BMJ Open COVID-19 among workers of a comprehensive cancer centre between first and second epidemic waves (2020): a seroprevalence study in Catalonia, Spain

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ABSTRACT

Objectives Patients with cancer are at higher risk for severe COVID-19 infection. COVID-19 surveillance of workers in oncological centres is crucial to assess infection burden and prevent transmission. We estimate the SARS-CoV-2 seroprevalence among healthcare workers (HCWs) of a comprehensive cancer centre in Catalonia, Spain, and analyse its association with sociodemographic characteristics, exposure factors and behaviours.

Design Cross-sectional study (21 May 2020–26 June 2020).

Setting A comprehensive cancer centre (Institut Català d'Oncologia) in Catalonia, Spain.

Participants All HCWs (N=1969) were invited to complete an online self-administered epidemiological survey and provide a blood sample for SARS-CoV-2 antibodies detection.

Primary outcome measure Prevalence (%) and 95% Cls of seropositivity together with adjusted prevalence ratios (aPR) and 95% Cl were estimated.

Results A total of 1266 HCWs filled the survey (participation rate: 64.0%) and 1238 underwent serological testing (97.8%). The median age was 43.7 years (p25–p75: 34.8–51.0 years), 76.0% were female, 52.0% were nursing or medical staff and 79.0% worked on-site during the pandemic period. SARS-CoV-2 seroprevalence was 8.9% (95% CI 7.44% to 10.63%), with no differences by age and sex. No significant differences in terms of seroprevalence were observed between onsite workers and teleworkers. Seropositivity was associated with living with a person with COVID-19 (aPR 3.86, 95% CI 2.49 to 5.98). Among on-site workers, seropositive participants were twofold more likely to be nursing or medical staff.

Strengths and limitations of this study

- ➤ Seroepidemiological study with a large sample size settled in a comprehensive cancer centre.
- Questionnaire completeness was very high, with no variables presenting more than 5% of missing values
- Recall bias is possible as the data for the correlates of SARS-CoV-2 infection rely on a self-administered questionnaire.
- ► The accomplishment of preventive measures might be overestimated: response and perception biases must be considered, as well as complacency bias.
- Answers reported in the questionnaire could be influenced by the participants' knowledge regarding their COVID-19 status.

Nursing and medical staff working in a COVID-19 area showed a higher seroprevalence than other staff (aPR 2.45, 95% CI 1.08 to 5.52).

Conclusions At the end of the first wave of the pandemic in Spain, SARS-CoV-2 seroprevalence among Institut Català d'Oncologia HCW was lower than the reported in other Spanish hospitals. The main risk factors were sharing household with infected people and contact with COVID-19 patients and colleagues. Strengthening preventive measures and health education among HCW is fundamental.

INTRODUCTION

Front-line healthcare workers (HCWs) dealing with COVID-19 have higher exposure



to SARS-CoV-2 than the general population, ¹ and they can contribute to the spread of COVID-19 as per their exposure to vulnerable patients. Since the beginning of the pandemic, several studies have been published on SARS-CoV-2 infections prevalence in HCW, although with diverse results. A meta-analysis of 49 studies, including 127 480 HCWs, showed that the overall seroprevalence of SARS-CoV-2 antibodies in the European region was 8.5%. ² HCW in Spain have been highly affected: a total amount of 154 636 cases among HCWs were already officially notified by 2 December 2021 at the onset of the sixth pandemic wave. ³⁴

Patients with cancer are vulnerable, presenting a high risk for COVID-19 infection and more severe outcomes due to their immunosuppression status.⁵ The pandemic has presented unprecedented professional and personal challenges for the oncology community.⁶ Data are lacking on the seroprevalence of SARS-CoV-2 among HCW in oncological centres, and small sample sizes limit the few published studies. This study aims to estimate the seroprevalence of SARS-CoV-2 and associated sociodemographic and behavioural risk factors among workers of the Catalan Institute of Oncology (ICO), a Comprehensive Cancer Centre comprised of four hospitals in Catalonia (Spain), covering around 40% of the adult population in Catalonia.⁷

PARTICIPANTS AND METHODS Study design and setting

A cross-sectional study including blood sample collection and a self-administered questionnaire was conducted between 21 May 2020 and 26 June 2020 in the four ICO centres (L'Hospitalet de Llobregat, Badalona, Tarragona/Terres de l'Ebre and Girona).

The study population were HCW delivering care and services to patients (directly or indirectly) and support staff, including those who do not deliver care but work in other tasks within the hospital. A total of 1969 employees of ICO were invited to participate in the study through an email that allowed access to the study information. The inclusion criteria were: (1) to be an active worker during the epidemic period, (1 February 2020-26 June 2020) and (2) to be aged ≥18 years. The participants filled in an online epidemiological questionnaire and were scheduled for serology testing by the Occupational Health Department. A total of 1266 HCW filled in the online epidemiological questionnaire (participation rate: 64.3%) and 1238 of them (97.8%) underwent a serology test. Three participants with inconclusive serological results were excluded. The final analysis included 1235 participants (figure 1).

