SARS-CoV-2 Community Transmission disproportionately affects Latinx population during Shelter-in-Place in San Francisco

Gabriel Chamie^{1*}, Carina Marquez^{1*}, Emily Crawford^{1,2}, James Peng¹, Maya Petersen³, Daniel Schwab⁴, Joshua Schwab³, Jackie Martinez¹, Diane Jones⁵, Douglas Black¹, Monica Gandhi¹, Andrew D. Kerkhoff¹, Vivek Jain¹, Francesco Sergi¹, Jon Jacobo⁶, Susana Rojas⁶, Valerie Tulier-Laiwa⁶, Tracy Gallardo-Brown⁶, Ayesha Appa¹, Charles Chiu¹, Mary Rodgers⁷, John Hackett Jr.⁷, CLIAhub Consortium², Amy Kistler², Samantha Hao², Jack Kamm², David Dynerman², Joshua Batson², Bryan Greenhouse¹, Joe DeRisi^{1,2}, Diane V. Havlir¹

*Note: Drs. Gabriel Chamie and Carina Marquez contributed equally to this article.

Affiliations

- 1. University of California, San Francisco, CA
- 2. Chan Zuckerberg Biohub, San Francisco, CA
- 3. University of California, Berkeley, CA
- 4. College of the Holy Cross, Worcester, MA
- 5. Unidos en Salud/United in Health San Francisco, CA
- 6. Latino Task Force for COVID-19, San Francisco, CA
- 7. Abbott Laboratories, Abbott Park, IL

© The Author(s) 2020. Published by Oxford University Press for the Infectious Diseases Society of America. All rights reserved. For permissions, e-mail: journals.permissions@oup.com. Corresponding author:

Gabriel Chamie, MD, MPH

Associate Professor of Medicine

Division of HIV, Infectious Diseases & Global Medicine

Repter

University of California, San Francisco / San Francisco General Hospital

INSCR 1

Box 0874, 995 Potrero Avenue

San Francisco, CA 94110

Email: Gabriel.Chamie@ucsf.edu

Phone: 415-476-4082, ext 445

Summary: By providing mass SARS-CoV-2 PCR/antibody testing, regardless of symptoms, in a San Francisco census tract, we determined that infections, most of which were asymptomatic, from diverse lineages continued circulating among low-income, Latinx persons 6-weeks into the city's shelter-in-place ordinance.

Accepted Manuscrit

ABSTRACT

Background: There is urgent need to understand the dynamics and risk factors driving ongoing SARS-CoV-2 transmission during shelter-in-place mandates.

Methods: We offered SARS-CoV-2 reverse transcription-PCR and antibody (Abbott ARCHITECT IgG) testing, regardless of symptoms, to all residents (≥4 years) and workers in a San Francisco census tract (population: 5,174) at outdoor, community-mobilized events over four days. We estimated SARS-CoV-2 point prevalence (PCR-positive) and cumulative incidence (antibody or PCR-positive) in the census tract and evaluated risk factors for recent (PCR-positive/antibody-negative) versus prior infection (antibody-positive/PCR-negative). SARS-CoV-2 genome recovery and phylogenetics were used to measure viral strain diversity, establish viral lineages present, and estimate number of introductions.

Results: We tested 3,953 persons: 40% Latinx; 41% White; 9% Asian/Pacific Islander; and 2% Black. Overall, 2.1% (83/3,871) tested PCR-positive: 95% were Latinx and 52% asymptomatic when tested. 1.7% of census tract residents and 6.0% of workers (non-census tract residents) were PCR-positive. Among 2,598 tract residents, estimated point prevalence of PCR-positives was 2.3% (95%CI: 1.2-3.8%): 3.9% (95%CI: 2.0-6.4%) among Latinx vs. 0.2% (95%CI: 0.0-0.4%) among non-Latinx persons. Estimated cumulative incidence among residents was 6.1% (95%CI: 4.0-8.6%). Prior infections were 67% Latinx, 16% White, and 17% other ethnicities. Among recent infections, 96% were Latinx. Risk factors for recent infection were Latinx ethnicity, inability to shelter-in-place and maintain income, frontline service work, unemployment, and household income <\$50,000/year. Five SARS-CoV-2 phylogenetic lineages were detected.

Conclusion: SARS-CoV-2 infections from diverse lineages continued circulating among low-income, Latinx persons unable to work from home and maintain income during San Francisco's shelter-inplace ordinance. Key Words: community-based SARS-CoV-2 testing; asymptomatic SARS-CoV-2 infection; shelter-in-

place; ethnic disparities; phylogenetic analysis

Accepted Manuschi

Introduction

In early 2020, multiple introductions of SARS-CoV-2 into the United States laid the foundation for the ongoing epidemic that has claimed over 100,000 U.S. lives in less than 6 months.[1] Some of the earliest clinical cases of COVID-19 were recognized in California,[2] and the state led the nation in issuing a state-wide shelter-in-place mandate on March 19.[3] San Francisco declared a local emergency on February 25 and issued a series of increasingly restrictive mandates on sizes of gatherings culminating in a shelter-in place order on March 16. Although peak hospitalization and death rates in San Francisco over the ensuing month were nearly 10-fold lower than hard-hit cities such as New York,[4] the pattern of disproportionately higher hospitalizations among Latinx communities was similar.[5] In San Francisco, 45% of reported COVID-19 cases are among Latinx people, who represent 15% of the city's population, and at San Francisco General Hospital, a large public hospital with a patient population that is 31% Latinx, 81% of COVID-19related hospitalizations from March-May 2020 were among Latinx people.[6, 7]

