

Original Article

Seroprevalence of IgG against SARS-CoV-2 and its determinants among healthcare workers of a COVID-19 dedicated hospital of India

Mala Mahto¹, Ayan Banerjee¹, Bijit Biswas², Sushil Kumar¹, Neeraj Agarwal², Prabhat Kumar Singh³

¹Department of Biochemistry, All India Institute of Medical Sciences, Patna, Bihar, India; ²Department of Community and Family Medicine, All India Institute of Medical Sciences, Patna, Bihar, India; ³Department of Anaesthesiology and Critical Care, Director, All India Institute of Medical Sciences, Patna, Bihar, India

Received December 24, 2020; Accepted January 27, 2021; Epub February 15, 2021; Published February 28, 2021

Abstract: Healthcare workers (HCWs) due to their job profile are at utmost risk of contracting severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) infection. Serological survey is an useful tool for vulnerability mapping in an infectious disease pandemic. The aim of the current study was to assess seroprevalence of IgG against SARS-CoV-2 and its determinants among HCWs of a tertiary healthcare facility of India. It was an observational study, cross-sectional in design conducted among 919 HCWs of All India Institute of Medical Sciences, Patna, Bihar, India during September, 2020. In results, IgG seroprevalence for SARS-CoV-2 among the study subjects was 13.3% [95% confidence interval (CI): 11.2-15.6%]. In univariate logistic regression analysis; gender, occupation, place of posting, use of full personal protective equipment (PPE), prior corona virus disease (COVID)-19 infection, influenza like illness (ILI), use of steam inhalation, consumption of azithromycin, zinc and vitamin C were the significant attributes which affected the IgG seropositivity for SARS-CoV-2. In the multivariable logistic regression model; occupation, place of posting, prior COVID-19 infection and ILI were significant determinants of IgG seropositivity for SARS-CoV-2. To conclude, majority of the HCWs were found to be IgG seronegative for SARS-CoV-2. Till availability of effective vaccine all of the HCWs should abide by infection prevention and control (IPC) measures to keep themselves and their contacts protected from SARS-CoV-2.

Keywords: COVID-19, immunoglobulin G, seroepidemiologic studies, health personnel, epidemiologic factors

Introduction

Since its emergence in late 2019 corona virus disease-19 (COVID-19) emerged as global public health emergency of the century affected about ninety million and claimed about one and two million lives till date [1, 2]. India reported its first case of COVID-19 in January, 2020 and currently it is second in terms of total number of reported cases following USA with over ten million cases and over hundred and fifty thousand reported deaths [1, 3].

Serum antibody response (IgM and IgG) to SARS-CoV-2 is detectable in between 10-21 days of infection in most of the cases with median seroconversion time of 11 and 14 days after symptom onset for IgM and IgG respectively [4]. The level of antibody production is

proportional to severity of symptoms and may reduce or even disappear after three months after the disease onset [4, 5]. The aim of a serological survey is to measure proportion of people in a community or group have detectable and moreover protective level of antibodies (especially IgG) against a particular disease of interest. It not only helps to track progress of an infectious disease pandemic like COVID-19 in a certain community or group moreover it also helps to quantify risk of the members of that particular community or group to subsequent infection [6-8].

Healthcare workers (HCWs) are backbone of any health care system more so during times of a global pandemic like COVID-19. During the ongoing COVID-19 pandemic, health-care workers are at a substantially increased risk of con-

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

tracting SARS-CoV-2. This is because of their more frequent contact with a confirmed or suspect COVID-19 case or their body fluids during discharging their duties [9, 10]. Some of the COVID-19 cases among HCWs may have gone undetected due to absence of symptoms or decision of not undergoing antigen test for the disease [11]. As per existing literature seroprevalence of IgG for SARS-CoV-2 among the HCWs varies between 0.8-13.6%. Previous studies have also reported that age, gender, place of posting, comorbidity status, personal protective equipment (PPE) use, level of exposure, prior SARS-CoV-2 infection and influenza like illness (ILI) significantly determines IgG seropositivity for SARS-CoV-2 among HCWs [5, 12-19]. Although there may be several other factors which might influence immunogenesis against SARS-CoV-2 like diet preference, steam inhalation, consumption of zinc, azithromycin and multivitamins [20-23]. Most of the prior studies in this regard were conducted in western countries like Belgium, Germany, Spain, Italy and USA [5, 13, 14, 16, 18, 19]. Among the Indian studies most were reported from western part of the country [12, 15, 17]. With this background and to bring about better understanding on the issue the current research was envisioned to assess IgG seropositivity for SARS-CoV-2 and its determinants among HCWs of a COVID-19 dedicated tertiary care health facility of India. The study has taken into account all the prior reported and postulated factors which could influence SARS-CoV-2 IgG seropositivity among the HCWs.