Epidemiological guestionnaire and study variables

An epidemiological questionnaire was programmed online to collect information regarding sociodemographic characteristics, working information, compliance of personal protective equipment (PPE) measures

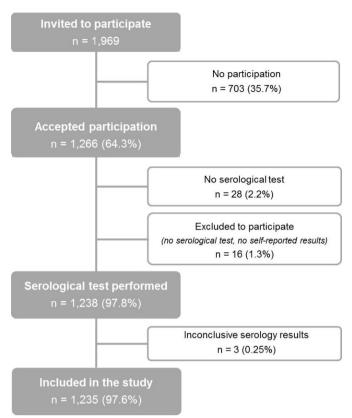


Figure 1 Participants' flow chart in the seroprevalence survey, Catalan Institute of Oncology. 21 May 2020–26 June 2020; Spain.

at work, at home and history of previous COVID-19 infection (online supplemental material). The question-naire was developed based on previous epidemiological studies conducted within the ICO centres, and a modified version was used in another seroprevalence study performed among university personnel of the University of Barcelona.⁸

Sociodemographic characteristics included information on age and sex, ICO centre of recruitment, presence of comorbidities, smoking history, pregnancy and cohabitants.

Work-related conditions included the professional category, teleworking status, type of shift, working on a COVID-19 area, contact with COVID-19 cases, contact with biological samples and reporting to be exposed to COVID-19.

Concerning PPE measures at work, participants were asked about feeling protected with PPE and compliance with PPE measures. Regarding the application of preventive measures outside the working setting, participants were asked if they got a shower after leaving the workplace or when arriving home, if they changed clothes after work or on home arrival, as well as about hand washing and use of face mask when shopping. Information about COVID-19 cases and protective measures were also collected among those participants reporting cohabitants. Participants were also asked about the type of transport used to go to work.



Participants were asked about a previous diagnosis of COVID-19 performed by rRT-PCR or serology test and date of diagnosis, as well as reporting COVID-19 compatible symptoms, and the type of symptoms.

SARS-CoV-2 laboratory testing

Serum samples from participants at L'Hospitalet, Girona and Tarragona/Terres de l'Ebre were studied at the Microbiology Department of Hospital de Bellvitge and samples from HCWs at ICO Badalona were analysed at the MetroNord Regional Clinical Laboratory, using the same procedures and techniques in both laboratories. Detection of SARS-CoV-2 antibodies was carried out using the quantitative SARS-CoV-2 S1/S2 IgG LIAISON test (DiaSorin, Vercelli, Italy) on the LIAISON XL platform, following the manufacturer's instructions. This test discriminates among negative (<12AU/mL; with 3.8 as IgG detection limit), equivocal (12.0-15.0AU/mL) and positive (>15.0AU/mL) subjects. In those cases in which (1) IgG anti S1/S2 quantification was higher than the limit of detection (ie, >3.8AU/mL) but did not reach the limit of discrimination (ie,<15AU/mL) and/or (2) when the HCW answered the questionnaire saying that he or she had been diagnosed of COVID-19 but IgG anti S1/S2 where lower than 15 AU/mL, an additional serological study was performed using a different antigen (N) as a target. In this case, a SARS-CoV-2 IgG test (Abbott Diagnostics, Sligo, Ireland) was run on an Architect i2000 platform. This test discriminates among negative (<1.4Index (S/C)) and positive (≥ 1.4 Index (S/C)) subjects.

Case definition

A seropositive case of SARS-CoV-2 was defined as seropositivity to IgG independently of previous self-reported results.

Patient and public involvement

No patient was involved in the study.

Statistical analysis

Crude global and by subgroups SARS-CoV-2 seroprevalence and 95% CIs were calculated. Differences in the distribution of study variables between seropositive and seronegative participants were assessed using χ^2 test for categorical variables, and parametric or non-parametric tests were performed for normal and non-normal continuous variables, respectively. Prevalence ratios (PR) and 95% CIs were estimated using Poisson regression models with robust variance. Adjusted PRs (aPRs) were used for statistically significant variables in the bivariate analysis and those considered relevant for the study design. Thus, adjusted models included sex, ICO centre of recruitment, age, type of HCW, teleworking and cohabitants. Linear trends for variables with ordinal categories was based in the likelihood ratio test of the model with the ordinal variable as a continuous one. P values were based on two-sided hypothesis tests and considered significant at p<0.05. All analyses were conducted using Stata V.16.0 (StataCorp).

