## Hospital infection and COVID-19: Do not put all your eggs on the "swab" tests

### Francesco Chirico MD<sup>1,2</sup> <sup>(i)</sup>, Gabriella Nucera MD<sup>3,4</sup> and Nicola Magnavita MD<sup>5,6</sup>

<sup>1</sup>Postgraduate School of Occupational Medicine, Università Cattolica del Sacro Cuore, Rome, Italy, <sup>2</sup>Health Service Department, State Police, Ministry of Interior, Milan, Italy, <sup>3</sup>Faculty of Nursing, University of Milan, Italy, <sup>4</sup>Fatebenefratelli Sacco, FatebeneFratelli Hospital, Milan, Italy, <sup>5</sup>Postgraduate School of Occupational Medicine, Università Cattolica del Sacro Cuore, Rome, Italy and <sup>6</sup>Department of Woman/Child and Public Health, Fondazione Policlinico "A.Gemelli" IRCCS, Rome, Italy

To the Editor—In healthcare settings, including long-term care facilities, hospital administrators have a legal obligation to set up a risk assessment strategy to carry out effective prevention and control measures during the management of suspected and confirmed cases of COVID-19 infection.<sup>1</sup> Hospitalized inpatients and residents in care homes are often elderly and immune-depressed patients with comorbidities; thus, they are at high risk of infection and mortality. Special attention and efforts to protect or reduce transmission should be also applied in healthcare providers because depletion of the healthcare workforce not only will affect health care but also will contribute to the spread of the outside hospitals.

According to European Centres for Disease Control and Prevention (EU-CDC) guidelines,<sup>1</sup> each hospital should constitute a 'COVID-19 preparedness and response committee' and should prepare a specific plan, including a number of administrative and structural measures for patient and healthcare management. Undoubtedly, the most important measure in reding the likelihood of nosocomial infection is early isolation of patients with COVID-19, or at least maintaining a safe distance between those who are awaiting diagnosis. However, a number of recent studies showed that patients with mild or nonspecific symptoms can escape isolation and thus introduce SARS-CoV-2 into hospitals, leading to clusters of nosocomial infections.<sup>2</sup>

To minimize the risk of spreading, mass testing with nasopharyngeal and oropharyngeal (NP/OP) swab of all patients has been proposed,<sup>2,3</sup> associated with mass testing of both symptomatic and asymptomatic healthcare workers.<sup>4</sup> Even the use of these expensive and demanding mass strategies, however, cannot be considered a measure of absolute guarantee.

Indeed, Xie et al<sup>5</sup> observed typical COVID-19 chest lesions via computed tomography (CT) scans in 5 patients with a negative or weakly positive swab test (RT-PCR test). Another patient with a chest X-ray showing interstitial pneumonia but with a negative RT-PCR test was reported by Winichakoon et al.<sup>6</sup> Kumar et al<sup>7</sup> reported the case of a patient with pneumonia and negative nasopharingeal swab who tested positive some days later with a bronchial lavage sample. Bandirali et al<sup>8</sup> found that asymptomatic or minimally symptomatic patients may have abnormalities in chest x-rays after 14 days of quarantine, with a sensitivity of RT-PCR testing of 59%.<sup>8</sup>

Author for correspondence: Prof Francesco Chirico, E-mail: medlavchirico@gmail.com

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Cao et al<sup>9</sup> observed that patients with negative to RT-PCR tests may have specific IgG and/or IgM for SARS-CoV-2 at recovery stage.<sup>9</sup> In reality, the sensitivity of an NP/OP swab in the course of disease ranges between 42% and 71%<sup>6</sup> and depends on sampling technique, timing within the clinical course of COVID-19, and viral loads detected in the swab.<sup>6</sup>

In conclusion, given the fact that negative NP/OP swabs do not rule out COVID-19 diagnosis, we propose that all the patients hospitalized with pneumonia be subjected to swab obtained by deep tracheal aspirate, which has a lower risk of aerosolization.<sup>10</sup> We further recommend that suspected infection be checked with a combination of repeated RT-qPCR tests and chest CT scan. All patients hospitalized without respiratory symptoms should also be checked with repeated RT-qPCR tests and chest X ray before admission in hospital wards.

Moreover, healthcare providers should be tested regularly with serological test and swabs and symptom monitoring. Finally, a policy of universal masking and eye shielding for all healthcare providers involved in direct patient care is needed.

