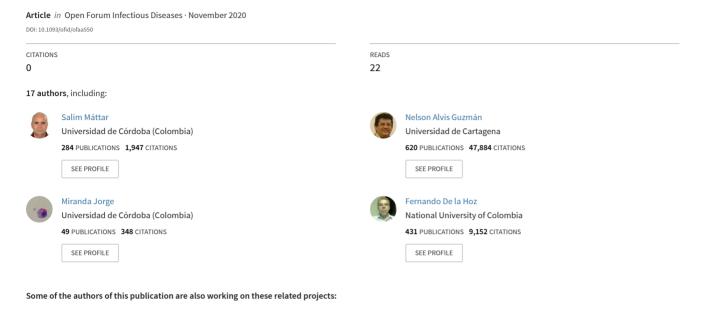
See discussions, stats, and author profiles for this publication at: https://www.researchgate.net/publication/346875102

SARS-CoV-2 seroprevalence among adults in a tropical city of the Caribbean area, Colombia: are we much closer to herd immunity than developed countries?



Jóvenes de hogares pobres en la subregión Montes de María, View project

Characterization of hepatitis B virus in Amerindian children and mothers from Amazonas State, Colombia View project

BRIEF REPORT

# Severe Acute Respiratory Syndrome Coronavirus 2 Seroprevalence Among Adults in a Tropical City of the Caribbean Area, Colombia: Are We Much Closer to Herd Immunity Than Developed Countries?

Salim Mattar,<sup>1,©</sup> Nelson Alvis-Guzman,<sup>2</sup> Evelin Garay,<sup>1</sup> Ricardo Rivero,<sup>1</sup> Alejandra García,<sup>1</sup> Yesica Botero,<sup>1</sup> Jorge Miranda,<sup>1</sup> Ketty Galeano,<sup>1</sup> Fernando de La Hoz,<sup>3</sup> Caty Martínez,<sup>1</sup> Germán Arrieta,<sup>1</sup> Álvaro A. Faccini-Martínez,<sup>1</sup> Camilo Guzmán,<sup>1</sup> Hugo Kerguelen,<sup>4</sup> Maria Moscote,<sup>4</sup> Hector Contreras,<sup>1</sup> and Veronica Contreras<sup>1</sup>

<sup>1</sup>Universidad de Córdoba, Montería, Córdoba, Colombia, <sup>2</sup>Universidad de Cartagena, Fundacion Alzack, Cartagena, Colombia, <sup>3</sup>Universidad Nacional de Colombia, Departamento de Salud Pública, Bogotá, Colombia, <sup>4</sup>Secretaria de Salud de Monteria, Cordoba, Colombia

A serological survey was carried out in Monteria (500 000 population), a mid-size city in Colombia. An overall prevalence of 55.3% (95% confidence interval, 52.5%–57.8%) was found among a sample of 1.368 people randomly selected from the population. Test positivity was related to economic characteristics with the highest prevalence found in the most impoverished areas, representing 83.8% of the city's population. We found a prevalence that might be associated with some important level of population immunity.

**Keywords.** coronavirus; ELISA; infectious disease transmission; poverty areas; socioeconomic status.

Colombia (503 million inhabitants) identified its first case of coronavirus disease 2019 ([COVID-19] severe acute respiratory syndrome coronavirus 2 [SARS-CoV-2] infection) on March 2, 2020, and by October 14, 2020 it had reported 930 159 cases (incidence, 17 539 per 100 000 people) and 28 306 deaths (mortality, 5339 per 100 000 people), with variations for incidence and mortality by department and region [1]. Monteria is a medium-size city (~500 000 people) in the Caribbean region, with a high proportion of the population living in poverty (~80%). Local SARS-CoV-2 circulation was first detected on March 15, 2020, and by October 14, 2020 it had reported 25 208 cases and 1585 deaths (incidence rate, 13 776 per 100 000 people; mortality rate, 15 258 per 100 000 people) (Figure 1).

