

UCLA

UCLA Previously Published Works

Title

SARS-CoV-2 Infection Among People Living With HIV Compared With People Without HIV: Survey Results From the MACS-WIHS Combined Cohort Study

Permalink

<https://escholarship.org/uc/item/8c34p2gk>

Journal

J AIDS Journal of Acquired Immune Deficiency Syndromes, 89(1)

ISSN

1525-4135

Authors

D'Souza, Gypsyamber
Tong, Weiqun
Gustafson, Deborah
[et al.](#)

Publication Date

2022

DOI

10.1097/qai.0000000000002822

Peer reviewed



Published in final edited form as:

J Acquir Immune Defic Syndr. 2022 January 01; 89(1): 1–8. doi:10.1097/QAI.0000000000002822.

SARS-CoV-2 Infection Among People Living With HIV Compared to People Without HIV: Survey Results From The MACS-WIHS Combined Cohort Study

Gypsyamber D'Souza¹, Weiqun Tong¹, Deborah Gustafson², Maria L. Alcaide³, Cecile D. Lahiri⁴, Anjali Sharma⁵, Audrey L. French⁶, Frank J Palella⁷, Mirjam-Colette Kempf⁸, Matthew J. Mimiaga⁹, Catalina Ramirez¹⁰, Seble Kassaye¹¹, Charles R. Rinaldo¹², Todd T. Brown¹³, Phyllis C. Tien¹⁴, Adaora A. Adimora¹⁵

¹Department of Epidemiology, Johns Hopkins Bloomberg School of Public Health, Baltimore MD, USA

²Department of Neurology, State of New York Downstate Health Sciences University, Brooklyn, NY, USA

³Department of Medicine, Division of Infectious Diseases, University of Miami Miller School of Medicine, Miami, FL, USA

⁴Department of Medicine, Division of Infectious Diseases, Emory University School of Medicine, Atlanta, GA, USA

⁵Department of Medicine, Albert Einstein College of Medicine, Bronx, NY, USA

⁶Department of Medicine, CORE Center/Stroger Hospital of Cook County, Chicago IL

⁷Department of Medicine, Northwestern University Feinberg School of Medicine, Chicago, IL, USA

⁸Schools of Nursing, Public Health and Medicine, University of Alabama at Birmingham, Birmingham, AL

⁹Department of Epidemiology, Fielding School of Public Health, UCLA, Los Angeles 90095-1772

Correspondence: Gypsyamber D'Souza, Johns Hopkins School of Public Health, 615 N Wolfe St., Baltimore, MD 21205, P: 410-502-2583, gdsouza2@jhu.edu.

Author Contributions:

Study conception: G DSouza, D Gustafson

Data Analysis: G DSouza, W Tong

Data Acquisition: D Gustafson, M Alcaide, C Lahiri, A Sharma, A French, F Palella, MC Kempf, M Mimiaga, C Ramirez, S Kassaye, C Rinaldo, T Brown, P Tien, A Adimora

Paper Writing: G DSouza, W Tong, D Gustafson, A Adimora

Data Interpretation: D Gustafson, M Alcaide, C Lahiri, A Sharma, A French, F Palella, MC Kempf, M Mimiaga, C Ramirez, S Kassaye, C Rinaldo, T Brown, P Tien, A Adimora

Manuscript Editing and approval of final version: D Gustafson, M Alcaide, C Lahiri, A Sharma, A French, F Palella, MC Kempf, M Mimiaga, C Ramirez, S Kassaye, C Rinaldo, T Brown, P Tien, A Adimora

Conflicts of Interest:

TTB has served a consultant for Merck, Gilead, Janssen, ViiV Healthcare, and Theratechnologies. FJP has served as a consultant and/or speaker for Merck, Gilead, Janssen, ViiV Healthcare, and Theratechnologies. All other authors had no conflicts

¹⁰Department of Medicine, UNC School Division of Medicine, The Infectious Diseases, University of North Carolina School of Medicine, University of North Carolina at Chapel Hill, Chapel Hill, NC, USA

¹¹Department of Medicine, Division of Infectious Diseases, Georgetown University, Washington DC, USA

¹²Department of Infectious Diseases and Microbiology, Department of Pathology, University of Pittsburgh, Pittsburgh, PA, USA

¹³Department of Medicine, Johns Hopkins University, Baltimore, MD, USA

¹⁴Department of Medicine, University of California, San Francisco and Department of Veterans Affairs, San Francisco, CA, USA

¹⁵Department of Epidemiology, UNC Gillings School of Global Public Health, University of North Carolina at Chapel Hill, Chapel Hill, NC, USA

Abstract

Background: SARS-CoV-2 infection and COVID-19 symptoms among people living with HIV (PLWH) are not well described.

Setting: Longitudinal survey within the MACS/WIHS Combined Cohort Study (MWCCS) of PLWH compared to similar HIV seronegative (SN) individuals.

Methods: Telephone-administered survey of MWCCS participants at 13 clinical research sites across the U.S. addressing COVID-19 symptoms, SARS-CoV-2 testing, and pandemic impact on social distancing and antiretroviral therapy (ART) use. Primary data collection occurred during May (wave 1), June-July (wave 2), and August-September, 2020 (wave 3).

Results: One-third of MWCCS participants were tested for SARS-CoV-2 infection; 10% were tested 2 times. Similar proportions of PLWH and SN participants were tested, but SARS-CoV-2 positivity was higher among PLWH than SN (9.4% vs 4.8%, $p=0.003$). Odds of SARS-CoV-2 positivity remained higher among PLWH after adjusting for age, sex, race/ethnicity, and study site (aOR=2.0, 95% CI=1.2-3.2). SARS-CoV-2 positivity was not associated with CD4 cell counts among PLWH. Among SARS-CoV-2 positive participants, 9% had no symptoms, 7% had 1–2 mild symptoms, and 84% had 3 symptoms. Most (98%) participants reported physical distancing during all survey waves; self-reported ART adherence among PLWH was not adversely impacted during the pandemic compared to the prior year (similar adherence in 89% of participants, improved in 9%, decreased in 2%).

Conclusions: Despite similar SARS-CoV-2 testing and physical distancing profiles by HIV serostatus among MWCCS participants, PLWH who reported SARS-CoV-2 testing were more likely to have a positive test. Additional studies are needed to determine whether and why PLWH are at increased risk for SARS-CoV-2 infection.