RESULTS

A total of 1235 HCWs with serological results (figure 1) were included in the analysis: 76.0% were female, the median age was 43.7 years (p25–p75: 34.8–51.0 years), 52.2% were nursing or medical staff and 18.6% of the participants teleworked full time during the study period (table 1). Up to 14.7% of the participants reported at least one comorbidity. Regarding smoking habits, 16.0% were current smokers and 28.2% reported to be former smokers (table 1). Seven women were pregnant, and none of them showed seropositivity.

The overall crude SARS-CoV-2 seroprevalence was 8.9% (95% CI 7.44% to 10.63%), with no statistically significant differences by neither age group nor sex, and the seroprevalence for nursing and medical staff was 11.6% (95% CI 9.37% to 14.34%). After fully adjustment, the main determinants of higher seroprevalence included working at ICO Girona compared with workers at ICO L'Hospitalet (aPR 1.52, 95% CI 0.97 to 2.38) and nursing or medical staff compared with other groups (aPR 2.04, 95% CI 1.33 to 3.14) (table 1).

Seroprevalence among on-site workers was 8.8% (95%) CI 7.15% to 10.71%) (table 2). Onsite workers were younger, assisting HCWs and reported more frequently rRT-PCR previous to serology than teleworkers, but no differences were observed in sex, self-reported comorbidities, smoking history, cohabiting with COVID-19 positive case between them and teleworkers (online supplemental material). Among this group (N=981) of professionals who never or occasionally teleworked SARS-CoV-2 seropositivity was not associated with not working in a COVID-19 area (aPR 1.29, 95% CI 0.81 to 2.06), nor being in contact with COVID-19 biological samples (aPR 1.30, 95% CI 0.77 to 2.20) nor being in contact with patients with COVID-19 (aPR 1.09, 95% CI 0.66 to 1.79) were associated with SARS-CoV-2 positivity (table 2). On-site nursing or medical staff who worked in a COVID-19 area had twofold SARS-CoV-2 seroprevalence than others who did not work in COVID-19 area (aPR 2.45, 95% CI 1.08 to 5.52). Seropositivity was higher among those whom referred being exposed by interacting with colleagues (aPR 3.26, 95% CI 1.49 to 7.15). On-site workers who self-reported symptoms of COVID-19 were almost 10-fold more likely to be seropositive than those who did not (aPR: 9.5, 95% CI 5.34 to 17.03). Most of the on-site workers were highly adherent to the recommendation of hand hygiene at work. Hand washing before eating or working, were followed by more than 97% of on-site workers, whereas around 24% of them reported not hand hygiene after working or a low frequency of handwashing during the workday. In relation to protective measures at work, 17.4% of the on-site workers did not feel protected with PPE, and 12.1% did not use PPE with confirmed or suspicious COVID-19 cases. About colleagues' behaviour, 2 m safety distance from colleagues when having lunch was reported to be unfollowed by 14.1% (table 2).

Concerning the correlates of seropositivity according to household factors for all participants (table 3),

Table 1 Sociodemographic characteristics associated with SARS-CoV-2 positive serology among study participants (N=1235)

