RESEARCH LETTERS

and mass reproduction of common voles in several parts of Europe, TULV should be considered as a threat to human health.

Acknowledgments

We gratefully acknowledge the expert technical assistance by Christine Stephan, Susanne Schwarz, and Dörte Kaufmann and helpful discussions with Stefan Drewes.

This work was supported by the German Federal Ministry of Public Health through Robert Koch Institute (grant nos. 1369-382/435 and 1362-924/980) and the Bundesministerium für Bildung und Forschung through the Research Network Zoonotic Infectious Diseases (grant nos. FKZ 01KI1721A and FKZ 01KI1721C).

About the Author

Dr. Hofmann is the chair of the National Consultation Laboratory for Hantaviruses, Institute of Virology, Charité– Universitätsmedizin Berlin, Germany. His primary research interest is human infections with viral pathogens.

References

- Kruger DH, Figueiredo LT, Song JW, Klempa B. Hantaviruses – globally emerging pathogens. J Clin Virol. 2015;64:128–36. https://doi.org/10.1016/j.jcv.2014.08.033
- Vapalahti O, Lundkvist A, Kukkonen SK, Cheng Y, Gilljam M, Kanerva M, et al. Isolation and characterization of Tula virus, a distinct serotype in the genus *Hantavirus*, family *Bunyaviridae*. J Gen Virol. 1996;77:3063–7. https://doi.org/10.1099/0022-1317-77-12-3063
- Schmidt S, Saxenhofer M, Drewes S, Schlegel M, Wanka KM, Frank R, et al. High genetic structuring of Tula hantavirus. Arch Virol. 2016;161:1135–49. https://doi.org/10.1007/ s00705-016-2762-6
- Klempa B, Meisel H, Räth S, Bartel J, Ulrich R, Krüger DH. Occurrence of renal and pulmonary syndrome in a region of northeast Germany where Tula hantavirus circulates. J Clin Microbiol. 2003;41:4894–7. https://doi.org/10.1128/ JCM.41.10.4894-4897.2003
- Zelená H, Mrázek J, Kuhn T. Tula hantavirus infection in immunocompromised host, Czech Republic. Emerg Infect Dis. 2013;19:1873–5. https://doi.org/10.3201/eid1911.130421
- Reynes JM, Carli D, Boukezia N, Debruyne M, Herti S. Tula hantavirus infection in a hospitalised patient, France, June 2015. Euro Surveill. 2015;20:30095. https://doi.org/ 10.2807/1560-7917.ES.2015.20.50.30095
- Klempa B, Fichet-Calvet E, Lecompte E, Auste B, Aniskin V, Meisel H, et al. Hantavirus in African wood mouse, Guinea. Emerg Infect Dis. 2006;12:838–40. https://doi.org/10.3201/ eid1205.051487
- Mertens M, Hofmann J, Petraityte-Burneikiene R, Ziller M, Sasnauskas K, Friedrich R, et al. Seroprevalence study in forestry workers of a non-endemic region in eastern Germany reveals infections by Tula and Dobrava–Belgrade hantaviruses. Med Microbiol Immunol (Berl). 2011;200:263–8. https://doi.org/10.1007/s00430-011-0203-4
- 9. Ulrich R, Meisel H, Schütt M, Schmidt J, Kunz A, Klempa B, et al. Prevalence of hantavirus infections in Germany [in

German]. Bundesgesundheitsblatt Gesundheitsforschung Gesundheitsschutz. 2004;47:661–70. https://doi.org/10.1007/ s00103-004-0858-8

 Sibold C, Meisel H, Lundkvist A, Schulz A, Cifire F, Ulrich R, et al. Short report: simultaneous occurrence of Dobrava, Puumala, and Tula hantaviruses in Slovakia. Am J Trop Med Hyg. 1999;61:409–11. https://doi.org/10.4269/ ajtmh.1999.61.409

Address for correspondence: Jörg Hofmann, Institute of Virology, Helmut-Ruska-Haus, Charité University Medicine, Charitéplatz 1, 10117 Berlin, Germany; email: joerg.hofmann@charite.de

Rapid Spread and Control of Multidrug-Resistant Gram-Negative Bacteria in COVID-19 Patient Care Units

Ashka Patel, Michele Emerick, Marie K. Cabunoc, Michelle H. Williams, Michael Anne Preas, Gregory Schrank, Ronald Rabinowitz, Paul Luethy, J. Kristie Johnson, Surbhi Leekha

Author affiliations: University of Maryland Medical Center, Baltimore, Maryland, USA (A. Patel, M. Emerick, M.K. Cabunoc, M.H. Williams, M.A. Preas); University of Maryland School of Medicine, Baltimore (G. Schrank, R. Rabinowitz, P. Luethy, J.K. Johnson, S. Leekha)

DOI: https://doi.org/10.3201/eid2704.204036

We describe rapid spread of multidrug-resistant gramnegative bacteria among patients in dedicated coronavirus disease care units in a hospital in Maryland, USA, during May–June 2020. Critical illness, high antibiotic use, double occupancy of single rooms, and modified infection prevention practices were key contributing factors. Surveillance culturing aided in outbreak recognition and control.