Keywords

coronavirus; testing; symptoms; PLWH; CD4; distancing; MWCCS

Introduction

Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), the virus responsible for coronavirus disease 2019 (COVID-19), manifests itself with a variety of clinical symptoms during infection. While most people who become infected are either asymptomatic or experience mild symptoms, the illness can be severe and/or life-threatening¹. Between March and September 2020, with infection levels surging across the U.S.², over 7 million people tested positive for SARS-CoV-2 and 200,000 people died^{3,4}. In an effort to mitigate the impact of the pandemic, many states mandated policies of social distancing and closures of businesses and other venues⁵.

COVID-19 severity increases among adults with each decade of advancing age⁶, and among those with certain underlying health conditions and comorbidities. It is unclear whether SARS-CoV-2 infection is higher among people living with HIV (PLWH), however some recent studies suggest PLWH may have increased disease severity and risk of hospitalization.⁷ A recent review of SARS-CoV-2 cases among PLWH indicated that although documented combined HIV and SARS-CoV-2 co-infection was uncommon globally, greater HIV-related immunosuppression predisposed PLWH to more severe COVID-19 disease.^{8,9} Analysis of 192 PLWH suggested that a CD4 count <200 cells/ μ L [vs 200 cells/ μ L] was associated with a 4.9-fold higher odds of progression to severe COVID-19.⁸ Currently, recommendations for prevention of SARS-CoV-2 infection and COVID-19 management among PLWH are no different than those for the general population. The U.S. Centers for Disease Control and Prevention (CDC) recommend that “until more is known, additional caution for all PLWH, especially those with advanced or poorly controlled HIV, is warranted” and that PLWH should follow CDC prevention recommendations.¹⁰ However, PLWH are more likely than the general population to have risk factors associated with greater SARS-CoV-2 exposure and/or worse COVID-19 outcomes, including higher prevalence of adverse social determinants of health (e.g., unstable housing, public transportation use) and comorbidities (e.g., cardiovascular disease, obesity, smoking).¹¹ PLWH in the U.S. are also disproportionately represented among racial minorities and groups hit hardest by the SARS-CoV-2 pandemic.

We previously reported similar prevalences and types of COVID-19 symptoms among PLWH and similar HIV-seronegative (SN) adults in the MACS/WIHS Combined Cohort Study (MWCCS) during Spring 2020.^{12(p1)} In the current analysis, we characterize the evolving pandemic among MWCCS participants by extending our initial report to include additional waves of data collection through September 2020, after the surge of COVID-19 that occurred in the U.S. in the Spring and Summer of 2020.

Methods

The MWCCS integrates two long-standing, prospective, multicenter U.S. cohorts of PLWH and SN people, the Multicenter AIDS Cohort Study (MACS) and the Women's Interagency HIV Study (WIHS). Clinical research sites are located in New York City (Brooklyn, Bronx), Mid-Atlantic (Baltimore, MD; Washington DC;), Southeast (Chapel Hill, NC; Atlanta, GA; Miami, FL), South (Birmingham, AL; Jackson MS), Midwest (Chicago, IL; Columbus, OH;

Pittsburgh, PA), and California (San Francisco, Los Angeles). Characteristics of participants are described in detail elsewhere.¹³ The MACS targeted enrollment of men who have sex with men (MSM) with some censoring of seronegative participants in 1995 to achieve an approximate 1:1 target ratio of PLWH to seronegative men for follow-up, and the WIHS enrollment targeted women living with HIV and similar seronegative women at a 3:1 ratio.

Three waves of an interviewer-administered telephone survey were conducted between April 8th and September 30th 2020, with the majority of surveys occurring in May (wave 1), June and July (wave 2), and August and September (wave 3). A subset of MWCCS sites conducted an optional 4th administration of the survey involving 70 participants in September 2020. These data were combined with wave 3 data for these analysis to report ever positive among those tested multiple times. Among those reporting symptoms, the severest symptoms reported across waves were reported. The survey was offered in English and Spanish. All MWCCS participants were eligible. Verbal consent was obtained prior to the telephone interview which was conducted by trained study staff. Participants were financially compensated for their time. The COVID-19 survey was reviewed and approved by each MACS and WIHS clinical research site's local Institutional Review Board.

The COVID-19 survey was developed by MWCCS investigators to capture self-reported information regarding COVID-19 symptoms, SARS-CoV-2 testing, preventive measures, and psychosocial effects of the pandemic (the latter are not included in this report). The survey was minimally revised after the first wave. Both versions of the survey are in the public domain and available at <https://statepi.jhsph.edu/mwccs/data-collection-forms/> (V100 COVID forms). Interviewers asked participants questions regarding COVID-19 symptoms, SARS-CoV-2 testing, antiretroviral therapy (ART) adherence, and adoption of social distancing measures. Questions addressing COVID-19 symptom severity (mild, moderate, or severe for each self-reported symptom) and cohabitation with someone who tested positive for SARS-CoV-2, were added after the first wave of the survey.¹²

This analysis includes three waves of COVID-19 symptom and self-reported SARS-CoV-2 testing data. SARS-CoV-2 testing included any self-reported viral diagnostic test via nasopharyngeal, nasal, or pharyngeal swabs, or saliva (serum antibody tests were excluded from this analysis). Information regarding the type of SARS-CoV-2 testing was not collected in wave 1 (when antibody testing was not available; all tests in that wave were assumed to be for diagnosis of active infection). Type of test was collected during subsequent waves of survey administration when SARS-CoV-2 antibody, antigen, and nucleic amplification tests became more widespread. SARS-CoV-2 positivity was defined as a self-reported positive viral diagnostic test. We attempted to confirm all self-reported SARS-CoV-2 testing results (positive and negative) with medical record review. Of 95 people with self-reported SARS-CoV-2 positive tests, 54 (57%) were confirmed by medical record abstraction, 41 (43%) were pending (did not have any record available or were still awaiting requested records at time of analysis), and none (0%) were ascertained to be negative after record review. There were 30 people (12 PLWH; 18 SN) who self-reported having been tested for SARS-CoV-2 but had unknown (pending) results at the time of analysis and were excluded from analysis.

Among PLWH, plasma CD4 and HIV RNA (viral load) and self-reported ART adherence data from the most recent in-person MACS or WIHS study visit, a median of 11.9 months (IQR=9.7–51.0) prior to the survey, were used.

