



Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.



Brief Report

Middle East respiratory syndrome coronavirus on inanimate surfaces: A risk for health care transmission



Raymond M. Khan MD ^{a,*}, Hasan M. Al-Dorzi MD ^a, Sameera Al Johani MD ^b,
Hanan H. Balkhy MD ^c, Thamer H. Alenazi MD ^d, Salim Baharoon MD ^a,
Yaseen M. Arabi MD ^a

^a Intensive Care Department, King Saud bin Abdulaziz University for Health Sciences, King Abdullah International Medical Research Center, Riyadh, Saudi Arabia

^b Pathology and Laboratory Medicine Department, King Saud bin Abdulaziz University for Health Sciences, King Abdullah International Medical Research Center, Riyadh, Saudi Arabia

^c Infection Control Department, King Saud bin Abdulaziz University for Health Sciences, King Abdullah International Medical Research Center, Riyadh, Saudi Arabia

^d Infectious Diseases, King Saud bin Abdulaziz University for Health Sciences, King Abdullah International Medical Research Center, Riyadh, Saudi Arabia

Key Words:

Environmental cleaning
environmental contamination
infection control
Middle East respiratory syndrome
coronavirus (MERS-CoV)
Saudi Arabia

The Middle East Respiratory syndrome coronavirus (MERS-CoV) has been responsible for multiple health care-associated outbreaks. We investigated whether high-touch surfaces in 3 rooms of laboratory-confirmed MERS-CoV patients were contaminated with MERS-CoV RNA. We found 2 out of 51 surfaces were contaminated with MERS-CoV viral genetic material. Hence, environmental contamination may be a potential source of health care transmission and outbreaks. Meticulous environmental cleaning may be important in preventing transmission within the health care setting.

© 2016 Association for Professionals in Infection Control and Epidemiology, Inc. Published by Elsevier Inc. All rights reserved.

In September 2012, the Middle East respiratory syndrome coronavirus (MERS-CoV) was identified from a patient in Saudi Arabia. As of March 29, 2016, the World Health Organization reported 1,698 laboratory-confirmed MERS cases in 26 countries, with 609 deaths (36%).¹ In its most recent report, the Centers for Disease

Control and Prevention has stressed the great importance of personal protective equipment (PPE), source control, and environmental infection control measures to help eliminate the threat of health care-associated outbreaks.²

Most health care-associated MERS-CoV outbreaks has occurred in Saudi Arabia. Although the precise mechanism of human-to-human transmission has not been elucidated, MERS-CoV can be recovered from plastic surfaces after 48 hours at 20°C and 40% relative humidity (RH), and the virus is viable for 8 hours at 30°C and 80% RH and for 24 hours at 30°C and 30% RH.³ Further, data from the South Korean outbreak (May 2015) demonstrated that several environmental surfaces frequently touched by laboratory-confirmed MERS patients and health care workers were contaminated by MERS-CoV.⁴ Additionally, viral shedding was detected by viral cultures from respiratory secretions up to 25 days postdisease onset.⁴

Although MERS-CoV was isolated from numerous high-touch surfaces in 2 Korean hospitals affected by MERS outbreak,⁴ such data are lacking in the Middle East. Therefore, the objective of this study was to examine the extent of environmental contamination with MERS-CoV during an outbreak in a Saudi hospital.

MATERIALS AND METHODS

The study was performed in the intensive care unit (ICU) at King Abdul-Aziz Medical City, Riyadh, during a MERS-CoV outbreak from

* Address correspondence to Raymond M. Khan, MD, Intensive Care Department, College of Medicine, King Saud bin Abdulaziz University for Health Sciences, King Abdullah International Medical Research Center, PO Box 22490, Mail code 1425, Riyadh, Saudi Arabia.

E-mail address: raymondkhan@yahoo.com (R.M. Khan).

Conflicts of Interest: None to report.

