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Letter to the Editor



61

Clinical characteristics and persistence of severe acute respiratory coronavirus virus 2 (SARS-CoV-2) IgG antibodies in 4,607 French Aq1 healthcare workers: Comparison with European countries

4 Chantal Delmas MD¹, Genevieve Plu-Bureau MD, PhD², Etienne Canouï MD³, Luc Mouthon MD, PhD⁴ and

AQ2 Jean-Francois Meritet MD⁵

AQ3 ¹Occupational Health Department, GH Paris Centre – Cochin, APHP, France, ²Epopee Team Inserm U1153 and Medical Gynecology Unit, GH Paris Centre – Cochin
 7 APHP, University of Paris, Paris, France, ³Antimicrobial Stewardship Team, GH Paris Centre – Cochin, APHP, Paris, France, ⁴Internal Medicine Department, GH
 8 Paris Centre – Cochin, APHP, University of Paris, Paris, France and ⁵Virology Department, GH Paris Centre – Cochin APHP, Paris France

9 To the Editor—The safety of healthcare workers (HCWs) is a 10 major challenge for healthcare systems. In the course of a severe 11 acute respiratory coronavirus virus 2 (SARS-CoV-2) infection, 12 immunoglobulin G (IgG) antibodies may be detected after a 13 median of 14–24 days (interquartile range [IQR], 10–18) after 14 onset of symptoms.¹

15 In France, the coronavirus disease 2019 (COVID-19) pandemic 16 reached a peak on April 7, 2020. HCWs had mobility and flexibility 17 inside the Paris Center university hospital, where there was a 18 cluster in the pandemic. We investigated the prevalence of IgG 19 antibodies against SARS-CoV-2 among all HCWs in this hospital. 20 We also sought to determine the correlation between RT-PCR test 21 and serology and to compare our seroprevalence with that of other 22 European countries.

23 From May 14, 2020, to June 17, 2020, all HCWs were asked 24 by the occupational health department to participate in serologic screening. The Abbott-Architect test (Abbott Laboratories, Abbott 25 Park, IL) was used to detect IgG anti-SARS-CoV-2. During blood 26 sampling, clinical information was recorded using a standardized 27 self-questionnaire on presented symptoms, comorbidities, and the 28 29 reverse-transcriptase polymerase chain reaction (RT-PCR) test if 30 one had been previously performed. Blood samples were collected 31 >28 days after the first symptoms from those who were 32 symptomatic.

33 The seroprevalence and 95% confidence interval were esti-34 mated using the Fisher exact method. The t test and the χ^2 test were performed to compare quantitative and qualitative variables, 35 respectively. Simple and multivariate logistic regressions were 36 performed to assess risk and symptoms associated with seropre-37 38 valence respectively. Statistical analyses were performed using SAS software (SAS Institute, Cary, NC). The local institutional 39 40 review board approved this study. All subjects participated volun-41 tarily under pseudonyms.

Author for correspondence: Chantal Delmas, E-mail address: chantal.delmas@aphp.fr Cite this article: Delmas C, et al. (2020). Clinical characteristics and persistence of severe acute respiratory coronavirus virus 2 (SARS-CoV-2) IgG antibodies in 4,607 French healthcare workers: Comparison with European countries. Infection Control & Hospital Epidemiology, https://doi.org/10.1017/ice.2020.1309 Of 5,021 workers present during the study period, 4,607 42 (91.8%) were included in the study. The mean age was 41.8 years 43 (SD, 12.6), and 75% were female. Furthermore, 45% were para-44 medical staff members, 36% were physicians (including medical 45 students), and 19% were in administrative and other professions. 46

Overall, the prevalence of IgG antibodies was 11.5% (95% 47 confidence interval [CI], 10.6-12.4), and it was significantly higher 48 (ie, 13%) for paramedical staff (P = .04). Age and gender did not 49 differ significantly according to seroprevalence. Furthermore, 50 5 clinical symptoms were independently associated with positive 51 serology: asthenia, fever, myalgia, ageusia, and anosmia, for which 52 the highest odd ratio (OR) was observed (OR, 11.1; 95% CI, 53 7.4-16.6). Notably, although anosmia appeared to be the most spe- 54 cific factor, 64.3% of subjects with antibodies did not experience 55 this symptom. The proportion of asymptomatic subjects with a 56 positive serology was 21.4%. When considering comorbidities, 57 positive serology was significantly associated with a lesser preva-58 lence in smokers (OR, 0.41; 95% CI, 0.29-0.58) and a higher preva-59 lence of diabetes (OR, 1.78; 95% CI, 1.04-3.03), 60

Discordance between RT-PCR and serology

In our study, 19.4% of the study participants had had a RT-PCR. 62 Among individuals with negative a RT-PCR, 51 of 662 (7.7%) 63 had detectable SARS-CoV-2 antibodies, whereas 29 of 233 64 (12.4%) of RT-PCR-positive participants also had no detectable 65 antibodies. The former result could be explained either by difficul- 66 ties implementing RT-PCR tests or by the delay between the time of 67 the test and the effective date of infection. For the latter finding, in 68 addition to participants who did not develop antibodies, the time lag 69 between PCR and serology should be mentioned (mean, 64.0 days), 70 which implies that the serology is often realized long after the IgG 71 peak. Indeed, the mean of antibody prevalence in this group 72 (0.28 ± 0.32) was higher than in the negative RT-PCR group 73 $(0.05 \pm 0.08; P < .001)$. More generally, this group with positive 74 RT-PCR and negative antibody tests had specific characteristics: 75 younger age $(38.3 \pm 12.8 \text{ vs } 43.3 \pm 12.4; P = .04)$, more likely a smoker 76 $(31.0\% \text{ vs } 7.4\%; P < 10^{-4})$, and male (37.9% vs 18.1%; 77)P = .01) compared with those with positive RT-PCR and positive 78 serology tests. 79