Bacterial colonization and secondary infection have been described in patients hospitalized with coronavirus disease (COVID-19) (1,2). We report a singlecenter experience with spread of multidrug-resistant (MDR) gram-negative bacteria (GNB) in COVID-19 patients in Maryland, USA, during May-June 2020. This investigation was determined to be non-human subjects research by the University of Maryland's Institutional Review Board.

At University of Maryland Medical Center (Baltimore, MD, USA), an 800-bed tertiary-care hospital, since early April 2020, critically ill COVID-19 patients had been housed in 3 dedicated units (3), which included 2 intensive care units (ICUs) (units A and B, unit A providing extracorporeal membrane oxygenation support) and 1 intermediate-care unit (unit C). Units were designed as closed, negative-pressure areas where staff remained in the same personal protective equipment while providing care to multiple patients. To accommodate the COVID-19 surge, single-patient ICU rooms in units A and B frequently housed 2 patients. Unit C rooms remained singleoccupancy and received patients for step-down care from units A and B. Hospital policy required staff to change gloves and perform hand hygiene (or glove hygiene if wearing 2 layers of gloves) between patients and to wear 2 layers of gowns for patients with resistant organisms and remove the outer gown before moving to the next patient. A team nursing model was used, in which multiple nurses shared responsibilities for each patient during a shift.

For routine surveillance, the hospital defined MDR GNB as Enterobacterales, *Acinetobacter baumannii*, or *Pseudomonas aeruginosa* nonsusceptible to ≥ 2 of piperacillin/tazobactam, cefepime, and a carbapenem. Before COVID-19, we performed admission and weekly surveillance for MDR Enterobacterales and *A. baumannii* using perirectal swab specimens on medical and surgical ICU patients and monitored hospitalwide MDR GNB incidence by using the first positive clinical or surveillance culture >48 hours postadmission.

In mid-May 2020, a cluster of 4 patients with MDR Escherichia coli was identified on unit A. Hospitalwide data showed increase in MDR GNB incidence from baseline (Figure, panel A) (weeks 9-11), driven by E. coli cases on units A and B (Figure, panel B). Further review also revealed several patients with cefepime-resistant E. coli (not meeting institutional MDR criteria), MDR P. aeruginosa, and MDR A. baumannii. Surveillance screens (perirectal swab specimens on all and sputum on ventilated patients) in the 3 units in week 12 identified 18/29 (62%) additional patients with resistant GNB (MDR GNB, cefepime-resistant E. coli, or both). Public health authorities were notified and observations of practice and discussions with leadership were conducted. Twice-weekly surveillance culturing among patients still negative for resistant GNB was instituted (Figure).



Figure. Incidence of patients with a clinical or surveillance culture-positive result indicating MDR or cefepime-resistant Escherichia coli, MDR Acinetobacter baumannii, or MDR Pseudomonas aeruginosa >48 hours after admission to a hospital in Maryland, USA, by week, March 1-July 31, 2020. A) Overall hospitalwide incidence (118 total cases, with 98 positive cultures belonging to outbreak units). Narrow white bars represent the number of surveillance cultures obtained during the outbreak and shaded bars show positive cultures by organism. Arrows show timing of relevant events for transmission and control. B) Incidence of outbreak cases (n = 98) stratified by the 3 units affected by the outbreak. Organisms nonsusceptible to ≥2 of piperacillin/tazobactam, cefepime, or carbapenem are considered MDR. Patients are included for the first positive culture per organism and therefore might be included more than once. MDR, multidrug-resistant.

During April 16–July 15, a total of 71 unique patients had positive clinical or surveillance cultures for resistant GNB, including 44 *E.coli* (33 MDR and 11 cefepime-resistant), 27 MDR *P. aeruginosa*, and 27 MDR *A. baumannii* (Appendix Table 1, https://wwwnc.cdc. gov/EID/article/27/4/20-4036-App1.pdf). Twentyfour patients (34%) were co-colonized with >1 resistant GNB. Of the 71 patients, 69 (97%) had received antibiotics before first positive resistant GNB culture, 30 (42%) required extracorporeal membrane oxygenation support, 27 (38%) required renal replacement therapy, 52 (73%) received corticosteroids, 25 (35%) received remdesivir, and 14 (20%) received tocilizumab. Twenty-three (32%) patients ultimately died.

