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**RESEARCH ARTICLE** 

COVID-19 self-testing using antigen rapid diagnostic tests: Feasibility evaluation among health-care workers and general population in Malawi

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# Abstract

# Background

COVID-19 testing is critical for identifying cases to prevent transmission. COVID-19 selftesting has the potential to increase diagnostic testing capacity and to expand access to hard-to-reach areas in low-and-middle-income countries. We investigated the feasibility and acceptability of COVID-19 self-sampling and self-testing using SARS-CoV-2 Antigen-Rapid Diagnostic Tests (Ag-RDTs).

# Methods

From July 2021 to February 2022, we conducted a mixed-methods cross-sectional study examining self-sampling and self-testing using Standard Q and Panbio COVID-19 Ag Rapid Test Device in Urban and rural Blantyre, Malawi. Health care workers and adults (18y+) in the general population were non-randomly sampled.

# Results

Overall, 1,330 participants were enrolled of whom 674 (56.0%) were female and 656 (54.0%) were male with 664 for self-sampling and 666 for self-testing. Mean age was 30.7y (standard deviation [SD] 9.6). Self-sampling usability threshold for Standard Q was 273/333 (82.0%: 95% CI 77.4% to 86.0%) and 261/331 (78.8%: 95% CI 74.1% to 83.1%) for Panbio. Self-testing threshold was 276/335 (82.4%: 95% CI 77.9% to 86.3%) and 300/332 (90.4%: 95% CI 86.7% to 93.3%) for Standard Q and Panbio, respectively. Agreement between self-sample results and professional test results was 325/325 (100%) and 322/322 (100%) for Standard Q and Panbio, respectively. For self-testing, agreement was 332/333 (99.7%: 95% CI 98.3 to 100%) for Standard Q and 330/330 (100%: 95% CI 99.8 to 100%) for Panbio.

Tropical Medicine Data Compass <u>https://</u> datacompass.lshtm.ac.uk/.

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Odds of achieving self-sampling threshold increased if the participant was recruited from an urban site (odds ratio [OR] 2.15 95% CI 1.44 to 3.23, P < .01. Compared to participants with primary school education those with secondary and tertiary achieved higher self-testing threshold OR 1.88 (95% CI 1.17 to 3.01), P = .01 and 4.05 (95% CI 1.20 to 13.63), P = .02, respectively.

# Conclusions

One of the first studies to demonstrate high feasibility and acceptability of self-testing using SARS-CoV-2 Ag-RDTs among general and health-care worker populations in low- and middle-income countries potentially supporting large scale-up. Further research is warranted to provide optimal delivery strategies of self-testing.

# Introduction

Only around 0.2% of people in Africa had tested for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) the infection that causes COVID-19 in August 2020 [1] By contrast, 19.5% of Americans had tested by the same time [1] since COVID-19 emergence in December 2019 [2, 3]. These contrasting trends have continued to exist with widening unequal access to testing, treatment and vaccination between high income countries and low- and middleincome countries (LMICs) despite four or more global epidemic waves [4]. However, the access gap has become narrower as the pandemic has grown older [5]. Testing remains the most critical step for identification and isolation of COVID-19 cases to prevent transmission [6]. In many resource-limited settings, demand for tests often exceeds supply [7]. SARS-CoV-2 rapid antigen tests (Ag-RDTs) are recommended to complement nucleic acid amplification tests (NAAT) for diagnosis [8], which in resource-limited settings are often hard to implement because they require specialised skills and limited centralized laboratory capacity, associated with long turnaround times, and high costs to both the health system and patients [9, 10].

COVID-19 self-testing was strongly recommended by the World Health Organization (WHO) in March 2022 as an additional strategy to complement professionally administered testing services [11]. Self-sampling and self-testing is a process by which a person collects his or her own specimen using a simple device, performs a diagnostic test and interprets the results usually in a setting, and time of their choice [12]. Self-testing is not a new paradigm with pregnancy self-testing and HIV self-testing being successful examples [13–16]. In general, COVID-19 self-testing has the potential to increase diagnostic capacity for COVID-19 and reduce access barriers as well as prevailing inequalities due to ease of distribution and being extremely convenient [17]. However, COVID-19 self-testing has so far been widely implemented and made available in high income countries with reported high feasibility and acceptability [17–21]. According to our knowledge, there were no reports on COVID-19 self-testing feasibility in LMICs prior to this study. As with HIV self-testing, lack of linkage for next steps with COVID-19 is a potential concern due to stigma, loss of economic opportunities due to isolation implications, and fear of complications including death.