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# The value of hospital personnel serological screening in an integrated COVID-19 infection prevention and control strategy

Filippo Quattrone MD<sup>1,2,3,a</sup> (), Marco Vabanesi MD<sup>4,a</sup> (), Alice Borghini MD<sup>5</sup>, Giuseppe De Vito MD<sup>6</sup>,

### Michele Emdin PhD, MD<sup>1,3</sup> and Claudio Passino MD<sup>1,3</sup>

<sup>1</sup>Institute of Life Sciences, Scuola Superiore Sant'Anna, Pisa, Italy, <sup>2</sup>Department of Translational Research and New Technologies in Medicine and Surgery, University of Pisa, Pisa, Italy, <sup>3</sup>Fondazione Toscana Gabriele Monasterio, Pisa, Italy, <sup>4</sup>Department of Neurology, San Raffaele Scientific Institute and University Hospital, Milan, Italy, <sup>5</sup>Institute of Management, Scuola Superiore Sant'Anna, Pisa, Italy and <sup>6</sup>Fondazione Stella Maris, Pisa, Italy

To the Editor—Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is highly infectious in healthcare-related settings, both among patients and healthcare workers (HCWs).<sup>1</sup> Hospital personnel have shown an increased risk of coronavirus disease 2019 (COVID-19) compared to the general population, possibly associated with repeated exposures and, in the current emergency context, frequent lack of adequate personal protective equipment (PPE). The spread of SARS-CoV-2 has also been dramatically efficient in long-term care facilities (LTCFs), where the combination of asymptomatic or paucisymptomatic occupational carriers and a highly fragile elderly population have produced numerous outbreaks, greatly contributing to the total burden of COVID-19–related deaths.<sup>2</sup>

An integrated COVID-19 infection and prevention control (IPC) strategy must be promptly adopted by healthcare facilities to prevent further outbreaks. This strategy should involve HCWs, patients, visitors, and support personnel (eg, administrative and ancillary services workers) due to their reported role in nosocomial outbreaks.<sup>3</sup> Current recommendations<sup>4</sup> include the adoption of general IPC measures (eg, hand hygiene, physical distancing, universal use of surgical masks, and triage at entrance for fever, respiratory symptoms, and history of exposure to the virus), environmental measures (eg, enhanced surface cleaning, control of indoor air, proper linen, laundry, and waste management), administrative measures (eg, limiting visitor access and promoting remote work and telemedicine), and patient management measures (eg, dedicated pathways and isolation wards for patients with fever and respiratory symptoms, consistent use of adequate PPE, and universal SARS-CoV-2 screening for inpatients).

Systematic screening of HCWs and support personnel plays a key role in limiting the intrahospital spread of SARS-CoV-2. The most described approach is screening with viral genome real-time polymerase chain reaction (RT-PCR) on nasopharyngeal swabs. RT-PCR tests should be offered whenever an HCW presents with any symptom suggestive of COVID-19; initial screening is warranted for all new employees. Universal RT-PCR screening protocols<sup>1,5</sup> have shown promising results. A limitation of this approach is the short-term RT-PCR positivity, with consequent need of repeated testing, sustained usage of intensive laboratory resources, exposure risks for the operators involved in screening, and possibly reduced compliance with repeated swab testing. To overcome some defects of traditional PCR-based testing, novel kits for point-of-care rapid PCR testing are currently being developed, with as yet uncertain yield.

Antibody response to SARS-CoV-2 has not been completely characterized; however, from the best available data,<sup>6</sup> it appears that the detection of serum anti-SARS-CoV-2 IgG antibodies with appropriate methods (ie, chemiluminescence enzyme immuno-assay, CLIA) is observable in almost all infected subjects within 20 days from symptom onset. Data on IgM appear less conclusive, and currently, these data do not support the classic sequential IgM–IgG transition; therefore IgM should not be the only target of antibody search.

Many regulatory institutes have assessed rapid, point-of-care antibody tests based on lateral flow immunoassay (LFIA), produced by multiple manufacturers. Although tempting for practical reasons, these tests have not met expectations for use in clinical settings due to unsatisfactory sensitivity and specificity.<sup>7</sup>

Serial serological screening with a validated technique, such as CLIA, could provide a significant contribution to IPC in hospitals and LTCFs, considering its lower cost, easier repeatability, and sustainability in the medium term, compared with swab-based molecular assays. Although serological tests have limited utility in diagnosing individual acute infections, they can inform actions to protect the hospital community. A serum antibody

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Author for correspondence: Filippo Quattrone, E-mail: filippo.quattrone@santannapisa.it <sup>a</sup>Authors of equal contribution.

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