Statistical Methods

Baseline MWCCS participant characteristics were obtained from the last in-person MACS and WIHS visit (2018–2019), stratified by HIV serostatus and sex. SARS-CoV-2 test positivity was explored overall, by HIV serostatus, CD4 count (≥ 500 vs <500 cells/ μL , and <350 vs 350–499 cells/ μL), calendar month of testing, and by sex.

COVID-19 symptom prevalence and severity were explored among participants reporting SARS-CoV-2 positive infection status. Among PLWH, ART adherence was compared among persons reporting SARS-CoV-2 positive testing vs SARS-CoV-2 negative testing vs no report of being tested. ART adherence reported during the pandemic was also compared to ART adherence reported the prior year (last WIHS/MACS visit). Social distancing measures self-reported at the time of the surveys were described. Differences among categorical variables were tested using Chi-square analyses (or Fisher's exact test when values in subgroups were <5). Significance was evaluated using 2-tailed tests, at $p < 0.05$. We decided *a priori* to adjust analyses for age, sex, race, and study site given prior published literature establishing these as potentially associated with SARS-CoV-2 infection. SAS version 9.4 was used for all analyses.

Results

Of 4,330 MACS and WIHS participants, 3,671 (85%) completed at least one survey, 500 (11%) could not be reached for the telephone interview, and 159 (4%) refused (Supplemental Table 1). Among the 3671 who completed at least one survey, the majority (89%) completed three surveys. For each survey wave, the range of dates and number of participants included are further described in Supplemental Table 1.

Participants who completed 1 survey were 53% female, 41% Black, 14% Hispanic, and 35% White (Table 1). Participants' median age was 58 years (IQR 50–65 years). Most PLWH reported taking ART (97%) and had an undetectable HIV viral load (73%) with a median CD4 cell count of 670 cells/ μL (IQR 468–907 cells/ μL) at their last study visit.

SARS-CoV-2 Testing

Characteristics of participants who reported having tested for SARS-CoV-2 infection were similar to those who did not (Table 1). Among those who reported testing, characteristics of PLWH and SN participants were similar (Table 1). Over one-third of MWCCS participants ($N=1234$, 34%) were tested for SARS-CoV-2 infection at 1 time during the study period (April 1st to September 31st, 2020), including 391 (11%) participants who were tested 2 times. A similar proportion of PLWH and SN participants were tested for SARS-CoV-2 (34.3% vs 33.5%, respectively, Table 2).

Most SARS-CoV-2-tested participants reported undergoing nasopharyngeal swab (84%) or oropharyngeal swab (15%), with a lower proportion reporting saliva (0.8%) or "other

type''(0.2%) The proportions of both PLWH (6.7%) and SN (4.5%) who reported living with someone who tested positive for SARS-CoV-2 were <10% (Table 1); 100 participants reported living with someone who tested positive for SARS-CoV-2, and of these 71% (71/100) reported getting tested and 41% (29/71) of those reported testing positive.

SARS-CoV-2 Positivity, HIV serostatus and CD4 cell count

The percent SARS-CoV-2 positive among tested persons was higher in wave 1 (9.3%) than waves 2 (4.4%) or 3 (4.5%), $p=0.001$. Among 1,248 MWCCS participants who were tested, SARS-CoV-2 positivity was higher among PLWH than SN (9.4% vs 4.8%, $p=0.003$) overall and at each visit wave: wave 1 (11.1% vs 6.7%, $p=0.13$); wave 2 (5.4% vs 3.1%, $p=0.21$); and wave 3 (6.1% vs 2.3%, $p=0.02$). The proportion of positive tests was also higher among PLWH, both men and women, than SN: among 658 women tested (8.6% vs 5.9%, $p=0.24$) and among 590 men tested (10.5% vs 4.0%, $p=0.003$).

Among PLWH, the proportion of positive SARS-CoV-2 tests was similar among those with CD4 count <500 cells/ μ L vs those with CD4 count \geq 500 cells/ μ L (9.8% vs 9.2%, $p=0.79$; Figure 1). The proportion of positive SARS-CoV-2 tests was not statistically different in PLWH with CD4 counts 350–499 vs <350 cells/ μ L (11.4% vs 8.3%, $p=0.43$) or vs 92 participants with CD4<200 (7.5%, $p= 0.47$). Only 109 participants (43 men, 66 women) with CD4 counts <350 cells/ μ L were tested for SARS-CoV-2, of whom 11.6% men and 6.1% women were SARS-CoV-2 positive (p for sex difference= 0.31). The proportion of positive SARS-CoV-2 tests was also similar by HIV viral load (among 21 PLWH with detectable vs 48 with undetectable: 2.8% vs 6.4%, $p=0.54$).

As shown in Figure 2, across study sites there was heterogeneity in the proportions of people with positive SARS-CoV-2 tests and differences in proportions of PLWH versus SN who had positive tests (Figure 2). The prevalence of SARS-CoV-2 positivity was higher among PLWH than SN participants in 8 of the 13 study sites (both California sites, all Midwestern sites, and the Brooklyn site), was modestly higher among PLWH than SN participants in 3 study sites (both Mid-atlantic and Atlanta sites) and was lower in PLWH than SN participants at 2 sites (Bronx and the Mississippi/Alabama site, which each reported <100 people tested).

After adjusting for age, sex, race, and study site, odds of SARS-CoV-2 test positivity among those tested remained higher among PLWH than SN participants (aOR=2.0, 95% CI=1.2–3.2).

COVID-19 symptoms

As shown in Table 3, most (N=80/95; 84%) SARS-CoV-2 positive participants reported >3 symptoms and more than one-third (35%) of SARS-CoV-2 positive participants reported >3 symptoms that were severe. Symptom severity was similar by HIV serostatus. Symptom severity increased with number of symptoms reported (Table 3). The most common symptoms reported as moderate or severe were myalgias (50%), headache (49%), loss of taste or smell (47%), cough (41%), chills (39%), shortness of breath (37%), and feeling feverish (33%); table 3. However, no single symptom was reported by all SARS-CoV-2

positive participants and many of the individual symptoms were reported as none or mild by the majority of participants (Table 3).

Among PLWH who were SARS-CoV-2 positive, lower CD4 cell count was not associated with higher symptom severity. There was also no difference by CD4 cell count category (<350, 350–499, and ≥500 cells/μL) in the proportion of PLWH with ≥1 severe symptom (56% vs 58% vs 51%, respectively, $p=0.89$) and ≥3 severe symptoms (33% vs 42% vs 31%, respectively, $p=0.79$).