Received 28 September 2020; editorial decision 31 October 2020; accepted 5 November 2020. Correspondence: Salim Mattar, PhD, Universidad de Córdoba, Instituto de Investigaciones Biológicas del Trópico, Córdoba, Colombia (smattar@correo.unicordoba.edu.co).

Open Forum Infectious Diseases<sup>®</sup>2020



Seroprevalence studies are essential because they allow a more reliable estimation of the population infected after COVID-19 epidemics and estimate the likelihood of a second peak of transmission [2]. Colombia, according to the website https://serotracker.com/data, has no reported seroprevalence studies of SARS-CoV-2.

## METHODS

This study aimed to estimate the prevalence of SARS-CoV-2 past infections in Monteria. A population-based serological study was carried out over a random sample (n = 1368) of the population selected from all city areas. The neighborhood population size weighted the sample. The steps to select the sample were as follows: (1) blocks of houses were randomly selected from every neighborhood (10 neighborhoods); (2) a list of houses in every selected block was developed, identifying every house with a particular number; and (3) house-to-house visits were made in the city's neighborhoods, taking the even numbers. Adjoining houses were not visited. No attempt was made to identify whether participants had antecedents of suspected or confirmed COVID-19 infection or disease, nor whether they had been a contact of the confirmed case(s). The number of samples taken by neighborhoods fluctuated between 55 and 428. The blood samples were taken by health staff trained for epidemiological fieldwork. Once the blood samples were obtained, the tubes were sent to the University of Cordoba laboratory, where they were centrifuged. The enzyme-linked immunosorbent assay (ELISA) test was carried out by microbiologists who are experts in this type of test.

Sera were analyzed using INgezim COVID 19 DR test (Ingenasa; Eurofins, Madrid, Spain), a dual recognition ELISA detecting semiquantitatively total SARS-CoV-2 virus N-protein-specific antibodies (immunoglobulin [Ig]G, IgM, and IgA). The assay was previously validated by both the manufacturer and us [3], with a 90%–93% sensitivity and specificity of 91%–99% (Table 1).

## **Patient Consent Statement**

The research committee of the Institute of Tropical Biological Research of the University of Cordoba approved the ethics protocol, and informed consent was obtained from all patients. Patients were registered using an anonymous numeric code. The study incorporated procedures, management, and conservation of samples, and technical-administrative procedures for health research required by resolution 8430 of the Ministry of Health of Colombia, in 1993 and declaration of Helsinki for ethical and medical research in human subjects. The study was considered as minimal risk.

Published by Oxford University Press on behalf of Infectious Diseases Society of America 2020. This work is written by (a) US Government employee(s) and is in the public domain in the US. This Open Access article contains public sector information licensed under the Open Government Licence v3.0 (http://www.nationalarchives.gov.uk/doc/open-government-licence/version/2/). DOI: 10.1093/ofid/ofaa550

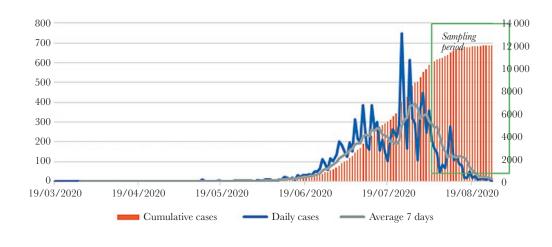


Figure 1.

### RESULTS

From 1368 people (male n = 577, n = female 791), we found an overall prevalence of 55.3% (95% confidence interval [CI], 52.5–57.8), with the most massive prevalence among people between 20 to 60 years old (53.9%; 95% CI, 49.4%–58.3%). Prevalence varied by neighborhood, from 11.5% to 75% (Table 2). Three neighborhoods had prevalence above the average (75%, 70.9%, and 57.1%), and all of them were placed in disadvantaged areas of the city. People living in rural areas had a prevalence under the city average (46.8%; 95% CI, 38.6–68.5) (Table 2). Three were no differences by sex: males had 57.2% of seroprevalence (95% CI, 53.1%–61.3%), and females had 54.4% of seroprevalence (95% CI, 50.8%–58.0%).