Hospitalizations and deaths represent a small fraction of total SARS-CoV-2 infections.[8-10] Estimates of the burden of community SARS-CoV-2 infections from direct measurements have been difficult to obtain and compare because symptomatic testing programs capture only a proportion of cases,[11] the recognized symptoms associated with COVID-19 have expanded over time,[12] the assays used to identify infection have variable performance characteristics, and easily accessible testing programs are not in place for some of the most highly affected communities. Data on community transmission and ethnic disparities in SARS-CoV-2 infection, as opposed to COVID-19 disease[13], as well as systematic efforts to determine factors driving these disparities remain limited.[14]

To estimate the point prevalence and cumulative incidence of infection, determine risk factors for ongoing versus prior infection and characterize ongoing transmission and viral strain diversity with phylogenetic analyses, we offered population-based, universal PCR and antibody testing for SARS-CoV-2 infection, regardless of symptoms, to all residents of a densely populated census tract within a majority Latinx community in San Francisco.

Methods

Unidos en Salud is a longitudinal study to characterize SARS-CoV-2 epidemiology and assess impact of public health measures within a US census tract in San Francisco. Six-weeks into the city's shelter-in-place ordinance, we provided cross-sectional, mass, low-barrier SARS-CoV-2 reverse transcription-PCR (RT-PCR) and antibody testing, regardless of symptoms, to residents (\geq 4 years) and people who work but may not reside in the census tract.

Study Setting and Community Mobilization

U.S. census tract 022901 is a population-dense, 16-square-block (0.1 square-mile) area in San Francisco's Mission District, with 5,174 residents of whom 58% are Latinx, 34% White/Caucasian, 5% Asian/Pacific Islander, and 1% Black/African American. Median per capita income was \$40,420/year in 2018, with 34% of households earning <\$50,000/year and 20% earning >\$200,000/year.[15] In partnership with the Latino Task Force for COVID-19, an umbrella organization coordinating local Latinx community-based organizations, we distributed flyers, mobilized the community on local and social media, and offered online and door-to-door preregistration within the census tract for testing appointments during the week prior to the testing campaign. From April 25-28, 2020, we offered outdoor testing at public parks and schools to those who provided an address in the census tract or worked in the tract. On April 28, we expanded eligibility to residents of neighboring city blocks (two blocks north, three blocks east and one block south), responding to high community demand. During pre-registration, we conducted a brief survey. At the time of testing, we obtained verbal consent for participation and conducted COVID-19 symptom screening. Medical staff performed a fingerstick blood collection (500μL) for antibody testing and an oropharyngeal/mid-turbinate nasal swab for quantitative RT-PCR. Participants could opt out of either test. We contacted all PCR-positive persons to disclose results and perform a clinical assessment. We provided household support via a community-led team for PCR-positive participants and evaluated symptoms among all PCR-positive participants over 2 weeks following testing.

Laboratory assays

Swabs were collected in DNA/RNA Shield (Zymo Research) to inactivate virus and preserve RNA stability. RT-PCR of viral N and E genes and human RNAse P gene was performed on extracted RNA at a CLIA-certified laboratory operated by UCSF and the Chan Zuckerberg Biohub using a Laboratory Developed Test with a limit of detection of log₁₀ 4.5 viral genome copies/mL. SARS-CoV-2-positive RNA samples were subjected to Primal-Seq Nextera XT version 2.0,[16] using the ARTIC Network V3 primers,[17] followed by paired-end 2 x 150bp sequencing on an Illumina NovaSeq platform. For antibody testing, the ARCHITECT SARS-CoV-2 IgG Emergency Use Authorization (EUA) assay (Abbott Laboratories, Abbott Park, IL, USA)[18] was performed on participants' plasma from the fingerstick collections, which is a research use of the test.

Study Outcomes

Outcomes included the estimated point prevalence of all PCR-positive infections, recent infections (PCR-positive and antibody-negative) and prior infections (antibody-positive and PCR-negative). Cumulative incidence of infection was defined as any PCR or antibody-positive result. Phylogenetics were used to measure strain diversity.

Statistical analyses

Proportions were compared using chi-squared tests and medians compared using Wilcoxon rank-sum tests. Cumulative incidence of infection was adjusted for RT-PCR and antibody test characteristics; 95% confidence intervals incorporating uncertainty in test characteristics were based on bootstrap. For census tract residents, we further adjusted for differences in age, sex, and race/ethnicity of participants compared to 2018 census estimates (**Supplementary Methods**). We used multivariate logistic regression for the dependent outcome of PCR-positivity among participants tested and included the independent variables sex, frontline worker status, household income and reporting a COVID-19 contact, based on *a priori* interest and univariate results. We did not adjust confidence intervals for multiple testing.

Bioinformatics and genomic analyses

A phylogenetic tree was constructed containing all 123 SARS-CoV-2 genomes from San Francisco County on GISAID[19] on May 22, 2020 together with the high-quality consensus genomes assembled from this study, using the nextstrain toolkit.[20] Global clade identification and naming follows the Nextstrain proposal,[21] and significance of population structure was computed by a permutation test was used for Hudson's FST (**Supplementary Methods**).[22]

Ethics Statement

The UCSF Committee on Human Research determined that the study met criteria for public health surveillance.