	SARS-CoV-2 Total participants seroprevalence				
		n (%)	Prevalence (95% CI)	P value*	aPR (95% CI)‡‡
Study participants	1235	110	8.91 (7.44 to 10.63)		(55 /5 5 -/ -
Sex	1200	110	0.51 (7.44 to 10.00)		
Male	291 (23.6)	27 (24.5)	9.28 (6.44 to 13.20)		REF
Female	939 (76.0)	83 (75.5)	8.84 (7.18 to 10.83)	0.82	0.82 (0.53 to 1.28)
Age (median, (p25–p75))	43.7 (34.8–51.0)	42.8 (32.0–50.1)	0.01 (1110 to 10.00)	0.62	0.99 (0.97 to 1.01)
<35 years	313 (25.3)	33 (30.0)	10.54 (7.59 to 14.46)	0.02	REF
35–49 years	566 (45.8)	47 (42.7)	8.30 (6.29 to 10.88)		0.85 (0.55 to 1.34)
>49 years	356 (28.8)	30 (27.3)	8.43 (5.95 to 11.80)	0.5	0.88 (0.53 to 1.46)
ICO centre	,	,	,		,
ICO L'Hospitalet	885 (71.7)	73 (66.4)	8.25 (6.61 to 10.25)		REF
ICO Girona	204 (16.5)	29 (26.4)	14.22 (10.06 to 19.72)		1.52 (0.97 to 2.38)
ICO Badalona	134 (10.9)	7 (6.4)	5.22 (2.51 to 10.56)		0.54 (0.25 to 1.19)
ICO Tarragona/Terres de l'Ebre	. ,	1 (0.9)	8.33 (1.16 to 41.38)	0.02	1.07 (0.15 to 7.83)
Professional category					
Nursing staff‡	380 (30.8)	43 (39.0)	11.32 (8.50 to 14.92)		REF
Medical staff§	265 (21.5)	32 (29.1)	12.08 (8.67 to 16.58)		1.07 (0.65 to 1.76)
Middle and superior technicians	285 (23.1)	14 (12.7)	4.91 (2.93 to 8.13)		0.41 (0.22 to 0.77)
Service staff¶	114 (9.2)	2 (1.8)	7.02 (3.55 to 13.42)		0.69 (0.31 to 1.54)
Porter	21 (1.7)	8 (7.3)	9.52 (2.39 to 31.16)		0.74 (0.17 to 3.24)
Administrative	129 (10.4)	8 (7.3)	6.20 (3.13 to 11.92)		0.54 (0.25 to 1.16)
Other	20 (1.6)	1 (0.9)	5.00 (0.70 to 28.26)	0.03	0.50 (0.07 to 3.71)
Nursing or medical staff**	645 (52.2)	75 (68.2)	11.63 (9.37 to 14.34)	<0.001	2.04 (1.33 to 3.14)
Other staff††	569 (46.1)	33 (30.0)	5.80 (4.15 to 8.05)		REF
Telework					
Never/occasionally	981 (79.4)	86 (78.1)	8.77 (7.15 to 10.71)		REF
Always	230 (18.6)	23 (20.9)	10.00 (6.72 to 14.63)	0.56	1.60 (0.98 to 2.59)
Shift work					
Morning	545 (44.1)	49 (45.0)	8.99 (6.86 to 11.7)		REF
Evening	140 (11.3)	10 (9.1)	7.14 (3.88 to 12.77)		0.56 (0.34 to 0.93)
Split shift (morning- evening)	417 (33.8)	38 (34.5)	9.11 (6.7 to 12.28)		0.88 (0.57 to 1.37)
Night	88 (7.1)	10 (9.1)	11.36 (6.22 to 19.86)		0.95 (0.46 to 1.96)
Other	25 (2)	3 (2.7)	12 (3.92 to 31.32)	0.83	1.15 (0.35 to 3.75)
Comorbidities**					
None	1054 (85.3)	99 (90.0)	9.39 (7.77 to 11.31)		REF
Yes	181 (14.7)	11 (10.0)	6.08 (3.4 to 10.64)	0.15	0.67 (0.36 to 1.25
Smoking history					
Never	650 (52.6)	80 (72.7)	12.31 (9.99 to 15.07)		REF
Past	348 (28.2)	22 (20.0)	6.32 (4.20 to 9.42)		0.57 (0.35 to 0.93)
Current	198 (16.0)	8 (7.3)	4.04 (2.03 to 7.87)	0.0002	0.38 (0.18 to 0.79)

Continued



Table 1 Continued

	Total participant	SARS-CoV-2 s seroprevalence			
	n (%)	n (%)	Prevalence (95% CI)	P value*	aPR (95% CI)‡‡
Cohabitants					
Yes	1119 (90.6)	95 (86.0)	8.49 (6.99 to 10.27)		REF
No	104 (8.4)	15 (13.6)	14.42 (8.88 to 22.57)	0.04	1.48 (0.83 to 2.66)

Numbers do not always sum up the total due to some missing values (none of the categories present more than 5% of missing values).

seropositivity was associated with living with a COVID-19 positive person (aPR 3.86, 95% CI 2.49 to 5.98). Up to 17.3% of the participants did not take a shower nor change clothes on home arrival, but the majority (99.0%) did hand hygiene. The least followed hand hygiene home practices were after money, phone and other personal tools manipulation and after nose blowing, coughing or sneezing (23.5% and 22.7%). However, not following protection measures or hand hygiene at home were associated with a higher SARS-CoV-2 seroprevalence.

Clinical characteristics were collected for those participants (N=469) who reported a rRT-PCR performed previous to serology (online supplemental material). The majority of the patients with a positive serology and reporting a positive rRT-PCR presented compatible COVID-19 symptoms (74.4%). Among seropositive patients, the most common symptoms were arthromyalgia, cough, headache, asthenia and anosmia. Reporting a positive rRT-PCR when presenting compatible symptoms was associated with a threefold higher prevalence of seropositivity (aPR 3.10, 95% CI 1.78 to 5.31). An increased number of compatible symptoms was also associated with a higher seroprevalence (aPR 7.4, 95% CI 1.78 to 5.31, for presenting four or more symptoms compared with no symptoms).