Pandemic impact

Self-reported social distancing practices were explored. Social distancing was consistently reported by 98% of participants across all surveys, while staying home “as much as possible” decreased modestly from 97% (Spring; wave 1) to 93% (Fall; wave 3), $p<0.001$. However, 13% of all participants in each wave reported not making any changes to their daily life and routine. In wave 1, 19% of PLWH and 13% of SN participants reported not making any changes. These findings did not change much over time, with 11–17% of PLWH and 7–14% of SN participants reporting not making changes in subsequent waves.

Participants reported not only social distancing, but also staying at home all the time (self-isolating) due to “experiencing symptoms or having a positive test” (1–3%), exposure to an infected person (1–2%), or uncertainty about infection status (2–5%, depending on wave). The prevalence of staying at home was similar by HIV serostatus (Supplemental Figure 1). When our survey was first administered in April 2020, 10% of participants reported self-isolating; this proportion decreased over time, and was 7% in May, 4% in June and July, and 3% in August and September (Supplemental Figure 1).

We explored use of ART among PLWH during the pandemic (Supplemental Table 2). The majority of participants (95%) reported taking ART ≥95% of the time, with most reporting an excellent (53%) or very good (31%) job of taking ART ‘the way they were supposed to’. Participants who reported being SARS-CoV-2 positive reported a lower prevalence of 100% ART adherence than those who tested negative (66% vs 76% $p=0.02$), and more often missed a dose of ART on ≥1 days per month (52% vs. 37%, $p=0.02$). However, the prevalence of being off ART for ≥1 week was similar by SARS-CoV-2 test result status (10% vs 11%, $p=0.088$, Supplemental Table 2). To further evaluate how pandemic-related disruptions may have changed ART adherence, we compared ART adherence reported in this MWCCS COVID survey to that reported when last measured as part of routine MACS and WIHS cohort data collection (approximately one year prior to the COVID survey). ART adherence reported in the COVID survey was similar to previously reported adherence in 89% of participants, improved in 9% of participants, and decreased in 2% of participants (none of whom were known to be SARS-CoV-2 positive) (Supplemental Table 2).

Discussion

Despite similar SARS-CoV-2 testing rates, a higher proportion of PLWH in our study tested positive for SARS-CoV-2 infection compared to SN participants. This suggests that PLWH may have increased susceptibility or have had greater non-HIV related risks for

SARS-CoV-2 infection. Our findings are similar to a large study of more than 280,000 people tested by the San Francisco Department of Public Health, which reported higher SARS-CoV-2 test positivity among PLWH than SN (4.5% vs 3.5%)¹⁴. However our findings differ from several other studies which reported similar risk of infection with SARS-CoV-2 in PLWH and SN.¹⁵⁻¹⁷.

The prevalence of self-reported COVID-19 symptoms and symptom severity was similar by HIV serostatus in the MWCCS. The proportion of SARS-CoV-2 positive participants who reported symptoms in our study was high (92%). However most of the months that our surveys were conducted, SARS-CoV-2 testing was of limited availability and sought primarily by persons who were symptomatic or had a known exposure. Reasons for SARS-CoV-2 testing were not collected in this study so could not be evaluated. COVID-19 symptoms did not differ by HIV serostatus, although some previous studies reported that immunosuppression in PLWH was associated with more severe COVID-19.⁸ Several large studies have reported higher rates of COVID-19 hospitalization and mortality among PLWH than among SN cases,^{7,18} raising concern that PLWH may be at increased risk of severe outcomes from COVID-19.

The impact of the pandemic on MWCCS participants was dramatic, prompting almost ubiquitous physical distancing and high reported self-isolation. Impacts on ART adherence were more modest, with disruptions in use reported by 9% of participants; however these appeared short-lived. Only 2% of PLWH reported lower ART adherence during the pandemic compared to pre-pandemic, and 9% reported higher ART adherence during the pandemic. This is consistent with studies reporting some disruptions in HIV care but care engagement not being seriously disrupted.^{19,20}, and adherence increasing in some PLWH during the pandemic.²¹

Our study had several strengths. First, the MWCCS includes study sites across the U.S. and collected data on PLWH and similar HIV SN individuals for comparison. In addition, the same survey was administered across all study sites by well-trained staff who knew the participants well. Third, a high proportion of participants completed the survey. Finally, the survey was administered multiple times over a 5-month timeframe and thus was able to capture updated, real-time, and changing information about social distancing practices, self-reported COVID-19 symptoms and SAR-CoV-2 testing. Our study also has some limitations. First, we relied on participants' self-report of COVID testing and test results, as medical records were not readily available for all cases. However, people generally are likely to recall experiencing a nasopharyngeal or oropharyngeal swab and during the early months of the COVID-19 pandemic these specimens were unlikely to have been obtained for reasons other than COVID-19 testing. Medical records were obtained for more than half of the self-reported SARS-CoV-2 positives. Among self-reported cases for whom test data were obtained, all cases were confirmed based on test results. MWCCS participants are used to reporting medical results and the study has a system in place for medical information collection, however the array of locations/testing sites and non-centralized testing methods/locations rolled out to provide access to SARS-CoV-2 testing, increased the challenges associated with this process. Furthermore, some rapid SARS-CoV-2 testing sites did not provide written results and/or did not have centralized documentation of test results that

could be requested, limiting confirmation. Second, we lacked information concerning access to and reasons for testing. Systematic differences in the distribution of reasons for testing could have affected the proportions of positive SARS-CoV-2 tests observed among PLWH and SN participants. We anticipate that most SARS-CoV-2 tests were prompted by the presence of COVID-19 symptoms, as there were few self-reported asymptomatic COVID-19 cases, however, asymptomatic cases are underreported everywhere. Another limitation was that we did not have confirmed information on COVID-related hospitalizations and deaths, though collection of those medical records is planned in the future. Additionally, as most participants are on effective therapies we had a limited number of severely immunosuppressed subjects so were not well powered to explore infection among those with very low CD4 cell counts.