Methods

Study type and design

It was a monocentric, observational study, cross-sectional in design conducted among HCWs of All India Institute of Medical Sciences (AIIMS), Patna, Bihar, India during the month of September 2020.

Study setting

AIIMS-Patna is one of the centres of excellence in terms of medical education and patient care in India. Since the emergence of the pandemic in the state of Bihar the institute has provided best possible care and treatment for the attending COVID-19 patients from not only Bihar but some adjoining states too. On 10th

July 2020, the institute was designated as COVID-19 dedicated hospital by the Government of Bihar. Currently the institute has about 460 general and 60 intensive care unit (ICU) beds for the attending COVID-19 patients manned by approximately 3150 HCWs.

Sample size, sampling and enrolment

Assuming that at least 11.1% (IgG seroprevalence among HCWs reported by a prior Indian study by Kumar et al. [17]) of the subjects in the study population will be IgG seropositive for SARS-CoV-2, adjusting for finite population size (3150), 20% relative precision (~2.2% absolute precision) and 95% confidence, the final minimum sample size for the study was calculated to be 624. The sample size was calculated using 'statulator', an online sample size calculator. The study was envisioned for HCWs of AIIMS-Patna only. Thus, any HCW who was working in AIIMS-Patna during the study period were included while those who were not working in the institute and unwilling to participate were excluded. For enrolment in the study, email and short message service (SMS) invitation to all the working staff in the institute during the study period was sent which contained their scheduled date and venue of blood sample collection for antibody testing. Before 5ml blood sample collection for SARS-CoV-2 IgG testing they were self-administered a structured schedule along with a consent form to obtain their background characteristics and consent for the study respectively. The serum IgG report of the study subjects were made available in health management information system of AIIMS-Patna and a designated report dispensing counter in outpatient department (OPD) within 24 hours of blood sample collection. In total 967 study subjects participated in the study which is about 30% of our total workforce. Out of these 919 study subjects met the eligibility criteria. Data for all the variables were available in case of 689 study subjects. The details of recruitment process of the study subjects is depicted in **Figure 1**.

Study variables

The structured schedule comprised of their sociodemographic details [age in completed years, gender (male/female)], occupational characteristics [occupation (doctor/nurse/technician/account staff/attendant/sanitary staff/

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

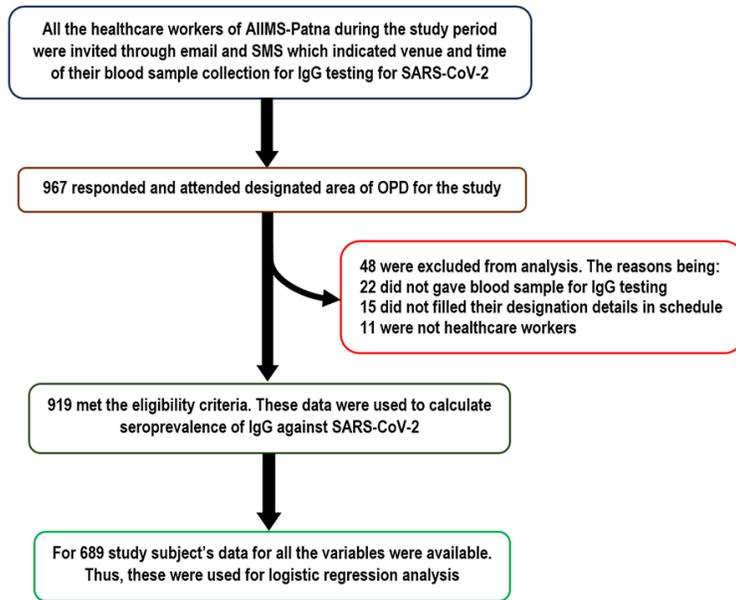


Figure 1. Flowchart showing recruitment of the study subjects.

others), place of posting (triage area/wards/ICUs/laboratories/others), whether exposed to confirmed COVID-19 case or their body fluids during duty, if yes average duration of exposure per duty shift (in hours), personal protective equipment (PPE) used during duty], personal history [currently smokes (yes/no), consumes alcohol (yes/no), known chronic co-morbidity status (yes/no), If yes name of co-morbidity suffering from, diet preference (vegetarian/non-vegetarian), history of prior COVID infection detected by reverse-transcriptase-polymerase chain reaction (RTPCR) or rapid antigen test for SARS-CoV-2 (yes/no), history of ILI in last 8 months (yes/no)] and practices related to the disease in last 8 months [used masks other than workplace (yes/no), sanitiser other than workplace (yes/no), steam inhalation (yes/no), hot beverages like hot water, tea and coffee (yes/no), consumed hydroxychloroquine (HCQ) (yes/no), azithromycin (yes/no), zinc (yes/no), multivitamin (yes/no), vitamin C (yes/no), vitamin E (yes/no)].