Role of the Funding Source

The study was supported by the Chan Zuckerberg Biohub, UCSF, and a Program for Breakthrough Biomedical Research award. ARCHITECT SARS-CoV-2 test kits were provided by Abbott Laboratories. The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Results

Testing Uptake and Coverage

Overall, 3,953 persons were tested during the campaign. Of all persons tested, 53% were male, and 40% identified as Latinx, 41% White, 9% Asian/Pacific Islander, 2% Black, and 7% other/mixed ethnicity (**Table 1**). A significantly higher proportion of Latinx compared to non-Latinx participants reported annual household income <\$50,000/year (62% vs. 19%), living with >5 persons/household (25% vs. 5%) and working frontline service jobs (43% vs. 17%, **Supplementary Table S1**). Estimated census tract testing coverage of adult residents (age ≥20 years) was 60%.

SARS-CoV-2 Infection by PCR testing

Among 3,871 tested by SARS-CoV-2 RT-PCR, 2.1% (83 people) tested PCR-positive: 1.7% (43/2,598; 95%CI: 1.2%-2.2%) of census tract residents, 6.0% (27/450; 95%CI: 4.0%-8.6%) of tract

workers and 1.6% (13/823; 95%CI: 0.8%-2.7%) of residents of neighboring city blocks. Among all persons tested, 237 (6.1%) reported symptoms compatible with COVID-19 of whom 31 (13.1%) tested PCR-positive. Twelve people (0.3% of all persons tested) reported having had a prior positive test for COVID-19, of whom five tested PCR-positive.

Among PCR-positive persons, 95% identified as Latinx, median age was 38 years (interquartile range [IQR]: 28-50 years), and 76% were male. Persons testing PCR-positive were significantly more likely than persons testing PCR-negative to identify as Latinx, report inability to shelter-in-place and maintain income, work frontline-service jobs or be unemployed, and live in households with income <\$50,000/year and \geq 3 persons/household (**Table 2**). Given that 95% of PCRpositive persons were Latinx, we limited our multivariate model to Latinx participants to evaluate risk factors for PCR-positivity within this group, and found significantly higher odds of PCR-positive infection if male (OR: 2.0, 95% CI: 1.1-3.6, p=0.02), working a frontline service job (OR: 2.6, 95% CI: 1.4-5.1, p=0.004, ref: non-frontline), household income <\$50,000/year (OR: 8.9, 95% CI: 1.9-158, p=0.03, ref: >\$100,000/year), or reporting a COVID-19 contact (OR: 3.6, 95%CI: 2.0-6.3, p<0.001).

Estimated point prevalence of PCR-positive infection in the census tract after adjusting for age and sex of participants in the testing campaign versus census demographics was 2.3% (95% CI: 1.2%-3.8%): 3.9% (95% CI: 2.0%-6.4%) among Latinx vs. 0.2% (95% CI: 0.0%-0.4%) among non-Latinx tract residents. Among Latinx people who worked in the census tract, unadjusted point prevalence of PCR-positive infection was 10.4% (95% CI: 7.0%-14.8%), compared to 0.0% (95% CI: 0.0%-2.0%) among non-Latinx workers.

Clinical characteristics of PCR-positive persons

Among 83 PCR-positive persons, 43 (52%) were asymptomatic at the time of testing. Twoweek follow-up was obtained for 41 participants who were asymptomatic at the time of testing:

11

8/41 (20%) recalled mild symptoms that had resolved by the time of testing, 10 (24%) developed symptoms after testing (pre-symptomatic) and 23 (56%) remained asymptomatic. Based on reclassified symptom status among PCR-positive people, 39/80 (49%) were symptomatic at time of testing, 8 (10%) were previously symptomatic, 10 (12.5%) were pre-symptomatic, and 23 (29%) remained asymptomatic throughout infection. One PCR-positive person (1.3%) required hospitalization.

SARS-CoV-2 Cumulative Incidence and Recent vs. Prior Infections

Among 3,861 participants tested for SARS-CoV-2 antibodies, 3.4% (131) tested Ab-positive: 3.1% (80/2,545; 95%CI: 2.5%-3.9%) among census tract residents compared to 7.7% (34/442; 95%CI: 5.4%-10.6%) among tract workers and 2.1% (17/829; 95%CI: 1.2%-3.3%) among adjacent city block residents. Estimated cumulative incidence (Ab or PCR-positive) among tract residents was 4.4% (95% CI: 3.2%-5.6%) after adjusting for test characteristics and 6.1% (95%CI: 4.0%-8.6%) after further adjusting for participation (**Supplementary Table S2**).