In our large, prospective, multi-center U.S. observational cohort of people with and without HIV infection, one-third of participants reported SARS-CoV-2 testing between April to September 2020, and the proportion of self-reported SARS-CoV-2 positive tests was twice as high among PLWH than SN participants. Higher SARS-CoV-2 positivity among PLWH was observed in each of the three administration cycles of the MWCCS COVID-19 survey and remained after adjusting for age, sex, race, and clinical research site. COVID-19 symptoms type, prevalence, and severity were similar by HIV serostatus. These data suggest that PLWH may be at increased risk of acquisition SARS-CoV-2 infection compared to SN persons, however since not all participants were tested, further research is needed to support this finding.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgement and Sources of Support:

The contents of this publication are solely the responsibility of the authors and do not represent the official views of the National Institutes of Health (NIH). MWCCS (Principal Investigators): Atlanta CRS (Ighovwerha Ofotokun, Anandi Sheth, and Gina Wingood), U01-HL146241; Baltimore CRS (Todd Brown and Joseph Margolick), U01-HL146201; Bronx CRS (Kathryn Anastos and Anjali Sharma), U01-HL146204; Brooklyn CRS (Deborah Gustafson and Tracey Wilson), U01-HL146202; Data Analysis and Coordination Center (Gypsyamber D'Souza, Stephen Gange and Elizabeth Golub), U01-HL146193; Chicago-Cook County CRS (Mardge Cohen and Audrey French), U01-HL146245; Chicago-Northwestern CRS (Steven Wolinsky), U01-HL146240; Northern California CRS (Bradley Aouizerat, Jennifer Price, and Phyllis Tien), U01-HL146242; Los Angeles CRS (Roger Detels and Matthew Mimiaga), U01-HL146333; Metropolitan Washington CRS (Seble Kassaye and Daniel Merenstein), U01-HL146205; Miami CRS (Maria Alcaide, Margaret Fischl, and Deborah Jones), U01-HL146203; Pittsburgh CRS (Jeremy Martinson and Charles Rinaldo), U01-HL146208; UAB-MS CRS (Mirjam-Colette Kempf, Jodie Dionne-Odom, and Deborah Konkle-Parker), U01-HL146192; UNC CRS (Adaora Adimora), U01-HL146194. The MWCCS is funded primarily by the National Heart, Lung, and Blood Institute (NHLBI), with additional co-funding from the *Eunice Kennedy Shriver* National Institute Of Child Health & Human Development (NICHD), National Institute On Aging (NIA), National Institute Of Dental & Craniofacial Research (NIDCR), National Institute Of Allergy And Infectious Diseases (NIAID), National Institute Of Neurological Disorders And Stroke (NINDS), National Institute Of Mental Health (NIMH), National Institute On Drug Abuse (NIDA), National Institute Of Nursing Research (NINR), National Cancer Institute (NCI), National Institute on Alcohol Abuse and Alcoholism (NIAAA), National Institute on Deafness and Other Communication Disorders (NIDCD), National Institute of Diabetes and Digestive and Kidney Diseases (NIDDK), National Institute on Minority Health and Health Disparities (NIMHD), and in coordination and alignment with the research priorities of the National Institutes of Health, Office of AIDS Research (OAR). MWCCS data collection is also supported by U11-TR000004 (UCSF CTSA), U11-TR003098 (JHU ICTR), U11-TR001881 (UCLA CTSI), P30-AI-050409 (Atlanta CFAR), P30-AI-073961 (Miami CFAR), P30-AI-050410 (UNC CFAR), P30-AI-027767 (UAB CFAR), and P30-MH-116867 (Miami CHARM). Support also from K23 AI124913 (Cecile Lahiri)

The authors gratefully acknowledge the contributions of the study participants and dedication of the staff at the MWCCS sites.

References

1. Gallo Marin B, Aghagoli G, Lavine K, et al. Predictors of COVID-19 severity: A literature review. *Rev Med Virol.* 2021;31(1):1–10. doi:10.1002/rmv.2146
2. COVID-19 United States Cases by County. Johns Hopkins Coronavirus Resource Center. Accessed January 22, 2021. <https://coronavirus.jhu.edu/us-map>
3. CDC. COVIDView, Key Updates for Week 42. Centers for Disease Control and Prevention. Published 10 23, 2020. Accessed January 22, 2021. <https://www.cdc.gov/coronavirus/2019-ncov/covid-data/covidview/past-reports/10232020.html>
4. Track Testing Trends. Johns Hopkins Coronavirus Resource Center. Accessed January 22, 2021. <https://coronavirus.jhu.edu/testing/tracker>
5. Impact of Opening and Closing Decisions in California, New Cases - Johns Hopkins. Johns Hopkins Coronavirus Resource Center. Accessed January 22, 2021. <https://coronavirus.jhu.edu/data/state-timeline>
6. CDC. COVID-19 and Your Health. Centers for Disease Control and Prevention. Published 2 11, 2020. Accessed February 25, 2021. <https://www.cdc.gov/coronavirus/2019-ncov/need-extra-precautions/older-adults.html>
7. Tesoriero JM, Swain C-AE, Pierce JL, et al. COVID-19 Outcomes Among Persons Living With or Without Diagnosed HIV Infection in New York State. *JAMA Netw Open.* 2021;4(2):e2037069. doi:10.1001/jamanetworkopen.2020.37069 [PubMed: 33533933]
8. Kanwugu ON, Adadi P. HIV/SARS-CoV-2 coinfection: A global perspective. *J Med Virol.* 2021;93(2):726–732. doi:10.1002/jmv.26321 [PubMed: 32692406]
9. Sun J, National COVID Cohort Collaborative. CROI 2021 Oral: COVID-19 HOSPITALIZATION AMONG PEOPLE WITH HIV OR SOLID ORGAN TRANSPLANT IN THE US. In: ; 2021. Accessed March 11, 2021. <https://ww2.aievolution.com/cro2101/index.cfm?do=abs.viewAbs&abs=1882>
10. Interim Guidance for COVID-19 and Persons with HIV COVID-19 and Persons with HIV (Interim Guidance). AIDSinfo. Accessed June 8, 2020. <https://aidsinfo-nih.gov.proxy1.library.jhu.edu/guidelines/html/8/covid-19-and-persons-with-hiv--interim-guidance-/554/interim-guidance-for-covid-19-and-persons-with-hiv>
11. HIV: COVID-19 Real Time Learning Network Summary by CDC and IDSA. Accessed January 22, 2021. <https://www.idsociety.org/covid-19-real-time-learning-network/special-populations/hiv/>
12. D'Souza G, Springer G, Gustafson D, et al. COVID-19 symptoms and SARS-CoV-2 infection among people living with HIV in the US: the MACS/WIHS combined cohort study. *HIV Res Clin Pract.* 2020;21(5):130–139. doi:10.1080/25787489.2020.1844521 [PubMed: 33211636]
13. D'Souza G, Bhondoekhan F, Benning L, et al. Characteristics of the MACS/WIHS Combined Cohort Study: Opportunities for Research on Aging With HIV in the Longest US Observational Study of HIV. *American Journal of Epidemiology.* 2021;190(8):1457–1475. doi:10.1093/aje/kwab050 [PubMed: 33675224]
14. Sachdev D, Mara E, Hsu L, et al. COVID-19 Susceptibility and Outcomes Among People Living With HIV in San Francisco. *J Acquir Immune Defic Syndr.* 2021;86(1):19–21. doi:10.1097/QAI.0000000000002531 [PubMed: 33044323]
15. Saag M Special section: COVID-19 among people living with HIV. *AIDS.* 2020;34(12):1755–1756. doi:10.1097/QAD.0000000000002648 [PubMed: 32889850]
16. Charre C, Icard V, Pradat P, et al. Coronavirus disease 2019 attack rate in HIV-infected patients and in preexposure prophylaxis users. *AIDS.* 2020;34(12):1765–1770. doi:10.1097/QAD.0000000000002639 [PubMed: 32889852]
17. Park L, Rentsch C, Sigel K, Rodriguez-Barradas M. COVID-19 in the largest US HIV cohort. In: ; 2020:Late-breaking poster LBPE023. <https://cattendee.abstractsonline.com/meeting/9289/presentation/3924>