Some operational definitions used in the study were as following

Full PPE: Those who reported to use goggles, N-95 mask, gown covering the whole body except hand, foot and front of face, double layer

gloves and shoe cover were considered to be using full PPE.

Serum IgG level for SARS-CoV-2: It was estimated using chemiluminescent immunoassay (CLIA) named 'ADVIA Centaur COV2G' which is a qualitative and semi-quantitative assay with excellent reported sensitivity (100.0%) and specificity (99.8%) by the manufacturer [24].

IgG seropositive for SARS-CoV-2: Those with serum IgG level of 1.00 or higher was considered as IgG seropositive for SARS-CoV-2 [24].

Ethical Issues

Ethical clearance of the Institutional Ethics Committee (IEC) of AIIMS-Patna (Ref. No. -AIIMS/Pat/IEC/2020/575) was taken before conducting the research. Informed written consent of each study subject was obtained before their enrolment in the study. The data analysis and manuscript drafting were done ensuring anonymity of the study participants. The study was designed, conducted and reported abiding by declaration of Helsinki.

Statistical analysis

IBM statistical package for social sciences (SPSS) (Chicago, USA) (version 22) was used for analysis of the data. At first, descriptive analysis using number, percentage and 95% confidence interval (CI) was performed. This has shown distribution of the study subjects as per their background characteristics and IgG seropositivity for SARS-CoV-2. Then to find out univariate and multivariable determinants of IgG seropositivity among the study subjects logistic regression analysis was performed. Attributes which were found to be significant ($P < 0.05$) in univariate logistic regression were only entered in multivariable logistic regression model using forced entry method. The strength of association was reported in terms of odds ratio (OR). Insignificant Hosmer-Lemeshow test ($P \geq 0.05$) indicated multivariable logistic regres-

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

sion model fit. For all the analysis minimum acceptable confidence level was $\alpha=0.95$.

Results

Background characteristics and seroprevalence

The median age of the study subjects was 29 years with interquartile range (IQR) of 26-32 years (range: 20-56 years). There was almost equal representation of both the sexes. Majority of the study subjects (72.8%) reported direct exposure to confirmed COVID-19 cases or their body fluids during performance of their duties with median duration of exposure per duty of 7 hours with IQR (6-9 hours) (range: 1-12 hours). Considering comorbidities, 4% of the HCWs reported to have it with hypothyroidism (2.0%) being most common co-morbidity reported followed by diabetes (0.7%) and hypertension (0.5%). Out of 919 HCWs, 13.3% (11.2-15.6%) were found to be IgG seropositive for SARS-CoV-2. The seropositivity was almost double in males (17.6%) in comparison to females (8.7%). Considering occupation attendants were most likely (31.4%) to be IgG seropositive followed by account staffs (30.0%), sanitary staffs (25.8%) and technicians (24.4%). Intensive care unit (ICU) staffs were least likely (5.2%) and laboratory staffs were most likely (28.6%) to be IgG seropositive (**Table 1**).

Predictors of seroprevalence

In univariate logistic regression analysis; gender, occupation, place of posting, use of full PPE, prior COVID-19 infection, ILI, use of steam inhalation, consumption of azithromycin, zinc and vitamin C were the significant attributes affecting IgG seropositivity for SARS-CoV-2. In multivariable logistic regression analysis; occupation, place of posting, prior COVID infection and ILI were significant determinants of IgG seropositivity for SARS-CoV-2. Overall, the independent variables in the multivariable logistic regression model predicted 30.8% variability of the SARS-CoV-2 IgG seropositivity of the HCWs with high predictive accuracy rate (PAR) (88.1%) (**Table 2**).

Discussion

The study was aimed to assess IgG seropositivity for SARS-CoV-2 and its determinants among

HCWs of a COVID dedicated tertiary healthcare facility of India.

We found that 13.3% of our HCWs were seropositive for SARS-CoV-2. It was similar with the findings of two Indian studies by Prakash et al. [12] (13.6%) and Kumar et al. [17] and a study conducted in Belgium by Martin et al. [25] (11.0%). Although an prior Indian study by Baveja et al. [15] (6.9%), three European country studies (1 Germany, 2 Italy, 1 Spain) by Schmidt et al. [19] (2.9%), Amendola et al. [14] (5.1%), Sotgiu et al. [26] (7.4%), Garcia-Basteiro et al. [16] (7.6%) and an American study by Mughal et al. [18] (0.8%) reported it to be less compared to us. The variability of the finding may be attributed to many factors. Such as variation in study subject selection (Martin et al. [25] recruited only staffs working in COVID-19 units, Prakash et al. [12] and Amendola et al. [14] recruited both HCWs and non-HCWs, Sotgiu et al. [26] recruited apparently healthy HCWs and Mughal et al. [18] recruited only ICU staff which was unlike us); different techniques used for serum IgG for SARS-CoV-2 estimation (Prakash et al. [12], Kumar et al. [17], Martin et al. [25], Schmidt et al. [19], Amendola et al. [14] and Garcia-Basteiro et al. [16] used enzyme linked immunosorbent assay (ELISA) while Baveja et al. [15], Sotgiu et al. [26] and Mughal et al. [18] used rapid immunochromatography test which was unlike us), socio-cultural differences and moreover due to variation in immune responses which is likely to be influenced by genetic, ethnic and climatic factors [27, 28].

