

Clinical Infectious Diseases

Brief Report

Diagnosis and Management of First Case of COVID-19 in Canada: Lessons applied from SARS

Xavier Marchand-Senécal MD, FRCPC¹, Rob Kozak PhD¹, Samira Mubareka MD, FRCPC^{1,2},
Natasha Salt¹, Jonathan B Gubbay MBBS, MMedSc, FRCPC, FRACP,³ Alireza Eshaghi PhD³,
Vanessa Allen MD FRCPC^{2,3}, Yan Li PhD⁴, Natalie Bastien PhD⁴, Matthew Gilmour PhD⁴,
Omar Ozaldin MSc⁵, Jerome A. Leis MD, MSc, FRCPC^{1,6}

1. Sunnybrook Health Sciences Centre, Toronto, ON, Canada
2. Department of Laboratory Medicine and Pathobiology, University of Toronto, Toronto, ON, Canada
3. Public Health Ontario Laboratory, Toronto, ON, Canada
4. National Microbiology Laboratory, Public Health Agency of Canada, Winnipeg, MB, Canada
5. Communicable Disease Control, Toronto Public Health, Toronto, ON, Canada
6. Division of Infectious Diseases, Department of Medicine, University of Toronto, Toronto, ON, Canada

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Corresponding Author:

Jerome A. Leis, MD, MSc, FRCPC

Medical Director, Infection Prevention and Control, Sunnybrook Health Sciences Centre

Staff Physician, Divisions of Infectious Diseases and General Internal Medicine, Sunnybrook Health Sciences Centre

Associate Professor, Department of Medicine, University of Toronto

Jerome.Leis@sunnybrook.ca

+1-416-480-4243

Abstract (unstructured)

We report diagnosis and management of the first laboratory-confirmed case of coronavirus disease 2019 (COVID-19) hospitalized in Toronto, Canada. No healthcare-associated transmission occurred. In the face of a potential pandemic of COVID-19, we suggest sustainable and scalable control measures developed based on lessons learned from SARS.

Introduction

On December 31st 2019, China alerted the World Health Organization about a cluster of cases of pneumonia of unknown origin in Wuhan, Hubei, China (1). One week later, a novel coronavirus (provisional name: 2019-nCoV; definitive name: SARS-CoV-2) was identified as the causative agent of the coronavirus disease 2019 (COVID-19). As of February 28th, 2020, the outbreak has spread due to human-to-human transmission throughout China, and to over fifty countries (2).

While the majority of transmission has occurred in community settings, super-spreading events in healthcare settings have already been described. In a Wuhan hospital, a nosocomial acquisition is suspected among 29% of healthcare workers (HCW) and 12.3% of the 138 hospitalized patients infected (2). A patient admitted with unrecognized symptoms of the infection was reported to transmit the virus to more than 10 HCWs.

Our hospital in Toronto, Canada, has past experience caring for patients with SARS-CoV-1 where the majority of cases were healthcare-associated (3). We now report the first imported case of COVID-19 hospitalized in Canada while emphasising the control measures applied based on lessons learned from SARS.

Case presentation

On January 23rd 2020, a 56-year old man presented to our hospital Emergency Department in Toronto with a new onset of fever and non-productive cough following return from Wuhan, China the day prior (4). Based on high suspicion for COVID-19, throat and mid-turbinate swabs (FLOQ Swab by Copan, with UTM transport media) were collected. On clinical assessment, the patient remained well without ever requiring supplemental oxygen. The chest-x-ray confirmed the presence of pneumonia (4). The patient was admitted to hospital for close observation and supportive management, considering early reports of respiratory failure due to COVID-19 (1).

The specimens collected were negative for influenza A and B, parainfluenza, respiratory syncytial virus, adenovirus and human metapneumovirus. Coronavirus was detected in both mid-turbinate and throat swab by pan-coronavirus RNA dependent RNA polymerase (RdRp) PCR and confirmed as SARS-CoV-2 by sequencing (5).

On the inpatient unit, the patient remained stable. Symptom duration and viral studies are summarized in Figure 1. Following resolution of fever for 48-hours, the patient was discharged home on day 8 of hospital stay. Public health followed up with his wife at home who had cough only. She was also confirmed to have COVID-19. Both patients recovered fully with home isolation which was discontinued on February 20th (day X) following two-negative swabs within 24-hours.