18. Bhaskaran K, Rentsch CT, MacKenna B, et al. HIV infection and COVID-19 death: a population-based cohort analysis of UK primary care data and linked national death registrations within the OpenSAFELY platform. *The Lancet HIV*. 2021;8(1):e24–e32. doi:10.1016/S2352-3018(20)30305-2 [PubMed: 33316211]
19. Gwadz M, Campos S, Freeman R, et al. Black and Latino Persons Living with HIV Evidence Risk and Resilience in the Context of COVID-19: A Mixed-Methods Study of the Early Phase of the Pandemic. *AIDS Behav*. Published online 2 10, 2021. doi:10.1007/s10461-021-03177-0
20. Ballivian J, Alcaide ML, Cecchini D, Jones DL, Abbamonte JM, Cassetti I. Impact of COVID-19-Related Stress and Lockdown on Mental Health Among People Living With HIV in Argentina. *J Acquir Immune Defic Syndr*. 2020;85(4):475–482. doi:10.1097/QAI.0000000000002493 [PubMed: 33136748]
21. Kalichman SC, Eaton LA, Berman M, et al. Intersecting Pandemics: Impact of SARS-CoV-2 (COVID-19) Protective Behaviors on People Living With HIV, Atlanta, Georgia. *JAIDS Journal of Acquired Immune Deficiency Syndromes*. 2020;85(1):66–72. doi:10.1097/QAI.0000000000002414 [PubMed: 32530862]

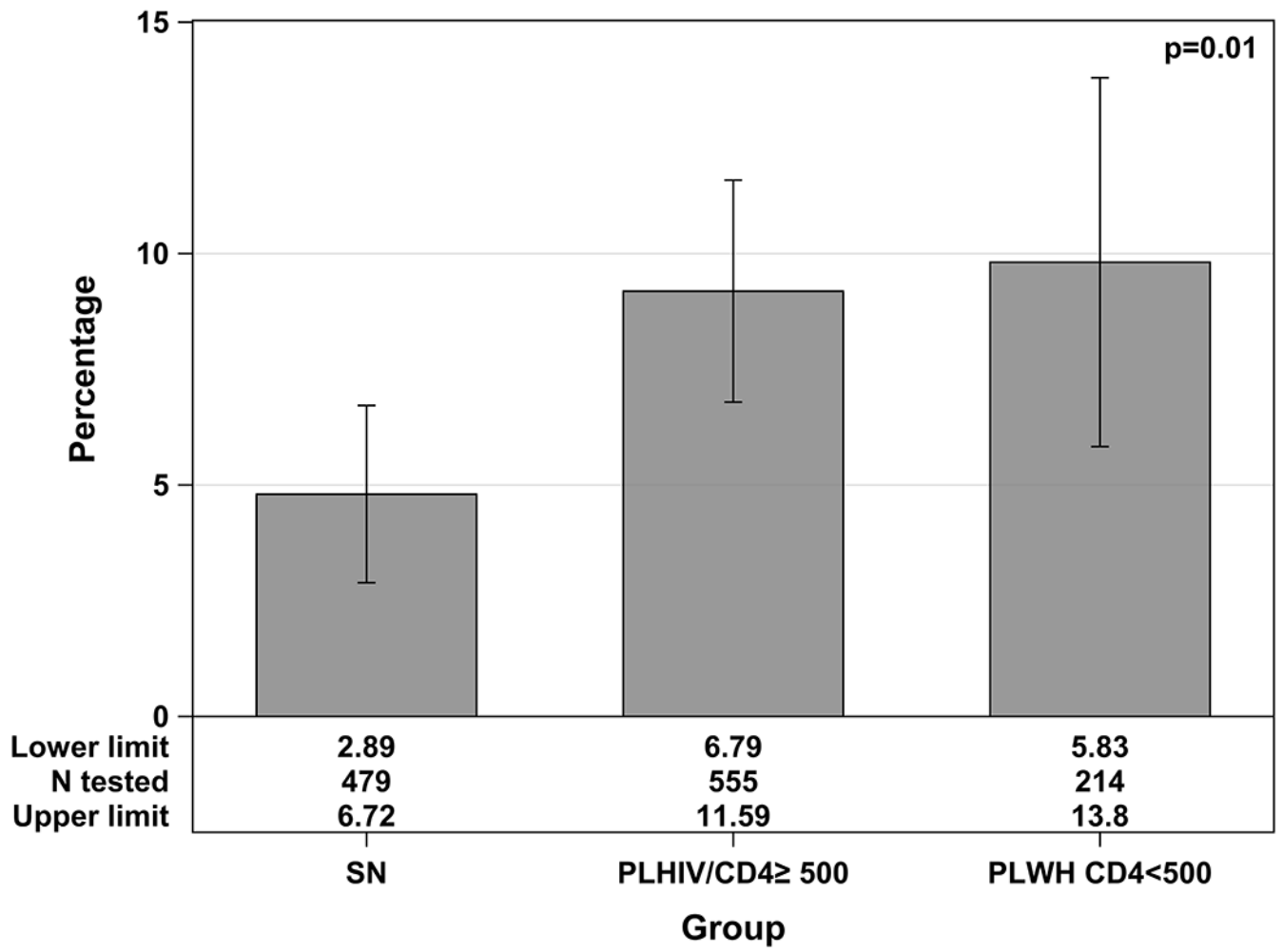


Figure 1. SARS-CoV-2 test positivity by HIV serostatus and most recent CD4 cell count, among persons tested for SARS-CoV-2.