DISCUSSION

Despite the impact of COVID-19 in oncological patients, ¹⁰ there are scarce SARS-CoV-2 seroprevalence studies in comprehensive cancer centres with large sample sizes. The global SARS-CoV-2 seroprevalence was 8.9% during the first wave of the COVID-19 pandemic, lower than expected, owing to the presumed higher risk among HCW. Also, it was lower than the reported estimates in two studies performed among HCW in Catalonia between March-April and May 2020, showing a seroprevalence of 11.2% ¹¹ and 10.3%, ¹² respectively. In all cases, the

seroprevalence was higher than in the general population, estimated to be of a maximum of 7.4% in the Barcelona metropolitan area when the study was conducted.¹³ Seroprevalence studies interpretation must be related to the average COVID-19 prevalence at the time of blood collection. Both of the mentioned studies were carried out earlier than ours, which was performed approximately 1 month later (21 June 2020 May-26 June 2020), and 2 months after the first-wave peak in Catalonia (23 March). ¹⁴ Another explanation for this lower seroprevalence in our Centre concerns the participation: all active HCW, regardless of their teleworking status during the previous months or work absenteeism, were invited to participate and most did (64%). In contrast, Garcia-Basteiro et al's¹¹ and Barallat et al's¹² studies comprised general hospitals¹⁰ 11 and primary healthcare centres¹² in which the incidence could be higher than in a monographic cancer centre.

Several studies regarding COVID-19 infections in HCW in Spain have been published, although showing diverse results. In a tertiary-care hospital in Mallorca, with low regional seroprevalence in the general population (<2%), the prevalence of infected HCW (n=2210) was 2.8%. 15 Varona et al performed a cross-sectional study evaluating 6038 employees from the healthcare system of 17 hospitals across four regions in Spain (Madrid, Catalonia, Galicia and Castilla-Leon), showing an 11% seropositivity for SARS-CoV-2 IgG. 16 Finally, other studies in Madrid reported a seroprevalence between 16.6% and 36.5% among HCW in areas with high COVID-19 prevalence. 17-19 These studies revealed seroprevalence of SARS-CoV-2 IgG antibodies in HCW tend to be higher than in the general population, at variance according to regional COVID-19 incidence.

The prevalence of SARS-CoV-2 antibodies among HCW has been increasingly investigated in many other countries showing a broad range of outcomes. So far, two systematic

^{*}Comorbidities: hypertension, obesity (BMI ≥30), heart disease, liver disease, diabetes, chronic respiratory disease, renal disease, cancer, autoimmune disorders and other immunological disorders.

[†] χ2 test for categorical variables (Fisher's exact test corrected for continuity) and median test for continuous variables.

[‡]Nursing staff: nurses and nursing assistants.

[§]Medical staff: resident physicians and specialists.

[¶]Service staff: security, maintenance, cleaning and kitchen.

^{**}Nurses, nursing assistants, resident physicians and specialists.

^{††}Middle and superior technicians, security, maintenance, cleaning, kitchen, porter, administrative and other.

^{‡‡} Adjusted for sex, age (continous), ICO centre, telework and cohabitants.

aPR, adjusted prevalence ratio; BMI, body mass index; ICO, Institute of Oncology; p25, 25% percentile; p75, 75% percentile.