Among all infections detected by PCR or Ab, 26% (48/182) were recent infection. 53% (96/182) were prior infection. Of the remaining infections, 18% (32/182) were PCR-positive/Abpositive, and 3% (6/182) had PCR or antibody testing alone (**Supplementary Table S3**). Whereas prior infections occurred across a range of ethnic groups, employment types and household income levels, recent infections were predominantly among persons who identified as Latinx, worked frontline service jobs and lived in households with income <\$50,000/year (**Figure 1**). Compared to individuals with prior infection, people with recent infection were significantly more likely to be of Latinx ethnicity (OR 10.1; 95% CI: 2.81-64.6, p=0.002, ref: non-Latinx ethnicity), report inability to shelter-in-place and maintain income (OR: 3.18; 95%CI: 1.10-11.6, p=0.048), work frontline service jobs (OR: 2.83; 95%CI: 1.21-6.93, p=0.019) or be unemployed (OR: 3.47; 95%CI: 1.08-11.3, p=0.035, ref: non-frontline service jobs), and live in households with income <\$50,000/year (OR: 19.3; 95%CI: 3.74-356, p=0.005; ref: household income >\$100,000/year; **Supplementary Table S4**).

SARS-CoV-2 RNA levels and phylogenetic analysis

Median levels of virus as estimated by RT-PCR cycle thresholds were significantly higher among PCR-positive/Ab-negative persons compared to PCR-positive/antibody-positive persons, supporting our classification of recent infection (**Figure 2**). Among recently-infected individuals, median levels of virus by RT-PCR cycle threshold did not differ significantly between symptomatic (24, IQR: 19-25, range 11-35; N=27) and asymptomatic (24, IQR: 19-26, range 16-32; N=10; p=0.98) persons (**Figure 2**); additional comparisons by subgroup are in **Supplementary Figure S1**.

We recovered SARS-CoV-2 genomes from 59% (49/83) of the PCR-positive RNA samples. The recovered genomes were diverse and phylogenetically intermixed with samples from across San Francisco, including representatives from five globally circulating clades, showing multiple independent introductions (**Figure 3, Panel A**). Overall, 58% of PCR-positive participants shared a home with another PCR-positive participant identified in the testing campaign: sequences from such households were consistent with within-household transmission (**Figure 3, Panel B**), with no variants detected in 65% of household links (N = 11/17, 95% Cl 41%-83%). We found no significant population structure separating the Mission district samples from the rest of San Francisco (p=0.19).

Discussion

We found stark ethnic and economic disparities in who is at risk for ongoing infection sixweeks into San Francisco's shelter-in-place ordinance. The estimated point prevalence of PCRpositive infection among Latinx residents (3.9%) was twenty times that of non-Latinx residents (0.2%). Perhaps even more striking was that recent infections were concentrated almost exclusively among low-income, Latinx people working frontline jobs, whereas prior infections occurred among more ethnically and economically diverse individuals. In addition, the majority of PCR-positive infections were asymptomatic at the time of testing, and recently infected individuals had high levels of virus regardless of symptoms. These data show that San Francisco's COVID-19 epidemic has continuing transmission in subgroups of the city population that require urgent attention.

Heterogeneity among populations most affected by the COVID-19 epidemic exists across the U.S., within states and cities, and even within neighborhoods as shown here. Population-level epidemiologic data coupled with phylogenetic analyses can help identify, track and inform testing strategies, public health policies and measures to mitigate health and economic effects. Low-barrier, community-mobilized testing is foundational to these efforts. We sought to overcome some of the testing barriers in this census tract through our partnership with the community-led Latino Task Force in San Francisco, who provided explanations about COVID-19 and communicated the importance of testing to the community. Through this approach, we were able to test a large proportion of the population in a short period of time. To date, there are limited published data on population-level PCR testing in the US. Our estimated point prevalence of PCR-positive infection among residents (2.3%) was higher than an estimate (1.74%) from a statewide random population sample of persons aged \geq 12 years in Indiana also obtained in late April 2020, [23] and notably higher than that of a community-wide testing study in the coastal town Bolinas, in the San Francisco Bay Area, in which no PCR-positive cases were identified.[24]

14

We determined that during shelter-in-place, COVID-19 transmission became increasingly concentrated in Latinx community members. The risk factors driving recent transmission among Latinx residents in our study were largely economic and highly correlated: low-income residents working frontline jobs who could not shelter-in-place and maintain their income. Unemployed residents may have experienced increased risk of PCR-positive infection as a consequence of seeking employment, particularly among day laborers. Given the evidence of within household transmission, confirmed by SARS-CoV-2 sequencing, we suspect that transmission was amplified in Latinx multigenerational and multi-family households – a byproduct of skyrocketing rental costs in the city.[25] These economic drivers and ethnic disparities observed at the community level here are reflected in COVID-19 hospitalizations and deaths widely reported in the US, in many settings disproportionately affecting Black/African American people.[13, 26, 27]

We observed high sequence diversity of SARS-CoV-2 infections in the census tract, similar to the diversity seen in San Francisco more broadly, suggestive of multiple independent introductions over time. Our data suggest that most recent infections during shelter-in-place are due to acquisition of virus when working in the census tract, with subsequent within-home transmission in highdensity, low-income households. These findings should help dispel common, dangerous pitfalls in interpreting ethnic disparities in infection, such as biological explanations, supposed community behaviors or stigmatizing communities as transmission "hot spots" about which others have cautioned.[14]

Our results also highlight the importance of SARS-CoV-2 testing in both symptomatic and asymptomatic individuals. Symptom-based testing would have failed to detect over 40% of PCR-positive infections in this community, many of whom had high levels of virus. Overall, 29% of PCR-positive infections never developed symptoms, slightly lower than the proportion found (43%) in a population-based SARS-CoV-2 screening study from Iceland.[28] In our study, recently infected people had high levels of virus that did not differ significantly based on symptoms, a finding in line

with a recent cohort study from the Republic of Korea.[29] The clear implication of these findings is that testing strategies limited to symptomatic individuals and those seeking testing at health centers alone, will fail to limit transmission.