Author Contributions: R.M.K.: Conception, acquisition of data, design, analytical plan, drafting of the manuscript and critical revision of the manuscript for important intellectual content, and approval of the final version to be published. H.M.A.: Conception, drafting of the manuscript and critical revision of the manuscript for important intellectual content, and approval of the final version to be published. S.A.: Conception, molecular analysis and critical revision of the manuscript for important intellectual content, and approval of the final version to be published. H.H.B.: Conception, critical revision of the manuscript for important intellectual content, and approval of the final version to be published. T.H.A.: Conception, critical revision of the manuscript for important intellectual content, and approval of the final version to be published. S.B.: Conception, and critical revision of the manuscript for important intellectual content, approval of the final version to be published. Y.M.A.: Conception, design, analytical plan, drafting of the manuscript and critical revision of the manuscript for important intellectual content, and approval of the final version to be published. All authors agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the article are appropriately investigated and resolved.

September 1-October 5, 2015. The ICU had strict environmental cleaning policies, which included cleaning the rooms at least twice daily using ammonium-based disinfectant and chlorine solution 1:10 or 5,000 ppm, having a checklist, and frequent inspection using fluorescent light or culturing of high-touch areas.

Table 1

Characteristic, physiologic, and laboratory variables for the patients in the rooms during environmental sampling

Variables	Patients		
	A	B	C
Age, y	35	85	30
Sex	Female	Male	Male
Body mass index, kg/m ²	28.7	24.9	37.3
APACHE II score	18	31	15
Time in room before environmental sampling, d	8	16	4
Time from last positive MERS-CoV to environmental sampling, h	24	24	72
MERS-CoV, C _p /C _T	E = 17 O = 18	E = 20 O = 20	E = 34 O = 35
FiO ₂ , %	55	30	40
Tidal volume, mL	250	380	450
PEEP, cm H ₂ O	16	5	8
PaO ₂ /FiO ₂ ratio	152.6	269.3	166.8
Leukocyte count, ×10 ⁹ cells/L	14.6	16.7	11.6
Platelet count, ×10 ⁹ cells/L	345	89	286
Lactate level, mmol/L	1.03	2.42	0.58
Creatinine concentration, μmol/L	65	117	399
Bilirubin level, μmol/L	13.3	31.9	12.7
AST level, U/L	41	879	39
ALT level, U/L	36	808	20

NOTE. C_p or C_T value is the cycle at which fluorescence achieves a defined threshold. The number of cycles needed for the amplification-associated fluorescence to reach a specific threshold level of detection (C_T or C_p value) is inversely correlated to the amount of nucleic acid that was in the original sample. C_T <29 is a strong positive reaction indicative of abundant target nucleic acid in the sample; C_T of 30-37 is a positive reaction indicative of moderate amounts of target nucleic acid, and C_T of 38-40 is a weak reaction indicative of minimal amounts of target nucleic acid.

ALT, alanine aminotransferase; APACHE II, Acute Physiology and Chronic Health Evaluation II; AST, aspartate aminotransferase; C_p, crossing point; C_T, threshold cycle; E, E-protein gene (upstream of the envelope gene); FiO₂, fraction of inspired oxygen; MERS-CoV, Middle East respiratory syndrome coronavirus; O, open reading frame 1b (ORF 1b); PaO₂/FiO₂, arterial oxygen partial pressure to fractional inspired oxygen; PEEP, positive end expiratory pressure.

Table 2

Fomites and different isolation reagents

Fomites	Room A (n = 51)			Room B (n = 51)			Room C (n = 51)		
	UTM Swab	1/4LR	PBS	UTM Swab	1/4LR	PBS	UTM Swab	1/4LR	PBS
Inside ICU room									
1 Bedrails 1 (head)	-	-	-	-	-	-	-	-	-
2 Bedrails 2 (side)	-	-	-	-	-	-	-	-	-
3 Vent	-	-	-	-	-	-	-	-	-
4 Vent tubing	-	-	-	-	-	-	-	-	-
5 Sink	-	-	-	-	-	-	-	-	-
6 Garbage bins	-	-	-	-	-	-	-	-	-
7 Monitors	-	-	-	-	-	-	-	-	-
8 Intravenous poles	-	-	-	-	-	-	-	-	-
9 Intravenous pumps	-	-	-	-	-	-	-	-	-
10 Telephone	-	-	-	-	-	-	-	-	-
11 Door knob	-	-	-	-	-	-	-	-	+
12 Floor	-	-	-	-	-	-	-	-	-
13 Drapes-blinds	-	-	-	-	-	-	-	-	-
14 Air vent	-	-	-	-	-	-	-	-	-
15 Surgical boom shelf	-	-	-	-	-	-	-	-	+
Outside ICU room									
16 Keyboards (computer)	-	-	-	-	-	-	-	-	-
17 Chart	-	-	-	-	-	-	-	-	-

NOTE. The results of real-time polymerase chain reaction for Middle East respiratory syndrome coronavirus viral RNA from various ICU environmental surfaces and eluents (solvents) used.