Being able to self-test rests on the assumption that individuals would be able to take their own sample (self-sampling) [22]. However, in settings with low exposure to technology and the ability to correctly follow instructions such assumptions may be faulty [23]. Thus, early work including optimization of instructions for use through iterative cognitive interviews is

essential to ensure correct use of self-tests [23]. Here we investigated the feasibility and acceptability of COVID-19 self-sampling and self-testing using SARS-CoV-2 Ag-rapid diagnostics tests (RDTs) in Malawi.

# Materials and methods

## Study design

A mixed-methods cross-sectional study examining self-sampling and self-testing for COVID-19 using STANDARD Q COVID-19 Ag Test (SD Biosensor) and Panbio COVID-19 Ag Rapid Test Device (Abbott Rapid Diagnostics). We conducted the study under five components. They were conducted serially as follows: cognitive interviews to refine instructions for use (IFUs) for self-sampling, observational cross-sectional study of self-sampling, cognitive interviews to refine instructions for use for self-testing, observational cross-sectional study of selftesting, and in-depth interviews (IDIs) to understand participant views on self-sampling and self-testing.

# Setting

Recruitment was conducted between July 2021 to February 2022 from Queen Elizabeth Central Hospital (QECH) from urban Blantyre, Malawi (Fig 1) and from Lirangwe Primary Health Centre from rural Blantyre (Fig 2). Sites were chosen to ensure inclusion of both rural and



Fig 1. Map of Blantyre with Queen Elizabeth Central Hospital marked. https://doi.org/10.1371/journal.pone.0289291.g001



Fig 2. Map of Blantyre with Lirangwe health centre marked. https://doi.org/10.1371/journal.pone.0289291.g002

urban participants whose literacy levels, technology exposure and perception to testing may differ. At the time of recruitment, COVID-19 testing in Blantyre was concentrated at QECH whereas Lirangwe health centre was only collecting samples to be tested at another facility.

### **Participants**

We recruited health care workers and members of the general public from the recruitment sites. To be eligible, participants needed to be 18 years or older, feeling well enough to comfortably conduct study activities, not having recent history of excessive nose bleeds, and having given consent. All health workers from the two health facilities were offered the choice to participate in the study with exclusion only done if ineligible. An additional eligibility criteria which was later relaxed due to scarcity of participants with waning wave concerned individuals being on the list to be tested for COVID-19 by the national systems. General public participants were non randomly sampled from outpatient departments. An anterior nasal swab for COVID-19 was done for both self-sampling and self-testing following a short in-person demonstration by a member of staff.

Participants were observed in-person during self-sampling and self-testing, and checklists (Tables A-D in <u>S1 text</u>) were completed to document whether each task was done correctly. A trained researcher then tested the collected sample using a SARS-CoV-2 Ag RDT during the self-sampling component of the study. Participants tested their own collected sample during the self-testing component of the study. The trained researcher collected and tested an anterior

nasal confirmatory sample using an Ag RDT during both the self-sampling and self-testing components.

#### Variables

For the cognitive interviews, the main output was to have refined IFUs in the local language (Chichewa) and in English. The first primary outcome was the percentage of participants who attained a usability threshold for self-sampling, defined as correct execution of all critical instructions during the self-sampling process for each kit. Correct self-sampling was referred to as self-sampling accuracy. The second primary outcome was the percentage of participants who attained a usability threshold for self-testing, defined as correct execution of all critical instructions during the self-testing process. Correct self-testing was referred to as self-testing accuracy. User views regarding self-sampling and self-testing were the main outcomes from the IDIs. Potential confounders for accuracy were age, sex, literacy and prior exposure to COVID-19 testing.