Table 2 Occupational factors associated with SARS-CoV-2 positive serology among on-site workers (N=981)						
	Total participants	SARS-CoV-2 seroprevalence				
	n (%)	n (%)	Prevalence (95% CI)	P value*	Adjusted PR (95% CI)	
On-site workers Type of transport to work	981 (79.4)	86 (78.1)	8.77 (7.15 to 10.71)	0.56		
Private	751 (76.6)	66 (76.7)	8.79 (6.96 to 11.04)		REF	
Public	154 (15.7)	15 (17.4)	9.74 (5.95 to 15.54)		1.32 (0.74 to 2.36)	
Private and public	35 (3.6)	2 (2.3)	5.71 (1.43 to 20.19)		0.63 (0.15 to 2.58)	
·			,	0.00	,	
Walking Working in a COVID-19 area	37 (3.8)	3 (3.5)	8.11 (2.63 to 22.34)	0.89	0.57 (0.14 to 2.35)	
	209 (40.6)	20 (22 7)	7.20 (5.11 to 10.20)		REF	
No Yes	398 (40.6)	29 (33.7)	7.29 (5.11 to 10.29)	0.14		
	545 (55.6)	55 (63.9)	10.09 (7.83 to 12.92)	0.14	1.29 (0.81 to 2.06)	
Type of and COVID-19 area‡	440 (45.4)	7 (0.0)	4.70 (0.07 0.0)		DEE	
Non-assisting HCW and never worked in a COVID-19 area	148 (15.1)	7 (8.0)	4.73 (2.27 to 9.6)		REF	
Non-assisting HCW and ever worked in a COVID-19 area	230 (23.4)	13 (15.1)	5.65 (3.31 to 9.5)		1.12 (0.44 to 2.82)	
Assisting HCW and never worked in a COVID-19 area	244 (24.9)	22 (25.6)	9.02 (6.01 to 13.32)		1.81 (0.77 to 4.26)	
Assisting HCW and ever worked in a COVID-19 area	311 (31.7)	40 (46.5)	12.86 (9.57 to 17.07)	0.006	2.45 (1.08 to 5.52)	
p-trend					0.26	
Contact with COVID-19 cases						
No	333 (33.9)	23 (26.7)	6.91 (4.63 to 10.18)		REF	
Yes	536 (54.6)	57 (66.3)	10.63 (8.29 to 13.54)	0.07	1.30 (0.77 to 2.20)	
Contact with COVID-19 biological sa	amples				, ,	
No	646 (65.9)	51 (59.3)	7.89 (6.05 to 10.24)		REF	
Yes	282 (28.7)	30 (34.9)	10.64 (7.54 to 14.81)	0.17	1.09 (0.66 to 1.79)	
Reporting to be exposed to COVID-	19 by interacting	with colleagues at v			,	
No	242 (24.7)	66 (76.7)	2.89 (1.38 to 5.95)		REF	
Yes	608 (62.0)	7 (8.1)	10.86 (8.62 to 13.59)	<0.0001	3.26 (1.49 to 7.15)	
Reporting COVID-19 compatible syr	nptoms	· ,	· · ·		, ,	
No	623 (63.5)	15 (17.4)	2.41 (1.46 to 3.96)		REF	
Yes	306 (31.2)	68 (79.1)	22.22 (17.91 to 27.23)	<0.0001	9.53 (5.34 to 17.03)	
Not following protection measures a		,	,		,	
Felt protected with PPE	132 (17.4)	12 (16.9)	9.09 (5.23 to 15.34)	0.83	0.98 (0.51 to 1.88)	
Colleagues cover themselves with their elbows when sneezing/ coughing	155 (15.8)	21 (24.4)	13.55 (9.00 to 19.90)	0.01	1.70 (1.01 to 2.87)	
2 m safety distance from colleagues during lunch	127 (14.1)	12 (15.6)	9.45 (5.44 to 15.91)	0.71	1.06 (0.56 to 1.99)	
Use of PPE with confirmed or suspicious COVID-19 patients	79 (12.1)	7 (10.45)	8.86 (4.28 to 17.46)	0.63	1.01 (0.45 to 2.26)	
PPE removal safety	48 (7.3)	3 (4.6)	6.25 (2.03 to 17.68)	0.33	0.54 (0.17 to 1.74)	
Personal use of mask	34 (3.5)	1 (1.2)	2.94 (0.41 to 18.17)	0.21	0.41 (0.06 to 2.99)	
Colleagues use of surgical mask	7 (0.7)	1 (1.2)	14.29 (1.96 to 58.12)	0.62	1.68 (0.23 to 12.29)	
Not following hand hygiene at work			,		,	
≤7 times during workday	233 (23.8)	15 (17.4)	6.44 (3.92 to 10.41)	0.13	0.71 (0.39 to 1.28)	
After money, phone and other personal tools manipulation	175 (17.8)	16 (18.6)	9.14 (5.67 to 14.41)	0.89	1.00 (0.58 to 1.74)	



Table 2 Continued

	Total participants	SARS-CoV-2 seroprevalence			
	n (%)	n (%)	Prevalence (95% CI)	P value*	Adjusted PR (95% CI)†
Every time entering in a new workspace	102 (10.4)	5 (5.8)	4.90 (2.05 to 11.25)	0.14	0.55 (0.22 to 1.37)
Before working	21 (2.1)	3 (3.5)	14.29 (4.67 to 36.17)	0.37	1.72 (0.54 to 5.47)
After finishing the workday	17 (1.7)	1 (1.2)	5.88 (0.82 to 32.09)	0.67	0.65 (0.09 to 4.72)
Before eating	9 (0.9)	2 (2.3)	22.22 (5.59 to 57.95)	0.16	2.67 (0.65 to 10.94)

Numbers do not always sum up the total due to some missing value (none of the categories present more than 5% of missing values). $^*\gamma 2$ test.

reviews estimated an overall seroprevalence of SARS-CoV-2 antibodies of 8.7% and 8.0% among 127 480 HCW and 168 200 HCW, respectively, before vaccination started. Seroprevalence was higher in studies conducted in North America (12.7%) compared with those conducted in Europe (8.5%), Africa (8.2) and Asia (4%).