In the present study we found no association between age and SARS-CoV-2 IgG seropositivity. This was in line with the findings of Martin et al. [13] and Kumar et al. [17]. We found that males were more likely to be IgG seropositive compared to females. This was in line with the findings of Amendola et al. [14] and Kumar et al. [17] which reported similar observations. This might be because Indian men due to their various outdoor activities (i.e. shopping of household goods) and high mobility in comparison to their female counterparts are at more risk of contracting SARS-CoV-2 infection. Moreover, Indian women are twice more likely to be anaemic in comparison to their male counterparts irrespective of their socio-economic status. Anaemia is a known influencer of immune response to any infectious agent [29, 30].

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

Table 1. Distribution of the healthcare workers as per their background characteristics and seropositivity for IgGn=919

Variable	Total			IgG seropositive against SARS-CoV-2		
	N	%	95% CI	N	%	95% CI
Age in years						
<30 (median 29 years)	515	56.0	52.8-59.2	67	13.0	10.4-16.2
≥30	404	44.0	40.8-47.2	55	13.6	10.6-17.3
Gender						
Male	471	51.3	48.0-54.5	83	17.6	14.4-21.3
Female	448	48.7	45.3-52.0	39	8.7	6.4-11.7
Occupation						
Doctor	124	13.5	11.4-15.8	16	12.9	8.1-19.9
Nurse	523	56.9	53.7-60.1	31	5.9	4.2-8.3
Technician	41	4.5	3.3-6.0	10	24.4	13.8-39.3
Account staff	20	2.2	1.5-3.5	6	30.0	14.5-51.9
Attendant	118	12.8	10.8-15.2	37	31.4	23.7-40.2
Sanitary staff	31	3.4	2.4-4.8	8	25.8	13.7-43.2
Others	62	6.7	5.3-8.5	14	22.6	14.0-34.4
Place of posting: (n=839)						
Triage	58	6.9	5.4-8.8	7	12.1	6.0-22.9
Wards	397	47.3	44.0-50.7	56	14.1	11.0-17.9
ICUs	249	29.7	26.7-32.9	13	5.2	3.1-8.7
Laboratories	42	5.0	3.7-6.7	12	28.6	17.2-43.6
Others	93	11.1	9.1-13.4	23	24.7	17.1-34.4
Exposure to confirmed COVID-19 cases or their body fluids during duty: (Yes)	669	72.8	69.8-75.6	74	11.1	8.9-13.7
PPE use: (n=840)						
Full PPE	610	72.6	69.5-75.5	68	11.1	8.9-13.9
Both N-95 and surgical mask with gloves	46	5.5	4.1-7.2	14	30.4	19.1-44.8
Both N-95 and surgical mask	41	4.9	3.6-6.5	12	29.3	17.6-44.5
N-95 mask and gloves	37	4.4	3.2-6.0	5	13.5	5.9-27.9
N-95 mask only	47	5.6	4.2-7.4	5	10.6	4.6-22.6
Others	59	7.0	5.5-8.9	5	8.5	3.7-18.3
Had prior COVID-19 infection: (Yes)	79	8.6	6.9-10.6	40	50.6	39.8-61.4
Had ILI in past few months: (Yes)	69	7.5	6.0-9.4	21	30.4	20.8-42.1
Had co-morbidity: (No)	882	96.0	94.5-97.1	113	12.8	10.8-15.2
Used to smoking: (Yes)	33	3.6	2.6-5.0	1	3.0	0.5-15.3
Used to alcohol drinking: (Yes)	31	3.4	2.4-4.7	4	12.9	5.1-28.8
Used to drink hot beverages: (Yes)	808	87.9	85.7-89.9	108	13.4	11.2-15.9
Used to take steam inhalation: (Yes)	112	12.2	10.2-14.5	27	24.1	17.1-32.8
Used mask other than workplace: (Yes)	899	97.8	96.7-98.6	118	13.1	11.1-15.5
Used sanitiser other than workplace: (Yes)	903	98.3	97.2-98.9	117	13.0	10.9-15.3
Diet preference: (n=808)						
Vegetarian	244	30.2	27.1-33.5	28	11.5	8.1-16.1
Non-vegetarian	564	69.8	66.5-72.9	83	14.7	12.0-17.9
Have consumed HCQ: (Yes)	106	11.5	9.6-13.8	13	12.3	7.3-19.9
Have consumed Azithromycin: (Yes)	138	15.0	12.8-17.5	40	29.0	22.1-37.0
Have consumed Zinc: (Yes)	49	5.3	4.1-7.0	15	30.6	19.5-44.5
Have consumed Multivitamin: (Yes)	96	10.4	8.6-12.6	21	21.9	14.8-31.1
Have consumed Vitamin C: (Yes)	182	19.8	17.3-22.5	45	24.7	19.0-31.5
Have consumed Vitamin E: (Yes)	43	4.7	3.5-6.2	10	23.3	13.1-37.7