ICU, intensive care unit; PBS, phosphate buffer solution; UTM, universal transport medium; 1/4LR, one-quarter lactate ringers; -, negative test result in the room; +, positive test result in the room.

Three negative-pressure rooms of laboratory-confirmed MERS patients (A, B, and C) were selected for this study (Table 1). The room temperature was 20.0°C-25.0°C, and RH was 30%-40%. The air exchange rate was 12 per hour, and the pressure gradient between the room and its anteroom ranged from 2.5-12.5 Pa. Sixteen high-touch surfaces were evaluated (Table 2): 14 in the patients' room (bedrails, mechanical ventilator, ventilator tubing, sink, garbage bin, monitor, intravenous poles, intravenous pumps, telephone, door knobs, floor, drapes-blinds, air conditioning vent, and shelf of the surgical boom) and 2 outside (computer and medical chart). Environmental samples were collected as described by Julian et al.⁵ Briefly, a sterile swab premoistened with viral transport media was used to swab each surface (at least 10 cm²) horizontally, vertically, and diagonally for 30 seconds. This procedure was repeated using eluents: 1/4 lactated ringer solution and phosphate buffer solution (PBS). Virus detection was performed using specific real-time reverse-transcription polymerase chain reaction (PCR) assays for the upstream of the envelope gene and the open reading frame 1A. Positive tests were reported as the cycle threshold value for both upstream of the envelope gene (E) and open reading (O) frame 1A.

RESULTS

The demographic for the patients are summarized in Table 1. All 3 laboratory-confirmed MERS patients were on mechanical ventilators, with an average PaO₂/FiO₂ ratio of 196. The mean ICU length of stay and time from last positive tracheal aspirate for MERS-CoV RNA to environmental sampling were 9.3 days and 40 hours, respectively.

Sixteen surfaces were evaluated in each of the 3 ICU rooms, with 153 environmental samples processed (Table 2). MERS-CoV viral nucleic acid was detected in 2 specimens (PBS as eluent) taken from room C. The positive surfaces were the shelf of the surgical boom and the inside door knob, with cycle threshold values E = 33/O = 31 (positive) and E = 36/O = 0 (intermediate), respectively.

DISCUSSION

Our study revealed that MERS-CoV viral RNA was isolated from the environmental surfaces of MERS patients.

Currently, much remains uncertain about the transmission mechanism responsible for MERS nosocomial outbreaks. It was postulated from the outbreak in Al-Hasa, Saudi Arabia, in May–June 2012 that respiratory droplet and airborne transmission during aerosol-generating procedures were the most likely transmission modes.⁶ However, genetic data from a cluster in Hafr Al-Batin, Saudi Arabia, showed that direct person-to-person contact could not account for all of their cases,⁷ therefore raising the likelihood of an alternate transmission mechanism. Studies on kinetics and patterns of viral excretion indicate that MERS-CoV RNA was isolated from urine and feces 13 and 16 days, respectively, after initial symptoms.⁸ Viral shedding from respiratory aspirates may persist up to 33 days after illness onset.⁹ Prolonged viral shedding^{8,9} and survival on surfaces for 48 hours³ make it difficult to ignore contaminated environmental surfaces as a potential etiology of hospital outbreaks.