These results have several implications moving forward as shelter-in-place restrictions are lifted. First, more efforts are needed to address uncontrolled epidemics among sub-populations, especially vulnerable populations such as the Latinx community highlighted here. Expanded lowbarrier testing, that is targeted, community-led and mobilized, and not limited to symptomatic persons, is needed. Testing needs to be coupled with social protection of job security and economic support for self-isolation and quarantine (i.e. "test and respond") and culturally responsive contact tracing. Our testing campaign contributed to policy change in San Francisco, with the mayor announcing on May 4, 2020 that essential workers would be eligible for free SARS-CoV-2 testing regardless of symptoms,[30] and then on May 28, 2020 that low-wage workers with COVID-19 would be provided funds to stay home and isolate ("Right to Recover").[31] In parallel, longitudinal, population-based cohorts that couple epidemiologic data with PCR and antibody testing and viral sequencing can provide evidence of effectiveness of public health measures and viral introductions over time, enabling evidence-based responses in a dynamic landscape.

Our study has several limitations. SARS-CoV-2 PCR tests do not detect all cases and antibody sensitivity may be lower in asymptomatic infection which could have resulted in underestimation of cumulative incidence. False positive antibody results could result in overestimation of cumulative incidence and misclassification of prior infections. Misclassification of prior infections could also result from infections that occurred, generated antibodies and resolved to PCR-negative over the six weeks following the shelter-in-place order. Fingerstick sampling could also impact antibody test performance. However, the EUA antibody test we used has been shown to have a high sensitivity (96.9-100%) at \geq 17-22 days post symptom onset, and high specificity (\geq 99.6%) with venous drawn plasma,[32, 33] and our estimates of cumulative incidence accounted for sensitivity and specificity of

16

the PCR and antibody assays used. Second, selection bias in who chose to test may have affected our estimates. Although we adjusted for demographic differences between participants and community composition based on 2018 American Community Survey data, these data may not fully reflect tract demographics in 2020. However, population-based testing in a census tract allowed for greater clarity in understanding who did not participate. Lastly, we relied on self-reported symptoms and survey responses, which may have resulted in misclassification. With follow-up of PCR-positive participants over two weeks, we were able to further explore symptom status, allowing for monitoring and reclassification.

In conclusion, improving access to SARS-CoV-2 testing, regardless of symptoms, through community-led, low-barrier testing programs in vulnerable communities is feasible. Adding economic support and protections for low-income workers during isolation and quarantine could further increase testing and contribute to reducing community transmission and the massive disparities in SARS-CoV-2 infection observed in the U.S.

cepter

NOTES

Acknowledgements

We thank the community members from the census tract studied for their generous participation in our study during San Francisco's shelter-in-place ordinance. We gratefully acknowledge the contributions of all of the study volunteers who made community-based testing possible, and all the members of the Latino Task Force for COVID-19 who led community mobilization efforts, community outreach and education and dissemination of our findings to the community. We thank Chesa Cox, MPH for logistical and administrative support, Stacie Powers for her generous donation of the Brava Theater space to operate as headquarters for the study volunteers and the Community Wellness Team. We thank San Francisco Recreation and Parks and the San Francisco Unified School District for generously allowing us to offer community-based testing at two public parks and two public schools. We thank Supervisor Hillary Ronen, the San Francisco Mayor's Office, Dr. Grant Colfax Director of San Francisco Department of Public Health, Elaine Forbes, Executive Director, Port of San Francisco as well as Naomi Kelly, SF City Administrator, Dr. Susan Ehrlich CEO of Zuckerberg San Francisco General Hospital and the Ward 86 Clinic staff for their efforts in support of this community-based testing project. We thank Andrew Kobylinski and Primary tech for assistance in their virtual support platform and on-line results reporting. We also wish to thank Ana Vallari, Ana Olivo, Barb Harris, and Chris Lark from Abbott Laboratories for helping conduct the antibody assays for this study.

Funding

This work was supported by the Chan Zuckerberg Biohub, the University of California, San Francisco (UCSF), and a Program for Breakthrough Biomedical Research award. ARCHITECT SARS-CoV-2 test kits were provided by Abbott Laboratories.

Potential Conflicts of Interest:

Certer

Mary Rodgers and John Hackett Jr. are employees and shareholders of Abbott Laboratories. Charles Y. Chiu is the director of the UCSF-Abbott Viral Diagnostics and Discovery Center (VDDC) and receives research support funding from Abbott Laboratories. M.G. reports grants from the NIH. V.J. reports grants from CDC/PEPFAR, outside the submitted work. D.H. reports grants from National Institutes of Health and The Gates Foundation, outside the submitted work. J.D. reports grants from Chan Zuckerberg Biohub, outside the submitted work. G.C. reports grants from National Institutes of Health and The Gates Foundation, outside the submitted work. All other authors have no potential conflicts.