ICU: intensive care unit, PPE: personal protective equipment, ILI: influenza like illness, HCQ: hydroxychloroquine, CI: confidence interval.

In our study occupation of the study subjects emerged as a significant influencer of IgG seropositivity to SARS-CoV-2. We found that staffs

other than doctors and nurses were more likely to be IgG seropositive for the disease. Here educational level of the study subjects might

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

Table 2. Univariate and multivariable logistic regression analysis showing determinants of serum IgG status of the healthcare workers n=689*

Variable	Total		IgG seropositive against SARS-CoV-2			
	N	%	N	%	COR (95% CI)	AOR (95% CI)
Age in years						
<30 (median 29 years)	381	55.3	52	13.6	1.0 (0.7-1.6)	-
≥30	308	44.7	41	13.3	Ref.	
Gender						
Male	358	52.0	62	17.3	2.0 (1.3-3.2)	1.1 (0.6-1.9)
Female	331	48.0	31	9.4	Ref.	
Occupation						
Nurse	388	56.3	23	5.9	Ref.	Ref.
Doctor	105	15.2	11	10.5	1.8 (0.9-3.9)	1.6 (0.7-3.8)
Technician	29	4.2	9	31.0	7.1 (2.9-17.4)	3.9 (1.3-12.4)
Account staff	15	2.2	5	33.3	7.9 (2.5-25.1)	9.3 (2.1-40.8)
Attendant	91	13.2	31	34.1	8.2 (4.5-15.0)	9.6 (4.4-20.9)
Sanitary staff	14	2.0	4	28.6	6.3 (1.8-21.8)	10.8 (2.8-41.0)
Others	47	6.8	10	21.3	4.3 (1.9-9.7)	6.3 (2.0-19.5)
Place of posting						
ICUs	197	28.6	8	4.1	Ref.	Ref.
Triage	48	7.0	6	12.5	3.4 (1.1-10.2)	1.7 (0.5-5.8)
Wards	338	49.1	50	14.8	4.1 (1.9-8.8)	1.9 (0.8-4.5)
Laboratories	30	4.4	10	33.3	11.8 (4.2-33.3)	6.0 (1.8-20.5)
Others	76	11.0	19	25.0	7.9 (3.3-18.9)	2.1 (0.7-6.7)
Exposure to confirmed COVID-19 cases or their body fluids during duty: (Yes)						
Used full PPE: (Yes)	493	71.6	58	11.8	0.6 (0.4-0.9)	0.6 (0.3-1.2)
Had prior COVID-19 infection: (Yes)	60	8.7	28	46.7	7.6 (4.3-13.4)	6.9 (2.9-16.5)
Had ILI in past few months: (Yes)	45	6.5	14	31.1	3.2 (1.6-6.3)	2.6 (1.2-5.8)
Had co-morbidity: (No)	659	95.6	86	13.1	0.5 (0.2-1.2)	-
Used to smoking: (Yes)	25	3.6	0	0.0	-	-
Used to alcohol drinking: (Yes)	24	3.5	3	12.5	0.9 (0.3-3.1)	-
Used to drink hot beverages: (Yes)	612	88.8	82	13.4	0.9 (0.5-1.8)	-
Used to take steam inhalation: (Yes)	94	13.6	22	23.4	2.2 (1.3-3.9)	1.0 (0.5-2.2)
Used mask other than workplace: (Yes)	677	98.3	92	13.6	1.7 (0.2-13.5)	-
Used sanitiser other than workplace: (Yes)	680	98.7	92	13.5	1.2 (0.2-10.1)	-
Diet preference						
Vegetarian	206	29.9	23	11.2	Ref.	
Non-vegetarian	483	70.1	70	14.5	1.3 (0.8-2.2)	
Have consumed HCQ: (Yes)	89	12.9	9	10.1	0.7 (0.3-1.4)	-
Have consumed Azithromycin: (Yes)	104	15.1	30	28.8	3.3 (2.0-5.5)	1.5 (0.7-3.3)
Have consumed Zinc: (Yes)	38	5.5	10	26.3	2.4 (1.1-5.2)	0.6 (0.2-1.6)
Have consumed Multivitamin: (Yes)	73	10.6	14	19.2	1.6 (0.9-3.0)	-
Have consumed Vitamin C: (Yes)	140	20.3	34	24.3	2.7 (1.7-4.3)	1.3 (0.6-2.7)
Have consumed Vitamin E: (Yes)	28	4.1	7	25.0	2.2 (0.9-5.4)	-
Nagelkerke R ²	-	-	-	-	-	.308
Hosmer Lemeshow test p-value	-	-	-	-	-	.919
Predictive accuracy rate (PAR)	-	-	-	-	-	88.1