The rate of detecting MERS-CoV in our environmental samples was low (1.3%) compared with recently published data (PCR positive = 20.3%; culture positive = 4.0%),⁴ but the current methods for isolating viruses from the environmental surfaces are not optimal.⁵ Based on reported methodologies, we used a polyester swab, 1/4 lactated ringer solution,⁵ PBS⁵ and viral transport media⁴ because they seem to give the best yield for isolating viruses from fomites. However, we did screening at the tail-end of our outbreak when the patients' viral load might have been low and our infection control practices might have been optimal. Additionally, MERS patients were managed in our ICU since 2013 and were usually cohorted in 1 unit where the staff became very meticulous about PPE use and environmental cleaning. Moreover, fairly weak disinfectants, such as povidone iodine, have a rapid virucidal activity (reduction in virus titer by $\geq 4 \log_{10}$) against MERS-CoV, with an exposure time of just 15 seconds.¹⁰ Further, Leclercq et al demonstrated that at relatively low temperatures of 56°C, only 25 minutes was needed to reduce the initial titer by 4 \log_{10} , while at 65°C virucidity dropped significantly to 1 minute.¹¹ This sensitivity to weak disinfectants could explain why our stringent environmental cleaning policies may have attenuated the recovery of viral genetic material on fomites within the patients' rooms.

CONCLUSIONS

Our finding of MERS-CoV RNA on environmental samples within our ICU shows that the viral material may contaminate fomites and can be a theoretical cause of nosocomial infections. However, we did not use viral cultures; therefore, we do not know if the positive PCRs correlate with live viruses or infectivity. Despite this, we believe that in addition to proper hand hygiene and correct PPE donning and doffing, meticulous environmental cleaning is of paramount importance to eliminate health care outbreaks.

References

1. World Health Organization. Middle East respiratory syndrome coronavirus (MERS-CoV). 2015. Available from: <http://www.who.int/emergencies/mers-cov/en/>. Accessed December 12, 2015.
2. Williams HA, Dunville RL, Gerber SI, Erdman DD, Pesik N, Kuhar D, et al. CDC's early response to a novel viral disease, middle east respiratory syndrome coronavirus (MERS-CoV), September 2012–May 2014. *Public Health Rep* 2015;130:307–17.
3. van Doremalen N, Bushmaker T, Munster VJ. Stability of Middle East respiratory syndrome coronavirus (MERS-CoV) under different environmental conditions. *Euro Surveill* 2013;18.
4. Bin SY, Heo JY, Song MS, Lee J, Kim EH, Park SJ et al. Environmental contamination and viral shedding in MERS patients during MERS-CoV outbreak in South Korea. *Clin Infect Dis* 2016;62:755–60.
5. Julian TR, Tamayo FJ, Leckie JO, Boehm AB. Comparison of surface sampling methods for virus recovery from fomites. *Appl Environ Microbiol* 2011;77:6918–25.
6. Assiri A, McGeer A, Perl TM, Price CS, Al Rabeeah AA, Cummings DA, et al. Hospital outbreak of Middle East respiratory syndrome coronavirus. *N Engl J Med* 2013;369:407–16.
7. Memish ZA, Cotten M, Watson SJ, Kellam P, Zumla A, Alhakeem RF, et al. Community case clusters of Middle East respiratory syndrome coronavirus in Hafr Al-Batin, Kingdom of Saudi Arabia: a descriptive genomic study. *Int J Infect Dis* 2014;23:63–8.
8. Drosten C, Seilmaier M, Corman VM, Hartmann W, Scheible G, Sack S, et al. Clinical features and virological analysis of a case of Middle East respiratory syndrome coronavirus infection. *Lancet Infect Dis* 2013;13:745–51.
9. Poissy J, Goffard A, Parmentier-Decrucq E, Favory R, Kaut M, Kipnis E, et al. Kinetics and pattern of viral excretion in biological specimens of two MERS-CoV cases. *J Clin Virol* 2014;61:275–8.
10. Eggers M, Eickmann M, Zorn J. Rapid and effective virucidal activity of povidone-iodine products against middle east respiratory syndrome coronavirus (MERS-CoV) and modified vaccinia virus ankara (MVA). *Infect Dis Ther* 2015;4:491–501.
11. Leclercq I, Batejat C, Burguiere AM, Manuguerra JC. Heat inactivation of the Middle East respiratory syndrome coronavirus. *Influenza Other Respir Viruses* 2014;8:585–6.