References

- Schuchat A, Team CC-R. Public Health Response to the Initiation and Spread of Pandemic COVID-19 in the United States, February 24-April 21, 2020. Mmwr 2020; 69(18): 551-6.
- Heinzerling A, Stuckey MJ, Scheuer T, et al. Transmission of COVID-19 to Health Care Personnel During Exposures to a Hospitalized Patient - Solano County, California, February 2020. Mmwr 2020; 69(15): 472-6.
- <u>https://covid19.ca.gov/stay-home-except-for-essential-needs/</u>. Last accessed on June 1, 2020.
- 4. <u>https://data.sfgov.org/stories/s/San-Francisco-COVID-19-Data-and-Reports/fjki-2fab/</u>Last accessed on June 1, 2020.
- Wadhera RK, Wadhera P, Gaba P, et al. Variation in COVID-19 Hospitalizations and Deaths Across New York City Boroughs. JAMA 2020.
- Palomino J, Sanchez T. Latinos' coronavirus burden.
 <u>https://www.sfchronicle.com/bayarea/article/Bay-Area-Latinos-hit-hardest-by-coronavirus-</u> 15252632.php. Last accessed on June 4, 2020. San Francisco Chronicle. **2020** May 8, 2020.
- 7. <u>https://data.sfgov.org/stories/s/w6za-6st8</u>. Accessed August 8.
- Verity R, Okell LC, Dorigatti I, et al. Estimates of the severity of coronavirus disease 2019: a model-based analysis. The Lancet infectious diseases 2020; 20(6): 669-77.
- Rosenberg ES, Dufort EM, Blog DS, et al. COVID-19 Testing, Epidemic Features, Hospital
 Outcomes, and Household Prevalence, New York State-March 2020. Clin Infect Dis 2020.
- 10. <u>https://www.cdc.gov/coronavirus/2019-ncov/hcp/planning-scenarios.html</u>. Accessed August 8.
- Gandhi M, Yokoe DS, Havlir DV. Asymptomatic Transmission, the Achilles' Heel of Current Strategies to Control Covid-19. The New England journal of medicine **2020**; 382(22): 2158-60.

- 12. <u>https://www.cdc.gov/coronavirus/2019-ncov/symptoms-testing/symptoms.html</u>; last accessed on June 4, 2020.
- Price-Haywood EG, Burton J, Fort D, Seoane L. Hospitalization and Mortality among Black
 Patients and White Patients with Covid-19. The New England journal of medicine 2020.
- Chowkwanyun M, Reed AL, Jr. Racial Health Disparities and Covid-19 Caution and Context.
 The New England journal of medicine **2020**.
- 15. <u>https://censusreporter.org/profiles/14000US06075022901-census-tract-22901-san-</u> <u>francisco-ca/</u> Last accessed on June 1, 2020.
- Quick J, Grubaugh ND, Pullan ST, et al. Multiplex PCR method for MinION and Illumina sequencing of Zika and other virus genomes directly from clinical samples. Nat Protoc 2017; 12(6): 1261-76.
- 17. <u>https://artic.network/resources/ncov/ncov-amplicon-v3.pdf</u>; last accessed on June 3, 2020.
- 18. <u>https://www.fda.gov/media/137383/download;</u> last accessed on June 4, 2020.
- 19. Elbe S, Buckland-Merrett G. Data, disease and diplomacy: GISAID's innovative contribution to global health. Glob Chall **2017**; **1**(1): 33-46.
- 20. Neher RA, Bedford T. nextflu: real-time tracking of seasonal influenza virus evolution in humans. Bioinformatics **2015**; 31(21): 3546-8.
- 21. <u>https://virological.org/t/year-letter-genetic-clade-naming-for-sars-cov-2-on-nextstain-org/498</u>; last accessed on June 5, 2020.
- Hudson RR, Slatkin M, Maddison WP. Estimation of levels of gene flow from DNA sequencedata. Genetics 1992; 132(2): 583-9.
- Menachemi N, Yiannoutsos CT, Dixon BE, et al. Population Point Prevalence of SARS-CoV-2
 Infection Based on a Statewide Random Sample Indiana, April 25-29, 2020. Mmwr 2020;
 69(29): 960-4.

- 24. Appa A, Takahashi S, Rodriguez-Barraquer I, et al. Universal SARS-CoV-2 PCR and Antibody Testing for a Rural Community. International AIDS Society (IAS) COVID-19 Conference. San Francisco, **2020**.
- Joint Center for Housing Studies (JCHS). The state of the nation's housing 2017 (Report).
 Cambridge, MA: Harvard University, Joint Center for Housing Studies, 2017.
- 26. CDC COVID-NET. Characteristics of Laboratory-Confirmed COVID-19-Associated Hospitalizations. <u>https://gis.cdc.gov/grasp/COVIDNet/COVID19_5.html</u> Last accessed on August 3, 2020.
- 27. Suleyman G, Fadel RA, Malette KM, et al. Clinical Characteristics and Morbidity Associated With Coronavirus Disease 2019 in a Series of Patients in Metropolitan Detroit. JAMA Netw Open **2020**; 3(6): e2012270.
- Gudbjartsson DF, Helgason A, Jonsson H, et al. Spread of SARS-CoV-2 in the Icelandic Population. The New England journal of medicine 2020.
- 29. Lee S, Kim T, Lee E, et al. Clinical Course and Molecular Viral Shedding Among Asymptomatic and Symptomatic Patients With SARS-CoV-2 Infection in a Community Treatment Center in the Republic of Korea. JAMA Intern Med **2020**.
- 30. <u>https://www.nbcbayarea.com/news/local/san-francisco/san-francisco-now-offering-free-</u> covid-19-testing-for-all-essential-workers/2284712/; Last accessed on June 4, 2020.
- Fracassa D. SF to pay low-wage workers who get COVID-19 to stay home and isolate.
 https://www.sfchronicle.com/bayarea/article/SF-rolls-out-new-program-to-help-low-income-15299045.php; Last accessed on June 4, 2020. San Francisco Chronicle. 2020 May 28, 2020.
- Bryan A, Pepper G, Wener MH, et al. Performance Characteristics of the Abbott Architect SARS-CoV-2 IgG Assay and Seroprevalence in Boise, Idaho. Journal of clinical microbiology 2020.