*data for all the variables were available for 689 study subjects thus it was used for performing logistic regression analysis; ICU: intensive care unit, PPE: personal protective equipment, ILI: influenza like illness, HCQ: hydroxychloroquine, COR: crude odds ratio, AOR: adjusted odds ratio, CI: confidence interval.

have played a role as nurses and doctors by virtue of their professional training likely to be more aware of infection prevention and control (IPC) measures to be taken for contagious disease like COVID-19. Thus, they might have

taken more precaution in comparison to the other staffs to get themselves protected from SARS-CoV-2 infection. Similarly, we found place of posting as significant attribute affecting IgG seropositivity among the study subjects as in

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

comparison to those who were posted in ICUs all the other staffs were more likely to be IgG seropositive for the disease. This may be because those who were working in ICUs might have more served patients with severe form of the disease (i.e. multi organ failure, death) unlike other staffs. Moreover, HCWs deployed in ICUs oftenly conducts high-risk procedures (i.e. intubation, cardiopulmonary resuscitation) which are known to increase risk of the disease transmission. All these might have increased their perceived risk of contracting SARS-CoV-2 infection and enforced them to practice more stringent IPC measures compared to others. This was similar with the findings of Baveja et al. [15] which reported that working in COVID area as protective for IgG seroprevalence. Although Amendola et al. [14] reported that those who were posted in paediatric intensive care and surgery were more likely to be IgG seropositive against the disease. The variation of findings could be due to overall lower IgG seroprevalence was reported by Amendola et al. [9] (5.1%) and additionally the study has included non-HCWs in addition to HCWs in it which were unlike us. In our study, those who used full PPE during their duty were 40% less likely to be IgG seropositive for SARS-CoV-2 which was in line with the findings of Baveja et al. [15]. This re-establishes the importance of use of proper personal protective measures to reduce the risk of infection.

We found that those who had prior COVID infection were 6.9 times more likely to be IgG seropositive against the disease. This was in concordance with the findings of Garcia-Basteiro et al. [16] and Kumar et al. [17]. This was an obvious finding as acquiring infection of an infectious disease agent is the only way to develop immunity against that particular disease in absence of an effective vaccine. Similarly, those who had ILI symptoms in previous 8 months had 2.6 times higher odds for IgG seropositivity. Garcia-Basteiro et al. [16] and Kumar et al. [17] reported similar observations. The persons with ILI symptoms are the major focus of COVID testing strategy of India since very early stage of the pandemic due to higher probability of these persons to be SARS-CoV-2 positive [31]. These study subjects with history of ILI symptoms and seropositive excepting those who undergone testing might have acquired mild form of SARS-CoV-2 infection and devel-

oped immunity against it. These study subjects remained undiagnosed as they did not opt for testing for COVID-19. Some other factors like use of steam inhalation, consumption of azithromycin, zinc and vitamin-C although have shown significant association with IgG seropositivity in univariate analysis got neutralised in multivariable model which signifies their limited role in immunity development against SARS-CoV-2. Thus, these should be continued to use as supportive measure. Therapeutic role of these attributes in immunity development against SARS-CoV-2 is subject to further investigation.

Limitations

Self-reporting by the HCWs were the source of most of the study data thus there may be social desirability and reporting biases which were inevitable. Secondly, as we invited all the HCWs of our institute for the study so there might be chances that those with current or prior ILI symptoms, SARS-CoV-2 infection and working in more high-risk areas (i.e. ICUs) might have participated more to know their immunity status against the disease. So, there might be response bias which limited the generalisability of the study findings to other healthcare settings.

Conclusion

Majority of the HCWs were found to be IgG seronegative for SARS-CoV-2. Occupation, place of posting, prior SARS-CoV-2 infection and ILI were found to be significant multivariable determinants of IgG seropositivity for SARS-CoV-2 in the study subjects. Till availability of effective vaccine all of the HCWs should abide by infection prevention and control (IPC) measures to keep themselves and their contacts protected from SARS-CoV-2 as most them were found to be lacking protective antibody level against the disease. Serum IgG antibody surveillance for SARS-CoV-2 may be a useful strategy to track the progress of the COVID-19 pandemic by assessment of immunity level for the disease among population at increased risk such as HCWs.