Relatedness of early E. coli isolates was assessed by pulsed-field gel electrophoresis (PFGE) (n = 13, weeks 7–11) and genetic β -lactamase determination by Verigene gram-negative blood culture nucleic acid test (Luminex Corporation, https://www.luminexcorp.com) (n = 38, weeks 7-14) (4; Appendix). PFGE revealed 3 groups. Groups 1 and 2 (n = 7) were considered related and were negative for β-lactamases; these and 8/10 additional β -lactamase-negative isolates were from unit B. Group 3 (n = 6) isolates did not produce bands but were positive for CTX-M; these and 14/15 additional CTX-M positive isolates (including 10/11 phenotypically cefepime-resistant but not MDR) were from unit A and considered related, suggesting rapid patient-to-patient transmission (Appendix Table 1). MDR P. aeruginosa transmission occurred predominantly in unit A, whereas MDR A. baumannii was largely in unit B. Resistant GNB were likely introduced into unit C from both units A and B (Figure, panel B).

Key infection control findings (5) included tight physical spaces and close proximity of patients in double occupancy (6), multiple staff in contact with each patient in the team nursing model, and low compliance with hand and glove hygiene and gown changes between patients. To limit staff exposure to COVID-19 patients, the unit had less support from ancillary services; instead, daily room and equipment cleaning and stocking of medications and supplies were performed by unit-based clinical staff.

Outbreak control interventions included discontinuation of double occupancy, frequent infection prevention rounds to promote hand hygiene and glove and gown changes between patients, increased environmental services support, and attention to disinfection of reusable equipment and hightouch surfaces (Appendix Table 2) (7). Surveillance culturing showed a decrease in positive cultures over time (Figure). Prolonged critical illness, high antibiotic and corticosteroid use, double occupancy, the team nursing model, and modified infection prevention practice were considered contributors to transmission, underscoring the importance of vigilance to MDR organisms in this setting (5,7–10). Surveillance culturing aided with recognizing the extent of spread and informed early intervention.

Acknowledgments

We would like to thank Richard Brooks and Heather Saunders for guidance on outbreak management, and Gwen Robinson for assistance with creating the figure.

About the Author

Dr. Patel is a second-year infectious diseases fellow at the University of Maryland Medical Center. She is interested in infection prevention and hospital epidemiology and has worked on projects involving hospital- acquired *Clostridium difficile* infections as well as hospital-onset bloodstream infections.

References

- Rawson TM, Moore LSP, Castro-Sanchez E, Charani E, Davies F, Satta G, et al. COVID-19 and the potential longterm impact on antimicrobial resistance. J Antimicrob Chemother. 2020;75:1681–4. https://doi.org/10.1093/jac/ dkaa194
- Nori P, Cowman K, Chen V, Bartash R, Szymczak W, Madaline T, et al. Bacterial and fungal coinfections in COVID-19 patients hospitalized during the New York City pandemic surge. Infect Control Hosp Epidemiol. 2021;42:84– 8. https://doi.org/10.1017/ice.2020.368
- Centers for Disease Control and Prevention. Coronavirus disease 2019 (COVID-19). 2020 [cited 2020 Aug 20]. https://www.cdc.gov/coronavirus/2019-ncov/hcp/ infection-control-recommendations.html
- Hill JT, Tran K-DT, Barton KL, Labreche MJ, Sharp SE. Evaluation of the nanosphere Verigene BC-GN assay for direct identification of Gram-negative bacilli and antibiotic resistance markers from positive blood cultures and potential impact for more-rapid antibiotic interventions. J Clin Microbiol. 2014;52:3805–7. https://doi.org/10.1128/ JCM.01537-14
- Donà D, Di Chiara C, Sharland M. Multi-drug-resistant infections in the COVID-19 era: a framework for considering the potential impact. J Hosp Infect. 2020;106:198–9. https://doi.org/10.1016/j.jhin.2020.05.020
- Kaier K, Mutters NT, Frank U. Bed occupancy rates and hospital-acquired infections – should beds be kept empty? Clin Microbiol Infect. 2012;18:941–5. https://doi.org/ 10.1111/j.1469-0691.2012.03956.x
- Getahun H, Smith I, Trivedi K, Paulin S, Balkhy HH. Tackling antimicrobial resistance in the COVID-19 pandemic. Bull World Health Organ. 2020;98:442–442A. https://doi.org/ 10.2471/BLT.20.268573
- 8. Rawson TM, Moore LSP, Zhu N, Ranganathan N, Skolimowska K, Gilchrist M, et al. Bacterial and fungal

co-infection in individuals with coronavirus: a rapid review to support COVID-19 antimicrobial prescribing. Clin Infect Dis. 2020;71:2459–68.