The present work is the first serological study reported from Colombia, where SARS-CoV-2 has been circulating for more than 7 months. It yields a similar proportion of infection than the study conducted by Del Brutto et al [4] in Atahualpa (Ecuador), a small rural population, where they found a prevalence of 44%. It is remarkable that these Latin American results are higher than those reported from European cities [5, 6] and China [7], where positivity to serological tests ranged from 0.9% to 13%, suggesting a lower attack rate in the population.

Monteria's high prevalence suggests that there are many people entirely or partially immune to SARS-CoV-2, resulting in some level of population protection against the second wave of infection. During the first wave, Monteria detected 12 226 cases and 709 deaths, and the local hospital nearby could collapse. However, if 55% of the population is already nonsusceptible, the second wave may not occur because the level of population immunity to reach protection against COVID-19 outbreaks is approximately 65% [2].

Low adherence to quarantine recommendation is one of the main reasons behind this unexpected high prevalence. It is not surprising if the socioeconomic conditions of Monteria's population are reviewed. Employment in the informal sector is as high as 60% in the whole city but may be 83.8% in some areas, which forces people to remain on the streets struggling to gain enough to pay for the everyday food and shelter.

#### DISCUSSION

This study may help to clarify the true magnitude of COVID-19 mortality in Monteria. Currently, the city has reported 709 deaths, which amount to a case-fatality rate (CFR) of 5.7%, 2 times the national average CFR. However, if our results are valid, the true CFR would be 0.26% because ~275 000 people would have been infected in Monteria during the first peak of transmission.

The study also has some limitations. First, the recombinant antigen N was used for laboratory testing, which has excellent sensitivity; however, recombinant antigens are not usually specific (3). Although we indeed have a large circulation of SARS-CoV-2, other coronaviruses that could have cross-reactions with the N antigen cannot be ruled out (3). Moreover, crossreactions with arboviruses and SARS-CoV-2 were recently reported [3, 8, 9], and Monteria is an endemic region for dengue, Zika, and Chikungunya. Therefore, false positives could also be possible. In addition, the ELISA used by us detects total gamma globulins (IgM, IgG, IgA), and it is believed that IgA produces false positives [10]. Second, no information about antecedents of COVID-19 nor contact-confirmed cases was collected from participants, which precluded us from analyzing the performance of the test by the presence or absence of clinical symptoms.

The main strength of this study is that it shows, for the first time, the prevalence of SARS-CoV-2 in a small capital city of Colombia using a randomly selected sample. It allows us to extrapolate the sample results to the city's whole population and contribute epidemiological elements for decision making by municipal public health authorities. Vaccination strategies are already being discussed in Colombia, but a high prevalence of natural infection, such as observed in the present study, may preclude vaccines as a population strategy because it may not