Acknowledgements

We would like to acknowledge the sincere efforts of the laboratory staffs and other staffs of All India Institute of Medical Sciences, Patna

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

involved in the study. Without their unconditional support the study would not have been possible. We received no additional fund for the study. The institutional fund as part of healthcare workers health policy was utilised for this study.

Disclosure of conflict of interest

None.

Address correspondence to: Dr. Bijit Biswas, Department of Community and Family Medicine, All India Institute of Medical Sciences, Phulwarisharif, Patna 801507, Bihar, India. Tel: +917003881125; E-mail: drbijitbiswas@gmail.com

References

- [1] Coronavirus Update (Live): 88,618,548 Cases and 1,908,982 Deaths from COVID-19 Virus Pandemic-Worldometer [Internet]. [cited 2021 Jan 8]. Available from: <https://www.worldometers.info/coronavirus/>.
- [2] Gates B. Responding to Covid-19 - a once-in-a-century pandemic? *N Engl J Med* 2020; 382: 1677-1679.
- [3] Andrews MA, Areekal B, Rajesh KR, Krishnan J, Suryakala R, Krishnan B, Muraly CP and Santhosh PV. First confirmed case of COVID-19 infection in India: a case report. *Indian J Med Res* 2020; 151: 490-492.
- [4] Immune responses and immunity to SARS-CoV-2 [Internet]. *Eur. Cent. Dis. Prev. Control*. [cited 2020 Jan 8]. Available from: <https://www.ecdc.europa.eu/en/covid-19/latest-evidence/immune-responses>.
- [5] Patel MM, Thornburg NJ, Stubblefield WB, Talbot HK, Coughlin MM, Feldstein LR and Self WH. Change in antibodies to SARS-CoV-2 over 60 days among health care personnel in Nashville, Tennessee. *JAMA* 2020; 324: 1781-1782.
- [6] Coronavirus disease (COVID-19): Serology, antibodies and immunity [Internet]. [cited 2021 Jan 8]. Available from: <https://www.who.int/news-room/q-a-detail/coronavirus-disease-covid-19-serology>.
- [7] Metcalf CJ, Farrar J, Cutts FT, Basta NE, Graham AL, Lessler J, Ferguson NM, Burke DS and Grenfell BT. Use of serological surveys to generate key insights into the changing global landscape of infectious disease. *Lancet Lond Engl* 2016; 388: 728-730.
- [8] Serology in the context of COVID-19 [Internet]. [cited 2021 Jan 9]. Available from: <https://www.who.int/emergencies/diseases/novel-coronavirus-2019/serology-in-the-context-of-covid-19>.
- [9] Nguyen LH, Drew DA, Graham MS, Joshi AD, Guo CG, Ma W, Mehta RS, Warner ET, Sikavi DR, Lo CH and Kwon S. Risk of COVID-19 among front-line health-care workers and the general community: a prospective cohort study. *Lancet Public Health* 2020; 5: e475-e483.
- [10] Ng K, Poon BH, Kiat Puar TH, Shan Quah JL, Loh WJ, Wong YJ, Tan TY and Raghuram J. COVID-19 and the risk to health care workers: a case report. *Ann Intern Med* 2020; 172: 766-767.
- [11] Zhao D, Wang M, Wang M, Zhao Y, Zheng Z, Li X, Zhang Y, Wang T, Zeng S, Hu W, Yu W and Hu K. Asymptomatic infection by SARS-CoV-2 in healthcare workers: a study in a large teaching hospital in Wuhan, China. *Int J Infect Dis* 2020; 99: 219-225.
- [12] Prakash O, Solanki B, Sheth JK, Joshi B, Kadam M, Vyas S, Shukla A, Tiwari H, Rathod S, Rajput A and Trivedi T. Assessing seropositivity for IgG antibodies against SARS-CoV-2 in Ahmedabad city of India: a cross-sectional study. *BMJ Open* 2021; 11: e044101.
- [13] Martin C, Montesinos I, Dauby N, Gilles C, Dahma H, Van Den Wijngaert S, De Wit S, Delforge M, Clumeck N and Vandenberg O. Dynamics of SARS-CoV-2 RT-PCR positivity and seroprevalence among high-risk healthcare workers and hospital staff. *J Hosp Infect* 2020; 106: 102-106.
- [14] Amendola A, Tanzi E, Folgiori L, Barcellini L, Bianchi S, Gori M, Cammi G, Albani E and Zucconi GV. Low seroprevalence of SARS-CoV-2 infection among healthcare workers of the largest children hospital in Milan during the pandemic wave. *Infect Control Hosp Epidemiol* 2020; 41: 1468-1469.
- [15] Baveja S, Karnik N, Natraj G, Natkar M, Bakshi A and Krishnan A. Rapid volunteer-based SARS-Cov-2 antibody screening among health care workers of a hospital in Mumbai, India. *Indian J Med Sci* 2020; 72: 148-154.
- [16] Garcia-Basteiro AL, Moncunill G, Tortajada M, Vidal M, Guinovart C, Jiménez A, Santano R, Sanz S, Méndez S, Llupia A, Aguilar R, Alonso S, Barrios D, Carolis C, Cisteró P, Chóliz E, Cruz A, Fochs S, Jairoce C, Hecht J, Lamogliá M, Martínez MJ, Mitchell RA, Ortega N, Pey N, Puyol L, Ribes M, Rosell N, Sotomayor P, Torres S, Williams S, Barroso S, Vilella A, Muñoz J, Trilla A, Varela P, Mayor A and Dobaño C. Seroprevalence of antibodies against SARS-CoV-2 among health care workers in a large Spanish reference hospital. *Nat Commun* 2020; 11: 3500.
- [17] Kumar N, Bhartiya S, Desai S, Mutha A, Beldar A and Singh T. Seroprevalence of antibodies against SARS-CoV-2 among health care work-