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript

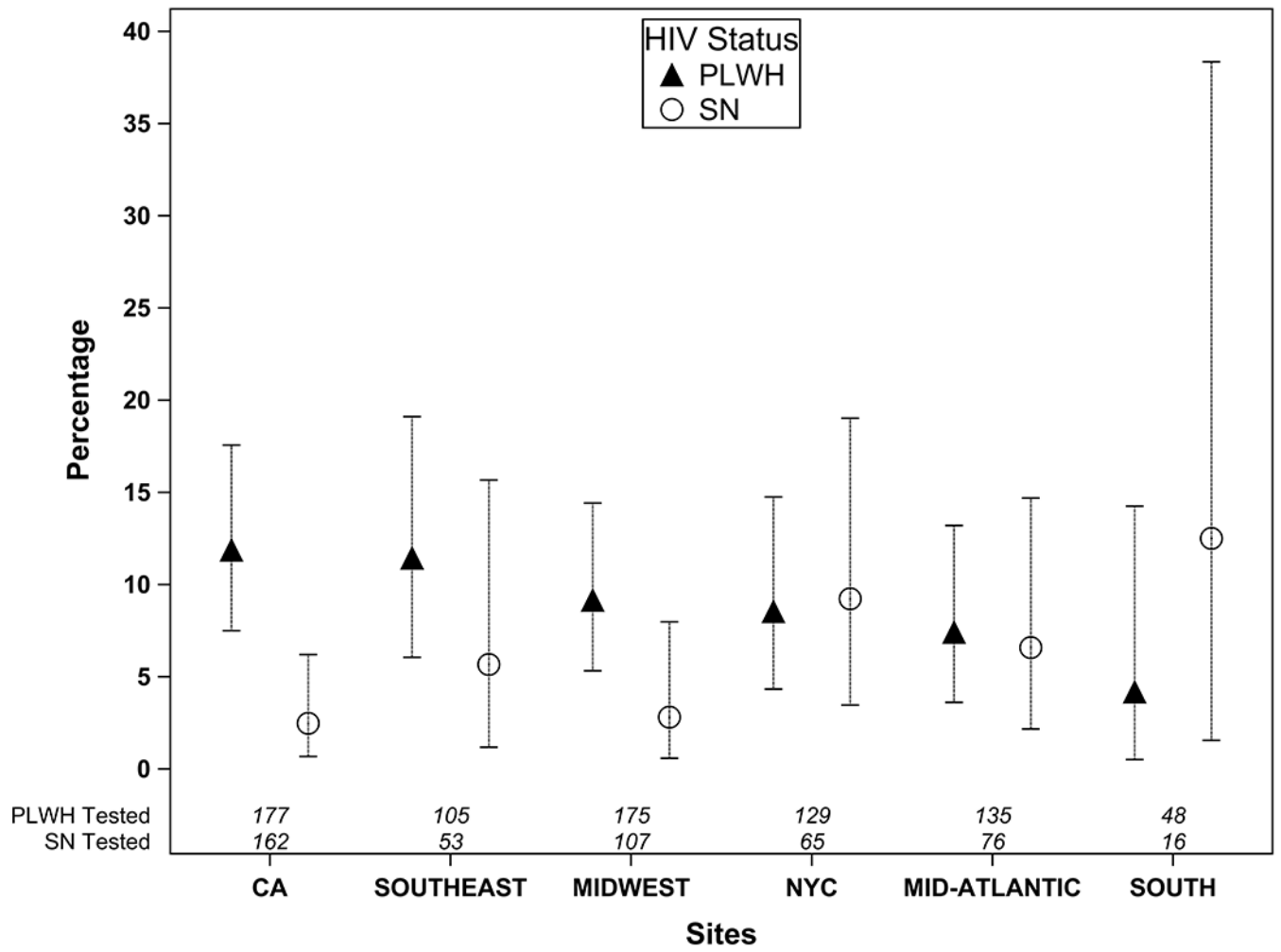


Figure 2. Percent of positive SARS-CoV-2 tests among persons tested by study site region and HIV serostatus

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript

Table 1. Characteristics of MWCCS participants completing the MWCCS COVID-19 survey, by SARS-CoV-2 testing and HIV serostatus.

Characteristics	No.		%		Column %						P value tested vs not
	All	N=3671	All	N=3671	All Tested	PLWH	SN	All Not Tested	PLWH	SN	
					N=1248	N=769	N=479	N=2423	N=1472	N=951	
Sex											0.95
Female	1938		53		53	59	42	53	62	39	
Male	1733		47		47	41	58	47	38	61	
Race and Ethnicity											0.34
Black, non-Hispanic	1516		41		42	47	34	41	46	32	
Hispanic, any race	496		14		15	16	13	13	16	9	
White, non-Hispanic	1299		35		34	27	44	36	28	50	
Other, non-Hispanic	360		10		10	11	9	10	10	9	
Region of US											0.004
California	873		24		27	23	34	22	20	25	
Mid-Atlantic (Wash DC, MD)	648		18		17	18	16	18	16	21	
Midwest (IL, PA, OH)	932		25		23	23	22	27	24	31	
Northeast (NY) ^{&}	539		15		16	17	14	14	16	11	
South (AL, MS) ^{&}	186		5		5	6	3	5	7	3	
Southeast (NC, GA, FL) ^{&}	493		13		13	14	11	14	17	10	
Number of people who live with you [~]											0.53
0	1241		36		37	37	37	36	37	34	
1-2	1896		56		54	54	55	56	55	58	
3	276		8		9	9	8	8	8	7	
Anyone you live with tested positive for coronavirus?	100		3		6	7	5	1	1	1	<.0001
Currently use tobacco? (smoke or vape)	829		24		23	25	21	25	27	22	0.42
Anyone you live with smoke tobacco? [^]	533		16		16	16	15	16	17	14	0.97