Pat	Patient's Group	z	Diagnosis Period	Positive Samples by ELISA COVID-19 Test (%)	Negative Samples by ELISA COVID-19 Test (%)
Patients from Córdoba, Colombia Col	Confirmed SARS-CoV-2 infection <sup>a</sup>	63	June-July 2020	58 (92)	5 (8)
	COVID-19 clinically diagnosed and asymp- tomatic contacts <sup>b</sup>	Ø	July 2020	6 (75)	2 (25)
Act	Acute dengue infection <sup>c</sup>	00	April–December 2019	0 (0)	8 (100)
Re	Recent dengue infection <sup>c</sup>	15	April–December 2019	0 (0)	15 (100)
Act	Acute Zika infection <sup>d</sup>	19	November 2015–February 2016	5 (26)	14 (74)
Pre	Previous Chikungunya infection <sup>e</sup>	6	November 2015–April 2016	0 (0)	9 (100)
Be	Recent exposure to spotted fever group <i>Rickettsiae</i> <sup>f</sup>	വ	February 2013	0 (0)	5 (100)
Data according assay manufacturers (Ingenasa; Co	Confirmed RT-qPCR SARS-CoV-2 infection	162	Ongoing SARS-CoV-2 pandemic	150 (92.6)	12 (7.4)
Eurofins, Madrid, Spain) <sup>g</sup>	Healthy patients/non-SARS-CoV-2 infection	249	Before SARS-CoV-2 pandemic	2 (0.8)	247 (99.2)
Se.	Seasonal coronavirus (229E, NL63, OC43 HKU1) infection	121	No data	0 (0)	121 (100)
Abbreviations: COVID-19, coronavirus disease 2019; ELISA, enzyme-linked immunosorbent assay; RT-qPCR, quantitative reverse-transcription polymerase chain reaction; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2.	zyme-linked immunosorbent assay; RT-qPCR, quantitativ	e reverse-trans	cription polymerase chain reaction; SARS-C	oV-2, severe acute respiratory synd	ome coronavirus 2.
usan manapatrian da ana ana ana ana ana ana ana ana an	or the first 2 weeks of symptoms in 1 nationt and 2 week	s after sympto	which is a nation to		

Table 1. Serology and Cross-Reactivity Using a SARS-CoV-2 Commercial ELISA Test in Patients With Tropical Endemic Diseases From Córdoba, Colombia, and According Assay Manufacturers

2

"Serum samples were collected on the same day as dengue-test (RT-qPCR, NS1 antigen, ELISA immunoglobulin [Ig]M) positive samples.

 $^{\mathrm{d}}\mathsf{Serum}$  samples were collected on the same day as Zika-RT-qPCR-positive samples.

<sup>e</sup>Serum samples were collected the on same day as Chikungunya-ELISA IgG-positive samples.

<sup>1</sup>Serum samples were collected the same day as spotted fever group Rickettsiae-immunofluorescence assay-IgG positive samples.

<sup>9</sup>See https://www.eurofins-technologies.com/ingezim-covid-19-dr.html.

Table 2. Neighborhoods, Socioeconomic Strata, Size of Population, and Results of Seroprevalence, Monteria, Co	olombia
---	---------

Neighborhoods (Socioeconomic Strata)	Population	Negative Tests	Positives Tests	Total Tests	Prevalence (%)	95% CI
1. (Low)	43.000	32	37	69	53.6	41.1-66.1
2. (Low)	17.000	40	19	59	32.2	19.4–43.9
3. (Low)	39.000	53	64	117	54.7	45.2-64.1
4. (Low)	102.000	13	39	52	75.0	62.2-87.7
5. (Middle)	23.000	201	72	273	26.4	20.9–31.7
6. (Low)	63.000	16	39	55	70.9	57.9–83.8
7. (Middle)	5.000	54	7	61	11.5	2.6-20.2
8. (High)	47.000	276	152	428	35.5	3.8–40.1
9. (Low)	15.000	63	84	147	57.1	48.8–55.4
10. Rural (Low)	110.000	55	52	107	48.6	38.6–58.5
Total Overall	464.000	803	565	1368	<sup>°</sup> 55.3	52.5–57.8

Bold text indicates the overall total confidence interval (95% CI).

Abbreviations: CI, confidence interval.

<sup>a</sup>Prevalence adjusted by population in all 10 neighborhoods.

be cost effective. However, the possibility of reinfection suggests that people that have been infected once cannot be considered to be immune. Although so far, long-established reinfection cases seem to be very scarce, and additional verification and longer supplement time are essential to apprehend the length of immunity, transmissibility, and the probability and consequences of reinfection [11]. The present assumption is that clinical management, infection prevention/control, and contact tracing concerns are not expected to diverge for a second infection related to people infected for the first time [11].