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121

Country, First Author	No. of Participants	Prevalence %	95% CI	Date of Blood Test	Population Type
Belgium, Blairon ⁶	1,494	1.6	NA	May 25–June 19	4 public hospitals
Belgium, Martin ⁷	326	11.0	NA	April 15– May 18	CHU Saint Pierre, Bruxels
UK, Bampoe ³	200	14.5	9.9–20.1	May 11–June 5	Maternity, London
Germany, Korth ²	316	1.6	NA	March 25– April 21	Essen Hospital, tertiary-care
Germany, Lackermair ⁹	151	2.6	0.8-7.1	April 2–6	Outpatient center, Dachau
Germany, Schmidt ¹	385	2.9	NA	April 20–30	Neurologic clinic
Spain, Garcia-Basteiro ⁴	578	7.6	NA	March 28–April 9	Hospital reference, Barcelona
Denmark, Iversen ⁸	28,792	2.7	2.5–2.9	April 15–23	Capital region
France, Delmas ^a	4,607	11.5	10.6-12.4	May 14–June 17	Paris Center, university hospital

Table 1. Comparison of Seroprevalence IgG in European Countries

Note. CI, confidence interval.

^aPresent study.

80 Comparison with European countries

81 In our literature review, we retained only studies with IgG antibody testing; we excluded those with IgA or IgM serologies. The 11.5% 82 83 prevalence of IgG in our HCWs is similar to the reported prevalences 84 in Belgium or the United Kingdom (Table 1). Different protective 85 measures, date of blood screening, and/or population structure in 86 each country could explain the variation in IgG serology from 87 1.6% reported by Korth et al² up to 14.5% reported by Bampoe 88 et al.³ In our hospital, masks are compulsory, and protective equip-

89 ment has been available since March 17.

90 Of 233 HCWs, 29 (12.4%) were RT-PCR positive with no 91 detectable antibodies. This result parallels that of Garcia-Basteiro 92 et al,⁴ who also reported 15% of individuals with positive 93 RT-PCR and negative serology. A recent study by Patel et al⁵ showed 94 the possibility of decreased antibodies over 60 days, which implies 95 transiently detectable antibodies.

96 Our study has some limitations. During the lockdown period, 97 some HCWs were isolated at home on a case-by-case basis for 98 reasons of severe personal or familial comorbidities. RT-PCR swab 99 tests were conducted at the time of suspected illness only in symp-100 tomatic or in individuals who had had contact with COVID-19 101 patients. Thus, 902 of 4,607 (19.6%) had this test at the time of 102 onset of symptoms.

103 The detection of asymptomatic cases by RT-PCR is essential 104 to isolating or avoiding quarantine of HCWs to prevent risk of 105 contamination for vulnerable patients and to reduce the risk of 106 interprofessional staff-to-staff transmission.

107 To limit virus transmission, we emphasize the necessity of 108 large-scale screening for exposed HCWs, even those who do not 109 present any symptoms. Further investigations are needed to 110 explore negative serology in subjects with positive RT-PCR for 111 understanding population immunity and the potential risks of 112 reinfection and disease in HCWs.

113 **Supplementary material.** To view supplementary material for this article, 114 please visit https://doi.org/10.1017/ice.2020.1309

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References

- Schmidt SB, Gruter L, Boltzmann M., Rollnik JD Prevalence of serum IgG 122 antibodies against SARS-CoV-2 among clinic staff. *PLoS One* 2020;15(6). 123 doi: 10.1371/journal.pone.0235417. 124
- Korth J, Wilde B, Dolff S, *et al.* SARS-CoV-2-specific antibody detection in 125 healthcare workers in Germany with direct contact to COVID-19 patients. 126 *J Clin Virol* 2020;128:104437. 127
- Bampoe S, Lucas DN, Neall G, *et al.* A cross-sectional study of immune seroconversion to SARS-CoV-2 in frontline maternity health professionals. 129 *Anaesthesia* 2020. doi: 10.1111/anae.15229. 130
- 4. Garcia-Basteiro AL, Moncunill G, Tortajada M, *et al.* Seroprevalence of antilodies against SARS-CoV-2 among healthcare workers in a large Spanish reference hospital. *Nat Comm* 2020;11. doi: 10.1038/s41467-020-17318-x.
 133
- Patel MM, Thornsburg NJ, Stubblefield WB, et al. Change in antibodies to 134 SARS-CoV-2 over 60 days among healthcare personnel in Nashville, 135 Tennessee. JAMA Network 2020. doi: 10.1001/jama.2020.18796.
 136
- Blairon L, Mokrane S, Wilmet A, Desilly G, *et al.* Large-scale, molecular, and 137 serological SARS-CoV-2 screening of healthcare workers in a 4-site public 138 hospital in Belgium after COVID-19 outbreak. *J Infect* 2020. doi: 10.1016/ 139 j.jinf.2020.07.033.
- Martin C, Montesinos I, Dauby N, *et al.* Dynamics of SARS-CoV-2 RT-PCR 141 positivity and seroprevalence among high-risk healthcare workers and hospital staff. *J Hosp Infect* 2020. doi: 10.1016/j.jhin.2020.06.028. 143
- Iversen K, Bundgaard H, Hasselbalch RM, *et al.* Risk of COVID-19 in healthcare workers in Denmark: an observational cohort study. *Lancet* 2020. doi: 145 10.1016/S1473-3099(20)30589-2. 146
- Lackermair K, William F, Grzanna N, *et al.* Infection with SARS-CoV-2 in 147 primary care healthcare workers assessed by antibody testing. *Fam Pract* 148 2020. doi: 10.1093/fampra/cmaa078. 149