Characteristics	No.		%		Column %				P value tested vs not		
	All	N=3671	All	N=3671	All Tested	PLWH	SN	All Not Tested		PLWH	SN
					N=1248	N=769	N=479	N=2423	N=1472	N=951	
Age in years: median (IQR)	3671	58 (50, 65)	97	57 (50, 65)	56 (49, 63)	60 (51, 67)	58 (50, 65)	56 (49, 62)	61 (51, 68)	0.46	
Currently taking antiretroviral medications*	2176	97	NA	98	97	NA	NA	97	NA	0.16	
Current CD4 cells/ μ L [†] :	2241	NA	NA	NA	NA	NA	NA	NA	NA	NA	
median (IQR)	670 (468, 907)	677 (477, 890)	667 (464, 913)	71%	72%	28%	71%	29%	0.65		
500	1602	71%	29%	28%	29%	29%	29%	29%	0.60		
<500	639	29%	29%	28%	28%	29%	29%	29%	0.71		
Current HIV RNA copies/ μ L [‡] ; median (IQR)	2200	Und (umd, 24)	NA	Und (umd, 25)	Und (umd, 24)	NA	NA	Und (umd, 24)	NA	0.71	

PLWH = People Living with HIV; SN= HIV Seronegative; Und= undetectable HIV viral load (<20 copies/ μ L)

[‡] “New York City”, “Southeast” and the “South” study regions only included women participants. Note: men in this paper were part of the former MACS cohort and women were part of the former WIHS cohort, now merged into the MACS/WIHS Combined Cohort Study (MWCSS).

* Current CD4 cell count, current HIV RNA and antiretroviral medication history data are from participants' last in-person visit, median of 11.9 months prior to when this survey was collected

[†] Analysis was among 3413 people as there were 258 people missing data (ie skipped or refused to answer) for number of people lived with

[‡] Analysis was among 3404 people as there were 267 people missing data (ie skipped or refused to answer) on tobacco use in those they lived with Chi-square statistics was used for categorical variables and Wilcoxon rank sum test was used for continuous variables, to test the variable differences between tested and not tested participants.

Table 2.

Cumulative prevalence of SARS-CoV-2 infection (April-September 2020), by HIV serostatus among MWCCS participants

Overall (Any Wave 1–3)							By Survey Wave [*]		
	Cum N	Cum Prev	PLWH	SN	P-value	1 (Spring 2020)	2 (Summer 2020)	3 [#] (Fall 2020)	
	N=3671		N=2241	N=1430		N=3415	N=3387	N=3325	
SARS-CoV-2 diagnostic Test [~]									
Tested 1 time	1248	34.0%	34.3%	33.5%	0.61	13.0%	15.6%	20.2%	
Tested 2 times	391	10.7%	11.1%	10.0%	0.31	0%	4.3%	7.3%	
SARS-CoV-2 diagnostic test result					0.003				
Positive [^]	95	7.6%	9.4%	4.8%		9.5%	4.6%	4.6%	
Negative	1153	92.4%	90.6%	95.2%		90.5%	95.4%	95.4%	

PLWH = People Living with HIV; SN= HIV Seronegative; Cum=cumulative

^{*} Results in each wave for PLWH and SN were similar and thus were combined in the table above for the wave-specific result presentation. For those interested, the proportion of PLWH and SN tested 1 time was: 13.5% vs 12.3% at wave 1 ; 16.1% vs 14.7% at wave 2; 20.5% vs 19.7% at wave 3. The proportion of tested PLWH and SN participants who were positive for SARS-CoV-2 was: 11.1% vs 6.7% at wave 1 ; 5.4% vs 3.1% at wave 2; 6.1% vs 2.3% at wave 3.

[~] During most of the time this survey was collected, SARS-CoV-2 antibody testing was not available outside of research settings. There were an additional 291 participants who reported having a blood test for SARS-CoV2; these antibody test results were not included in this analysis.

[#] 70 participants had wave 4 results which are included here as their most recent results were reported.

[^] Participants who had multiple test results are included as positive if ever positive in the “overall” description of test results across all waves above.

Table 3

COVID-19 symptom severity among 95 SARS-CoV-2 positive participants, by HIV serostatus

	Symptom Severity Among SARS-CoV-2 positive Participants					P-value PLWH vs SN
	All N=95			PLWH N=72	SN N=23	
	% mild or none	% moderate	% severe	% severe	% severe	
Number of symptoms reported						
0 (N=8)	100%	0 %	0 %	0 %	0 %	NA
1–2 (N=7) *	71%	29%	0 %	0 %	0 %	NA
3 (N=80) *	5%	26%	69%	63%	85%	0.16
<hr/>						
3 <i>severe</i> symptoms	NA	NA	35%	33%	39%	0.61
<hr/>						
Individual symptoms reported						
Headache	51%	34%	15%	17%	9%	0.43
Myalgias (muscle aches)	50%	24%	26%	24%	35%	0.57
Shortness of breath	63%	21%	16%	13%	26%	0.27
Chills	61%	26%	13%	10%	22%	0.30
Felt feverish	67%	21%	12%	8%	22%	0.13
Fever (Temp>100.4F)	81%	9%	10%	6%	22%	0.07
Loss of taste or smell	53%	21%	26%	28%	22%	0.66
Runny nose (rhinorrhea)	79%	19%	2%	3%	0 %	0.21
Cough	59%	23%	18%	18%	17%	0.63
Sore throat	74%	19%	7%	7%	9%	0.87
Diarrhea	70%	12%	18%	17%	22%	0.79
Nausea or vomiting	81%	14%	5%	6%	4%	0.43
Abdominal pain	84%	10%	6%	7%	4%	0.31

PLWH = People Living with HIV; SN= HIV Seronegative

* For symptom severity stratified by number of symptoms, individuals were categorized by the most severe symptom they had. So individuals with 3 symptoms, for example, were classified as “mild or none” if all symptoms were mild, as “moderate” if they had 1 moderate symptom and no severe symptoms, and as “severe” if they had 1 severe symptom.