In Europe, seroprevalence rates among HCW in Germany, Denmark and Belgium were low (1.6%, 4.0% and 6.4%,

respectively). ^{21–23} These studies were conducted during early stages of the epidemic, and therefore, they derived that infection was community acquired. Also, the Belgian study, with a sample size of almost 30 000 HCW, notes that the high availability of PPE, high standards of infection prevention and PCR screening in symptomatic staff, coupled with contact tracing and quarantine, might explain the relatively low seroprevalence. ²³ An study performed in Lombardy, Italy, ²⁴

Table 3 Household factors associated with SARS-CoV-2 positive serology among study participants (n=1235)							
	Total participants	SARS-CoV-2 seroprevalence			Adjusted PR (95%		
	n (%)	n (%)	Prevalence (95% CI)	P value*	CI)†		
Study participants	1235	110	8.91 (7.44 to 10.63)				
Cohabitants with COVID-19‡							
No	894 (79.9)	52 (54.7)	5.82 (4.46 to 7.56)		REF		
Yes	141 (12.60)	34 (35.8)	24.11 (17.76 to 31.86)	<0.0001	3.86 (2.49 to 5.97)		
Cohabitants cover themselves with the	Cohabitants cover themselves with their elbow when sneezing						
No	158 (14.1)	18 (18.9)	11.39 (7.29 to 17.37)		REF		
Yes	919 (82.1)	73 (76.8)	7.94 (6.36 to 9.88)	0.15	0.73 (0.43 to 1.22)		
Not following protection measures at	home§						
Use of face mask when shopping	17 (1.4)	2 (1.8)	11.76 (2.95 to 36.86)	0.67	0.98 (0.24 to 4.05)		
Shower and clothes changing afterwork or on home arrival	214 (17.3)	20 (18.2)	9.35 (6.11 to 14.05)	0.82	1.02 (0.62 to 1.69)		
Not following hand hygiene at home§							
On arrival	12 (1)	2 (1.8)	16.67 (4.19 to 47.76)	0.35	1.59 (0.39 to 6.60)		
Before eating	60 (4.9)	9 (8.2)	15.00 (7.99 to 26.4)	0.09	1.55 (0.77 to 3.12)		
After money, phone and other personal tools manipulation	290 (23.5)	27 (24.6)	9.31 (6.46 to 13.24)	0.71	1.01 (0.65 to 1.58)		
After cleaning	110 (8.9)	8 (7.3)	7.27 (3.68 to 13.88)	0.53	0.78 (0.38 to 1.61)		
After nose blowing	280 (22.7)	25 (22.7)	8.93 (6.1 to 12.88)	0.99	0.93 (0.58 to 1.48)		

Numbers do not always sum up the total due to some missing values (none of the categories present more than 5% of missing values). *72 test.

[†]Adjusted for sex, age (continuous), ICO centre, care staff, telework and cohabitants.

[‡]Assisting HCW: nurses, nursing assistants, resident physicians and specialists; otherwise, classified and non-assisting HCW.

HCW, healthcare worker; ICO, Institute of Oncology; PPE, personal protective equipment; PR, Prevalence Ratio.

[†]Adjusted for sex, age (continuous), ICO centre, care staff, telework and cohabitants.

[‡]Analyses performed among those participants who reported having cohabitants (n=1119).

[§]Unfollowing the measures of protection and hand hygiene recommendations.

ICO, Institute of Oncology; PR, Prevalence Ratio.

one of the Italian regions most hit by the first epidemic wave, showed a seroprevalence of 7.4% (3.8%–11.0%), similar to the observed in the Catalan studies. ^{11 12} Sweden and the UK were the two European countries reporting the highest seropositivity rates among HCW: 19.1% and between 18.0% and 45.3%, respectively. ^{25–27} In the UK, this high seroprevalence was settled in London during the week with the highest number of new cases in the city in the first wave, with around 15% seropositivity among the general population. In the USA, the prevalence of infection among HCW was 10.7%, despite high variation, as low as 1.1% in California ²⁸ to 13.7% in New York State. ²⁹

Despite SARS-CoV-2 seropositivity rate in oncological HCW has significant implications for oncological patients, scant research has been done. The only study published with a large sample size was in Tokyo, Japan, and it showed a very low seroprevalence of 0.67% among 1,190 HCW. It was performed at the end of the first wave in Japan, between the 3 August 2020 and the 30 October 2020, so this may explain the lower seroprevalence compared with our estimation. A French study performed among 663 HCW and 1011 patients with cancer, after the end of the first wave, showed also low seroprevalence both for HCW and patients (1.8% and 1.7%, respectively). Other studies that have been published were based on small sample sizes and showed very variable seroprevalence rates. 22 31-35

In our study, we found no differences in HCW seroprevalence according to sex, age and presence of comorbidities. Current or past smoking was however inversely associated to SARS-CoV-2 seroprevalence. Early studies in selected cohorts of COVID-19 patients showed a paradoxical higher risk of SARS-CoV-2 infection among non-smokers³⁶ while ever smokers showed higher risk of COVID-19 progression, including severity of the disease, intensive care unit admission and death.³⁷

It is worth mentioning that, unlike most of the other published seroepidemiological studies among HCW, this study was performed among all the HCWs of the institution, regardless they did full-time telework during the study period (21.6%). No differences by telework were found, and among all study participants the main factor associated with SARS-CoV-2 seropositivity was living with a COVID-19 case, with a 1.5 times higher probability, similarly to what has been described in other studies. This finding supports the importance of community dissemination of the infection also for HCWs.