Seroprevalence of IgG against SARS-CoV-2 among healthcare workers

- ers in Mumbai, India. *Asia Pac J Public Health* 2020; 1010539520977307.
- [18] Mughal MS, Kaur IP, Patton CD, Mikhail NH, Vareechon C and Granet KM. The prevalence of severe acute respiratory coronavirus virus 2 (SARS-CoV-2) IgG antibodies in intensive care unit (ICU) healthcare personnel (HCP) and its implications-a single-center, prospective, pilot study. *Infect Control Hosp Epidemiol* 2020; 1-2.
- [19] Schmidt SB, Grüter L, Boltzmann M and Rollnik JD. Prevalence of serum IgG antibodies against SARS-CoV-2 among clinic staff. *PLoS One* 2020; 15: e0235417.
- [20] Bleyzac N, Goutelle S, Bourguignon L and Tod M. Azithromycin for COVID-19: more than just an antimicrobial? *Clin Drug Investig* 2020; 40: 683-686.
- [21] Wessels I, Rolles B and Rink L. The potential impact of zinc supplementation on COVID-19 pathogenesis. *Front Immunol* 2020; 11: 1712.
- [22] Craddock JC, Neale EP, Peoples GE and Probst YC. Vegetarian-based dietary patterns and their relation with inflammatory and immune biomarkers: a systematic review and meta-analysis. *Adv Nutr* 2019; 10: 433-451.
- [23] Dowell AC and Turner N. Closing evidence to practice gaps: an end to an attack of the vapours? *Br J Gen Pract* 2016; 66: 118-119.
- [24] SARS-CoV-2 IgG Assay [Internet]. [cited 2020 Jan 8]. Available from: <https://www.siemens-healthineers.com/en-in/laboratory-diagnostics/assays-by-diseases-conditions/infectious-disease-assays/cov2g-assay>.
- [25] Martin C, Montesinos I, Dauby N, Gilles C, Dahma H, Van Den Wijngaert S, De Wit S, Delforge M, Clumeck N and Vandenberg O. Dynamics of SARS-CoV-2 RT-PCR positivity and seroprevalence among high-risk healthcare workers and hospital staff. *J Hosp Infect* 2020; 106: 102-106.
- [26] Sotgiu G, Barassi A, Miozzo M, Sadari L, Piana A, Orfeo N, Colosio C, Felisati G, Davi M, Gerli AG and Centanni S. SARS-CoV-2 specific serological pattern in healthcare workers of an Italian COVID-19 forefront hospital. *BMC Pulm Med* 2020; 20: 203.
- [27] Khalil I and Barma P. Sub-continental atmosphere and inherent immune system may have impact on novel corona virus' 2019 (nCovid-19) prevalence in South East Asia. *Myensingh Med J* 2020; 29: 473-480.
- [28] Yamamoto N and Bauer G. Apparent difference in fatalities between Central Europe and East Asia due to SARS-COV-2 and COVID-19: four hypotheses for possible explanation. *Med Hypotheses* 2020; 144: 110160.
- [29] Didzun O, De Neve JW, Awasthi A, Dubey M, Theilmann M, Bärnighausen T, Vollmer S and Geldsetzer P. Anaemia among men in India: a nationally representative cross-sectional study. *Lancet Glob Health* 2019; 7: e1685-e1694.
- [30] Jonker FA and Boele van Hensbroek M. Anaemia, iron deficiency and susceptibility to infections. *J Infect* 2014; 69 Suppl 1: S23-7.
- [31] Testing Strategy [Internet]. [cited 2020 Jan 9]. Available from: <https://www.icmr.gov.in/ctest-strat.html>.