sequence within hepatitis E virus (HEV) open reading frame 1 of the patient who visited Hong Kong in 2013 with corresponding, representative GenBank sequences. Included is a corresponding sequence from patient A, a 63-year-old Caucasian man, a resident of
Alabama, in whom acute, self-resolving hepatitis developed 5 weeks after he returned from a 2-week visit to Shandong, China. Numerals beside each sequence denote year of sequence reporting; alphanumerics denote GenBank accession numbers. GT, genotype; CH, China; HK, Hong Kong; MX, Mexico; US, United States. B) Chronology of changes in liver function and hepatitis E markers in the patient.