Our study shows that among on-site HCW in an oncological centre, working as medical care staff (nursing, nursing assistant, resident physicians and specialists) in COVID-19 areas stood out as one of the main factors associated with developing SARS-CoV-2 antibodies. Published results regarding the possibility of in-hospital infection among HCW and transmission at work are controversial. Some studies did not find any relation between working in COVID-19 unit or professional category with seropositivity whereas other studies reported that seroprevalence was strongly associated with patient related work. ¹⁶ ²¹ ²² ²⁵

Contact with colleagues at work is potentially a risky situation for transmission among HCW as well as the relaxation of protective measures at the end of the working day. In our study, the on-site HCW who reported being exposed to COVID-19 by other colleagues presented an almost fourfold probability of being seropositive. Most of the HCWs declared to follow the protective measures at the workplace, and no differences in seroprevalence were found according to protective measures and hand hygiene.

Protecting HWC health is of paramount importance for reducing morbidity and mortality, reducing transmission and maintaining the health system capacity. Thus, international health authorities recommend screening strategies for SARS-CoV-2 infection in exposed or high-risk HCW as well as massive COVID-19 vaccination.

Significant differences exist in SARS-CoV-2 testing between countries, and existing programmes focus on screening symptomatic rather than asymptomatic staff. Published studies point out the fact that screening should be performed regardless of the absence of typical symptoms for COVID-19 disease. It has been demonstrated that seroconversion can occur in HCW who have suffered no previous symptoms of SARS-CoV-2 infection 41 42 as asymptomatic transmission is very relevant in SARS-CoV-2 spread. 42 43 Thus, the approach for mass testing of both symptomatic and asymptomatic HCW could mitigate workforce depletion by unnecessary quarantine, reduce spread in atypical, mild or asymptomatic cases; and protect patients and healthcare workforce.

Among the potential limitations of the study, some recall bias is possible as the data for the correlates of SARS-CoV-2 infection rely on a self-administered questionnaire. Also, response and perception biases must be considered, as well as complacency bias. Results, especially those regarding the accomplishment of preventive measures, might be overestimated. Answers reported in the questionnaire could be influenced by the participants' knowledge regarding their COVID-19 status. However, this study is the first seroepidemiological study with such a large sample size settled in an oncological health centre. The sufficient sample size and high response rate (64.3%) are strengths of the study, although information regarding non-participants was not collected, and we cannot disregard a potential participation bias. However, the distribution by age and sex was similar between participants and non-participants and a possible reason for no participation is that professionals from ICO-Badalona had previously participated in an HCW county seroprevalence survey. 12 Also, the fact that the information of the study and the questionnaire was published online and sent by email, as well as the short period of time stablished to respond to it, could have limited the participation. Questionnaire completeness was very high, with no variables presenting more than 5% of missing values.

In conclusion, SARS-CoV-2 seroprevalence among ICO HCW at the end of the first wave of the pandemic was lower than the reported in other Catalan hospitals, but higher than among the general population living in the area. Whereas the main risk factor was living with infected people, among on-site workers, contact with colleagues was associated with SARS-CoV-2 infection. Knowing the seroprevalence rate and follow-up evaluation of persistence may help hospitals to characterise the staff at risk, rationalise their placement,



prioritise the use of PPE, thereby potentially reducing the risk of infection. Follow-up studies to evaluate long-term durability of antibodies among HCW will be of interest, after the introduction of COVID-19 vaccination among HCW, to better promote infection control in this group. Strengthening preventive measures and health education among HCW is fundamental, especially in oncological departments and centres.

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Contributors EF, DC-P, AP, CC, AC and AS-L contributed to study design. SC, AD, LG, IB, JT, MG, FS, JJ, DC, BC-C, DR-T, CG, AA-P and AP accrued participants and care for blood collection at ICO centres. Laboratory analyses were coordinated by MADL. The questionnaire was designed by DC-P and EF, and revised by PP-T, AS-L, YBM, DC, AP and LA. Questionnaire's implementation was done by EL, JM, JPR and CM-M. Data were analysed by YB and DC. PP-T, AS-L, YB, DC, LA, and EF interpreted the initial results and designed the tables. All authors contributed to interpretation of results. The first draft of the manuscript was prepared by PP-T and AS-L and EF. PP-T, AS-L, YB, DC, LA, DC-P and EF were the main contributors to the writing of the manuscript. All authors assisted in manuscript review. The co-senior authors had full access to all the data in the study for interpretation and had final responsibility for manuscript generation and review, and the decision to submit for publication. EF is the guarantor.

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