33. Ng DL, Goldgof GM, Shy BR, et al. SARS-CoV-2 seroprevalence and neutralizing activity in donor and patient blood from the San Francisco Bay Area. medRxiv (Preprint server) **2020**.

Accepted Manuschi

Tables

Table 1. Characteristics of persons participating in a population-based SARS-CoV-2 testing campaignin the study census tract (US Census tract 022901).

	Residents	Workers ¹	Adjacent City	Total
			Block Residents	
N	2,653	460	840	3,953
Sex at birth, N (%) ²				K
Male	1236 (53%)	284 (62%)	418 (51%)	1938 (53%)
Female	1118 (47%)	173 (38%)	408 (49%)	1699 (47%)
Age category (years)				
4-10	78 (3%)	12 (3%)	28 (3%)	118 (3%)
11-17	100 (4%)	18 (4%)	23 (3%)	141 (4%)
18-50	1695 (64%)	278 (60%)	559 (67%)	2532 (64%)
51-70	633 (24%)	133 (29%)	185 (22%)	951 (24%)
>70	147 (6%)	19 (4%)	45 (5%)	211 (5%)
Race/Ethnicity ²				<u> </u>
White/Caucasian	925 (40%)	112 (25%)	433 (53%)	1470 (41%)
Hispanic/Latinx	934 (40%)	265 (59%)	251 (31%)	1450 (40%)
Black/African	59 (3%)	15 (3%)	11 (1%)	85 (2%)
American				
Asian/Pacific	239 (10%)	38 (8%)	55 (7%)	332 (9%)
Islander				
Other	166 (7%)	22 (5%)	64 (8%)	252 (7%)

Occupation ²				
Frontline Service Jobs				
Food/Beverage	176 (8%)	125 (28%)	70 (9%)	371 (11%)
Healthcare	128 (6%)	21 (5%)	43 (5%)	192 (5%)
Tradesperson (e.g.	244 (11%)	76 (17%)	75 (9%)	395 (11%)
construction, plumbing)				X
or				Q
Cleaning (e.g. janitor,			C	
housekeeper)/personal			S	
services (e.g.			5	
hairdresser)				
Non-frontline Service Jobs		NO		
Education	119 (5%)	13 (3%)	53 (6%)	185 (5%)
Finance, Sales &	486 (21%)	31 (7%)	155 (19%)	672 (19%)
Technology	0			
Student	191 (8%)	34 (8%)	71 (9%)	296 (8%)
Retired/homemaker	219 (10%)	23 (5%)	76 (9%)	318 (9%)
Unemployed	160 (7%)	27 (6%)	50 (6%)	237 (7%)
Other occupation	542 (24%)	91 (21%)	230 (28%)	863 (24%)
Household (HH) informatior	2	1		
Number of people/HH				
1-2	807 (36%)	125 (31%)	338 (41%)	1270 (37%
3-5	1092 (49%)	212 (53%)	417 (51%)	1721 (50%

>5	313 (14%)	66 (16%)	68 (8%)	447 (13%)
Rooms/HH				
1-2	684 (32%)	154 (39%)	260 (32%)	1098 (33%)
3-4	898 (42%)	174 (44%)	338 (42%)	1410 (43%)
>4	535 (25%)	64 (16%)	208 (26%)	807 (24%)
Annual income/HH				
<\$50,000/year	819 (36%)	253 (58%)	205 (25%)	1277 (36%)
\$50,000-100,000/year	604 (27%)	105 (24%)	202 (25%)	911 (26%)
>\$100,000/year	831 (37%)	78 (18%)	407 (50%)	1316 (38%)
Homeless	43 (2%)	12 (3%)	5 (1%)	60 (2%)
Past Medical History ²				
Chronic lung disease	287 (12%)	54 (12%)	113 (14%)	454 (13%)
Chronic heart disease	120 (5%)	24 (5%)	35 (4%)	179 (5%)
Hypertension	325 (14%)	76 (17%)	81 (10%)	482 (13%)
Diabetes	148 (6%)	32 (7%)	33 (4%)	213 (6%)
Smoker ³	540 (23%)	103 (23%)	174 (21%)	817 (23%)
COVID-19-related history				
Reported prior positive test	5 (0.2%)	1 (0.2%)	6 (0.7%)	12 (0.3%)
for COVID-19				
Reported having a personal	175 (8%)	72 (16%)	69 (8%)	316 (9%)
contact diagnosed with				
COVID-19				

Reported being able to	993 (47%)	79 (19%)	377 (49%)	1449 (44%)
shelter-in-place & maintain				
income				

¹Includes workers and family members of people who reported working in the census tract.

ide ²Excludes testing campaign participants with missing responses (i.e. not provided) during the testing

 Table 2. Characteristics of SARS-CoV-2 testing campaign participants who tested PCR-positive

compared to PCR-negative, and factors associated with increased odds of PCR-positivity.