Diagnostic testing

All specimens were received at Public Health Ontario Laboratory and eluted in universal transport media (Copan, Italia, Brescia, IT)). Total nucleic acid extraction of all primary specimens was performed using a NucliSens easyMAG extraction system instrument (bioMérieux Canada Inc., Québec, Canada), according to the manufacturer's instructions. Reverse transcription and endpoint PCR was performed using hemi nested pan-CoV primers targeting the RdRp (5), and additional primers from this protocol adapted to be SARS-CoV-2 specific. Once validated for clinical testing, specimens were also tested using a published real-time PCR protocol targeting the envelope (E) and nucleoprotein (N) genes (6). Presumptive confirmation was obtained after detection of two gene targets by real-time PCR, or PCR and Sanger sequencing of the RdRp gene. Confirmation was conducted by the National Microbiology Laboratory (NML) of the Public Health Agency of Canada by using a Real-time PCR assay targeting the N gene (developed at NML). Conventional RT-PCR assay targeting the RdRp was

modified from elsewhere (7) and ORF3a (developed at NML) was also performed. Nucleotide analysis of the partial gene sequences of RdRp and ORF3a amplicons showed that they were more closely related to COVID-2019.

Infection prevention and control management

Based on presence of respiratory symptoms and travel history elicited at triage, this patient was immediately placed in private negative pressure room with airborne-droplet-contact precautions as per our screening protocol for novel high-consequence pathogens. The personal protective equipment (PPE) used included a long-sleeved gown, a pair of gloves, a fit-tested N95 respirator and a face-shield. The patient was cared for by the usual medical and nursing teams in the emergency department and on the unit. Since the PPE used is familiar to our hospital staff and no additional components used (eg. hood or powered air-purifying respirator), no specific training was undertaken other than audits to ensure correct donning and doffing technique.

Additional administrative control measures were put in place given the inherent risks of a novel high-consequence pathogen. First, we maintained a log of each healthcare worker involved in the patient's care. Second, we limited the number of individuals entering the isolation room to the most essential healthcare workers. For example, no visitors were allowed for the patient who were instead provided the option of using telephone communication. Disposable trays were used to avoid the need to enter and collect food items. Third, all equipment including vital signs machine were dedicated to this patient only. Finally, the critical care and code blue team were alerted on admission of the patient location and the need to abide to hospital policy for minimizing the risks of aerosol-generating medical procedures (AGMP). It was agreed upon by the clinical teams that should the patient require increasing supplemental oxygen, there would be

a low threshold to intubate using controlled and protected procedures including use of paralytics, video laryngoscopy, and PPE including N95 masks (8).

Following the patient's discharge, terminal cleaning of the patient's room was completed twice using 0.5% hydrogen peroxide. An IPAC professional and an Environmental Services Supervisor audited both stages of the room cleaning. The list of healthcare workers involved in the patient's care was shared with occupational health and safety, who were to immediately notify IPAC of any new reported specific or non-specific influenza-like illness within 14-days of discharge. By February 28th 2020, there were no probable or confirmed cases of COVID-19 among other patients or healthcare workers in the facility.

Discussion

We report a case of COVID-19 which was identified immediately and using transmission-based precautions, monitoring and administrative controls was safely cared for with no healthcare associated transmission.

While we expect many more cases of COVID-19 as this epidemic evolves, this first case highlights significant progress made since the SARS outbreak in our own hospital back in 2003. There were 375 cases of SARS throughout Toronto during this prior outbreak of which 271 resulted from healthcare-associated transmission (3). Following this event, additional resources in IPAC and public health were instituted in Canada to improve preparedness for managing high-consequence pathogens (9, 10). The key differences in preparedness and capacities at our hospital between SARS and COVID-19 epidemics are summarized in Table 1.

Additional lessons learned were applied in the management of this patient with COVID-19. First, we did not rely on a specific dedicated team, but instead built on IPAC capacities already in place among our usual staffing model that includes baseline preparedness of

management of novel high-consequence pathogens. This approach differed from SARS as well as Ebola in 2014, where many hospitals developed dedicated teams to care for these patients, which required significant investments in human resources and training (11). In the aftermath of those epidemics, this approach was difficult to sustain for hospitals resulting in loss of preparedness. Second, what allows our hospital to maintain baseline preparedness for novel high-consequence pathogens is that our standard PPE for managing these patients only includes components that are already familiar to our HCWs. This decision was made based on evidence demonstrating that additional PPE complexity is associated with risks of self-contamination during doffing (12). This approach is also consistent with prior evidence from SARS-CoV-1 suggesting that transmission occurs by the droplet or contact route (13). Finally, we instituted administrative controls for preventing super-spreading events around AGMP (3, 8). While our patient did not end up requiring any AGMP, coordinated communication and planning was in place to ensure these would be performed in a controlled fashion to minimize HCW risk.