- Vincent J-L, Sakr Y, Singer M, Martin-Loeches I, Machado FR, Marshall JC, et al.; EPIC III Investigators. Prevalence and outcomes of infection among patients in intensive care units in 2017. JAMA. 2020;323:1478–87. https://doi.org/10.1001/ jama.2020.2717
- Prestel C, Anderson E, Forsberg K, Lyman M, de Perio MA, Kuhar D, et al. *Candida auris* outbreak in a COVID-19 specialty care unit – Florida, July–August 2020. MMWR Morb Mortal Wkly Rep. 2021;70:56–7. https://doi.org/ 10.15585/mmwr.mm7002e3

Address for correspondence: Surbhi Leekha, University of Maryland School of Medicine, 10 South Pine St, MSTF 334F, Baltimore, MD 21201, USA; email: sleekha@som.umaryland.edu

Cetacean Morbillivirus and *Toxoplasma gondii* Co-infection in Mediterranean Monk Seal Pup, Italy

Antonio Petrella, Sandro Mazzariol, Iolanda Padalino, Gabriella Di Francesco, Cristina Casalone, Carla Grattarola, Giovanni Di Guardo, Camilla Smoglica, Cinzia Centelleghe, Claudia Gili

Author affiliations: Istituto Zooprofilattico Sperimentale della Puglia e della Basilicata, Foggia, Italy (A. Petrella, I. Paladino); University of Padova, Padua, Italy (S. Mazzariol, C. Centelleghe); Istituto Zooprofilattico Sperimentale dell'Abruzzo e del Molise, Teramo, Italy (G. Di Francesco); Istituto Zooprofilattico Sperimentale del Piemonte, Liguria e Valle d'Aosta, Turin, Italy (C. Casalone, C. Grattarola); University of Teramo Faculty of Veterinary Medicine, Teramo (G. Di Guardo, C. Smoglica); Stazione Zoologica Anton Dohrn, Naples, Italy (C. Gili)

DOI: https://doi.org/10.3201/eid2704.204131

A Mediterranean monk seal (*Monachus monachus*) pup from the southern Adriatic coast of Italy showed cetacean morbillivirus (CeMV) and disseminated *Toxoplasma gondii* co-infection, which probably resulted from CeMV-induced immunosuppression. These findings are of concern for the conservation of this critically endangered species. The Mediterranean monk seal (*Monachus mona-chus*), the most rarely occurring pinniped worldwide, ranks among the most endangered marine mammal species. A few breeding colonies remain along the shores of Greece, Turkey, and Cyprus as well as in Atlantic waters close to Cabo Blanco, Mauritania, and Madeira (1).

Monk seals are deemed to be officially extinct in many countries, including Italy. A monk seal pup was found alive along the southern Adriatic coast of Italy; it died after rehabilitation attempts. We performed a detailed necropsy on January 28, 2020, within 12 hours after death. Postmortem examination confirmed the animal was a female weaning pup; it had a poor body condition score. During necropsy, we collected samples from the animal's brain, spinal cord, lungs, liver, kidneys, lymph nodes, spleen, intestine, muscles, and tonsils for biomolecular analyses against viral and nonviral pathogens, with special emphasis on cetacean morbillivirus (CeMV) (2,3) and Toxoplasma gondii (4) (Appendix, https://wwwnc.cdc.gov/EID/ article/27/4/20-4131-App1.pdf). We fixed all the tissue samples promptly in 10% neutral buffered formalin and routinely processed them for conventional histology and for morbillivirus and T. gondii immunohistochemistry. We used a commercially available monoclonal antibody against canine distemper virus (CDV) nucleoprotein (Veterinary Medical Research and Development, https://vmrd.com) and a rabbit polyclonal antibody against T. gondii (MyBioSource, https://www.mybiosource.com) (5,6).

We found extensive multifocal brain hemorrhages, most likely caused by a severe arteritis that also involved major cardiac vessels. The brain showed a multifocal, severe, nonsuppurative meningoencephalitis, closely associated with extensive and multifocal hemorrhages. We detected a diffuse, bilateral, chronic, and moderate interstitial pneumonia associated with a marked bronchiolar epithelial hyperplasia; we observed positive immunohistochemistry labeling for morbilliviral antigen within hyperplastic epithelial cells (Figure). Round, variably sized protozoan cysts positively stained with the T. gondii antibody were visible in the lung, within myocardial inflammatory foci, and in the tunica media of the aorta and pulmonary vessels. Lymphoid tissues exhibited a widespread and severe immune cell depletion.

Through biomolecular analyses (2,3), we detected CeMV genetic fragments in brain, lung, and spleen tissues preserved in RNAlater solution (Thermo-Fisher, https://www.thermofisher.com) and frozen lung tissue. Fragments showed a strong homology with a CeMV isolate (complete genome GenBank