	PCR-	PCR-	Univariate risk of	P value
	positive	negative	PCR-positivity	
	(N=83)	(N=3,788)	(OR, 95% CI)	
Sex at birth (%) ¹				X
Female	20 (24%)	1638 (47%)	Ref	$\mathbf{S}_{\mathbf{r}}$
Male	63 (76%)	1845 (53%)	2.71 (1.64-4.69)	<0.001
Age category (years)			S	
4-10	4 (5%)	104 (3%)	Ref	
11-17	2 (2%)	127 (3%)	0.41 (0.06-2.14)	0.3
18-50	60 (72%)	2429 (64%)	0.64 (0.26-2.15)	0.4
51-70	14 (17%)	923 (24%)	0.39 (0.14-1.41)	0.11
>70	3 (4%)	205 (5%)	0.38 (0.07-1.76)	0.2
Race/Ethnicity ¹			Ref: non-	
White/Caucasian	1 (1%)	1441 (42%)	Hispanic/Latinx	
Hispanic/Latinx	79 (95%)	1348 (39%)		<0.001
Asian/Pacific Islander	2 (2%)	324 (9%)	28.3 (11.7-93.1)	
Other	1 (1%)	326 (9%)		
Occupation ¹				
Frontline Service Job	47 (64%)	902 (27%)	6.56 (3.86-11.6)	<0.001
Non-frontline Service Job	18 (24%)	2267 (67%)	Ref	
Unemployed	9 (12%)	220 (6%)	5.15 (2.18-11.3)	<0.001

Number of				
people/household ¹				
1-2	10 (14%)	1235 (37%)	Ref	
3-5	41 (58%)	1645 (50%)	3.08 (1.60-6.53)	0.002
>5	20 (28%)	420 (13%)	5.88 (2.79-13.2)	<0.001
Annual Household				X
Income ¹				5
>\$100,000	2 (3%)	1287 (38%)	Ref	
\$50-100,000	7 (9%)	889 (26%)	5.07 (1.22-34.1)	0.043
<\$50,000	65 (88%)	1182 (35%)	35.4 (11.1-216)	<0.001
Past Medical History ¹		. 0		
Any underlying conditions	22 (29%)	971 (28%)	1.02 (0.61-1.66)	0.94
COVID-19-related				
history ¹	Ċ			
Personal contact	25 (32%)	286 (8%)	5.29 (3.19-8.57)	<0.001
diagnosed with COVID-19			<i>Ref</i> : no contacts	
Able to shelter-in-place				
and maintain income				
Yes	5 (7%)	1417 (45%)	Ref	<0.001
Νο	64 (93%)	1756 (55%)	10.3 (4.58-29.6)	

¹Excludes testing campaign participants with missing responses (i.e. not provided) during the testing

survey.

FIGURE LEGENDS

Figure 1: Characteristics of prior (PCR-negative and antibody-positive) versus recent (PCR-positive, antibody-negative) SARS-CoV-2 infections among persons participating in a population-based SARS-CoV-2 testing campaign in a densely populated census tract in San Francisco.

Figure 2: Quantitative levels of virus among participants with PCR-positive SARS-CoV-2 infections (N=80) by classification as asymptomatic, pre-symptomatic, and symptomatic for COVID-19 disease as determined over longitudinal follow-up (2 weeks post-testing), and stratified by antibody status with PCR+/Ab- persons considered consistent with recent infection.

Figure 3. Viral genomic diversity among PCR-positive participants. Panel A: phylogenetic tree containing Mission district samples (red) and other San Francisco samples (grey), x-axis marks the number of mutations with respect to the reference genome from Wuhan. Yellow arrows mark introductions of five major global clades (right brackets) to the study population. Panel B: tree subset to Mission district samples. Shape indicates district resident or worker and color indicates antibody status. Households with multiple PCR-positive persons are drawn in green and include markers for samples from which genomes could not be recovered. Asterisk marks a household outside of the district in which an unhoused person in the district spent time.

Figure 1

Race/Ethnicity						Black	Asian	Oth
Prior Infection (PCR-, AB+)	Latinx		67%	v	Vhite 16%	4%	6%	7%
Recent Infection (PCR+, AB-)	Latinx		96%					Asian 4%
Employment type	2							
Prior Infection (PCR-, AB+)	Frontline Job	36%	Non-Frontline Job	53%			Unemp 12'	
Recent Infection (PCR+, AB-)	Frontline Job	53%		Non-Frontline Job		Unem	20%	
Household Incom	ie							
Prior Infection (PCR-, AB+)	< \$50K	47%	\$50K	-100K 27%	> \$1	.00К 2	6%	
Recent Infection (PCR+, AB-)	< \$50K		88%			\$!	50K-100 10%	K > \$10 3%

Receiption .

Flgure 2