When to discharge this patient was a decision that required us to carefully balance the risk that a stable patient who is yet early in the COVID-19 illness with potential to require medical intervention (2), against the ongoing risk of HCW exposure with each additional day of inpatient care. On the other hand, promoting early discharge of this patient might have resulted in transfer back to hospital if he deteriorated leading to additional risk of exposure during transport. We urgently need to develop evidence-based risk stratification of patients with COVID-19 to determine those best managed at home rather than in hospital as part of our strategy to keeping the number of exposures to a minimum.

How long patients with COVID-19 remain infectious represents a significant gap in our understanding. Our patient's symptoms lasted 13 days but viral shedding continued until day 28. Current CDC and Ontario's Ministry of Health recommendations are to maintain isolation until 2

negative repeat testing of respiratory tract specimens (14), pending more definitive assessment of bioaerosol studies.

Our report has important limitations. First, it represents the experiences of a single healthcare institution and public health unit around a single case, compared to the large number of patients being concurrently managed in the epicentre of the outbreak in Hubei province, China. Second, our patient remained stable and did not require any AGMP which would have increased the risk of transmission substantially. Finally, our patient presented quite typically and identified based on travel history; however, countries where local transmission has already occurred must rely on syndromic screening alone which may be more challenging given reports that some patients with COVID-19 may present atypically (2).

In the face of an impending pandemic of COVID-19, the diagnosis and management of this first Canadian case provides hope that we can limit the burden of healthcare-associated infection seen during SARS. We call for scalable and sustainable preparedness for COVID-19 and beyond.

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Table 1. Differences between 2003 and 2020 in preparedness for a novel coronavirus at a large academic hospital in Toronto, Canada.

| | 2003: SARS in Toronto* | 2020: COVID-19 Toronto case |
|--|--|---|
| Public health structures and infrastructures | | |
| Adequate funding and human resources | No | Yes |
| Protocols for information sharing among different levels of government | No | Yes |
| Link between public health and hospitals | Weak, Fragmented, Uncoordinated | Coordination and information sharing present |
| Rapid and accurate diagnostic testing | No | Yes |
| IPAC program structure and related hospital program | | |
| ICP staffing level | Understaffed 3 ICP for 1257 total beds (0.23 ICP/100 beds) | Adequate 13 ICP for 1355 total beds (0.96 ICP/100 beds) |
| ICP certification (Certification Board of Infection Control and Epidemiology) | Not universal | Required |
| Occupational Health & Safety | Disconnected from IPAC | Coordinated with IPAC |
| IPAC administrative controls | | |
| Syndromic triage in ED | No | Yes |
| Febrile respiratory illness surveillance | No | Yes |
| Isolation of all patients with acute respiratory symptoms | No | Yes Droplet & contact precaution; If COVID-19 suspected or confirmed: airborne, droplet & contact |
| Awareness of super-spreading events and individuals | No | Yes |
| Minimizing AGMP and Protected intubation policies | No | Yes |
| Hand hygiene program | No | Yes |
| Healthy Workplace Policy (Work restrictions for HCW with acute infectious symptoms) | No | Yes |
| Presence of a Pandemic plan | No | Yes |
| Engineering and environmental controls | | |
| Number of airborne infection isolation room | 20 (0 in ED) | 46 (8 in ED) |
| ED infrastructure | Shared air system; No protective barrier at triage | Isolated air system with negative pressure in each zone; Protective barrier at triage |
| Terminal disinfection completed twice at patient discharge for high-consequence pathogen | No | Yes |
| Personal protective equipment (PPE) | | |
| Regular N95 fit-testing of HCW | No | Yes |
| Clear recommendation on PPE for any novel high-consequence pathogen | No | Yes |

* Based on historical local data and the Naylor report (10); HCW=healthcare worker; IPAC=Infection Prevention and Control; AGMP= Aerosol Generating Medical Procedure; ED=Emergency Department; PPE=personal protective equipment; ICP=Infection Prevention and Control Professional.

Figure 1. Clinical trajectory and viral shedding of first confirmed case of Coronavirus Disease 2019. Timeline chart illustrates the symptoms duration and the cycle threshold (Ct) values of the real-time reverse-transcriptase-polymerase-chain-reaction (RT-PCR) evolution. Lower Ct values indicate higher viral loads.