### CONCLUSIONS

We do not know whether immunity provided by the natural infection is long-lasting. However, we will soon find out because on September 1, 2020, the national government opened trade, national and international road and air transportation, and restaurants. If the number of cases in Monteria maintains its current decreasing trend, it could be a sign that the city has reached some vital level of protection by natural infection even if it is below 65%, which is proposed as the cutoff point for herd immunity [2]. If we assume that approximately 50% of the population reached natural immunity in the city of Monteria, we could be moderately optimistic, and we are likely to have less occupancy in intensive care unit beds. Moreover, the possible reinfections could be clinically milder.

#### Acknowledgments

*Author contributions.* S. M., N. A.-G., E. G., R. R., A. G., Y. B., F. d. L. H., Á. A. F.-M., H. K., and M. M. design the study. J. M., K. G., C. M., G. A., C. G., H. C., and V. C. carried out the enzyme-linked immunosorbent assay tests and reviewed the article.

*Financial support.* This work was funded by Ministerio de Ciencia Tecnología e Innovación, Colombia: Research project "Fortalecimiento de capacidades instaladas de ciencia y tecnología de la Universidad de Córdoba para atender problemáticas asociadas con agentes biológicos de alto riesgo para la salud humana en el departamento de Córdoba" (Acta 76, May 5, 2020, to Alcaldia de Monteria for support on collecting the blood samples).

**Potential conflicts of interest.** All authors: no reported conflicts of interest. All authors have submitted the ICMJE Form for Disclosure of Potential Conflicts of Interest.

#### References

- De la Hoz-Restrepo F, Alvis-Zakzuk NJ, De la Hoz-Gomez JF, et al. Is Colombia an example of successful containment of the 2020 COVID-19 pandemic? A critical analysis of the epidemiological data, March to July 2020. Int J Infect Dis 2020; 99:522–9.
- Eckerle I, Meyer B. SARS-CoV-2 seroprevalence in COVID-19 hotspots. Lancet 2020; 396:514–5.
- Faccini-Martinez A, Rivero R, Garay E, et al. Serological cross-reactivity using a SARS-CoV-2 ELISA test in acute Zika virus infection, Colombia, Int J Infect Dis 2020; 101:191–3.
- Del Brutto OH, Costa AF, Mera RM, et al. SARS-CoV-2 in rural Latin America. A population-based study in coastal Ecuador. Clin Infect Dis 2020; doi: 10.1093/ cid/ciaa1055/5876901.
- Pollán M, Pérez-Gómez B, Pastor-Barriuso R, et al.; ENE-COVID Study Group. Prevalence of SARS-CoV-2 in Spain (ENE-COVID): a nationwide, populationbased seroepidemiological study. Lancet 2020; 396:535–44.
- Gudbjartsson DF, Helgason A, Jonsson H, et al. Spread of SARS-CoV-2 in the Icelandic population. N Engl J Med 2020; 382:2302–15.
- Xu X, Sun J, Nie S, et al. Author correction: Seroprevalence of immunoglobulin M and G antibodies against SARS-CoV-2 in China. Nat Med 2020; 26:1494.
- Lustig Y, Keler S, Kolodny R, et al. Potential antigenic cross-reactivity between SARS-CoV-2 and dengue viruses. Clin Infect Dis 2020; doi: 10.1093/cid/ciaa1207.
- Spinicci M, Bartoloni A, Mantella A, et al. Low risk of serological cross-reactivity between dengue and COVID-19. Mem Inst Oswaldo Cruz 2020; 115:e200225.
- Hanson KE, Caliendo AM, Arias CA, et al. Infectious Diseases Society of America guidelines on the diagnosis of COVID-19. Clin Infect Dis 2020; doi: 10.1093/cid/ ciaa760.
- 11. European Centre for Disease Prevention and Control. Reinfection with SARS-CoV: considerations for public health response: ECDC; 2020. Available at: https://www.ecdc.europa.eu/sites/default/files/documents/Re-infection-and-viral-shedding-threat-assessment-brief.pdf. Accessed 14 October 2020.