#### Data sources/ measurement

Qualitative data from cognitive interviews and IDIs were tape recorded before being translated and transcribed. Pre- and post-test questionnaires were administered in-person using Open Data Kit (ODK) loaded on tablets. A checklist (Tables A-D in S1 text) was completed by a member of staff to document whether each instruction was done correctly as a measure of accuracy. Results obtained by a trained researcher from testing the collected self-sample and a sample collected by the researcher were recorded on the checklist. For self-testing accuracy, participant's self-test selfread results were compared to RDT sampling and testing conducted by the researcher. Participants' reading of pre-made cassettes of negative, positive and invalid results was also recorded.

# Bias

The main source of bias is in the assessment by the research staff using a checklist of the performance of the participant on the IFU. A staff member who was more punitive may have harshly rated performance as incorrect while a more forgiving one may have rated performance differently. However, the fact that more than seven staff members were involved in the rating may have minimized such bias.

#### Study size

We aimed to recruit and purposively sample 120 participants for cognitive interviews for selfsampling and self-testing for both test kits. For self-sampling and self-testing, we conservatively assumed that 70% to 80% of participants will be able to correctly follow instructions and self-sample or self-test for COVID-19. For the sample proportion to be estimated to within +/-0.05 (5%) using the 95% confidence level, a sample of 323 participants were required. Thus, a total of 1,320 participants were needed: 330 per test kit for self-sampling and self-testing. A purposive sample of 120 participants was needed for the IDIs: 60 self-sampling and 60 self-testing participants.

#### Quantitative variables

A binary variable was generated for the first and secondary primary outcomes of achieving the threshold (accuracy) for either self-sampling or self-testing. This was coded as 1 for participants with a maximum score on the critical steps based on the checklist and 0 otherwise. Test result variables were coded as 1 for positive and 0 for negative.

### Statistical methods

Analysis used R [24] with 0.05 as an indicator of statistical significance. Frequencies were computed for categorical variables while mean and standard deviation (SD) or median and (inter quartile range) were computed for continuous variables that were normally distributed or skewed, respectively. We computed the proportion achieving accuracy along with Binomial Exact confidence intervals (CIs) for self-sampling and self-testing for each test kit. Similarly, we computed the proportion of self-test results that agreed with staff conducted RDT test results along with Binomial Exact confidence intervals (CIs) for each test kit. Logistic regression was used to examine factors associated with accuracy.

#### Ethics statement

The study was conducted according to the guidelines of the Declaration of Helsinki, and approved by the Malawi College of Medicine Research Ethics Committee of Kamuzu University of Health Sciences (Reg No: P.03/21/3277) and the World Health Organization Research Ethics Review Committee (Protocol ID: CERC.0104). Informed consent was obtained from all participants involved in the study.

### Results

#### Participants and descriptive data

A total of 1,510 participants were recruited across the five components of the study. A total of 120 participants were recruited for self-sampling and self-testing cognitive interviews (self-sampling (n = 76) and self-testing (n = 44)). Of 723 screened for eligibility 664 (91.8%) were recruited for self-sampling with mean age of 31.4y (standard deviation [SD]: 9.8) and 357/664 (53.8%) were male (Table 1). For self-testing, 666 (95.4%) were recruited of 698 screened for eligibility; mean age was 30.6y (standard deviation [SD]: 9.6) with 293/666 (44.0%) being male (Table 2). The main exclusion was being under 18 years. Sixty participants were recruited for IDIs.

## Outcome data

The cognitive interviews showed that participants in both rural and urban communities were able to follow the IFUs with no major suggestions for changes. Notable changes to IFUs included: making introductory text stand out to catch attention, enhancing clarity of IFUs such as by expanding text, adding labels on images, selecting words or phrases that could be well understood locally. Insertion of test swab to correct depth (1.5cm or 2cm) was illustrated by reference to inserting up to thumbnail depth.

#### Main results

Self-sampling accuracy was 273/333 (82.0%: 95% CI: 77.4 to 86.0) for Standard Q and 261/331 (78.8: 95% CI: 74.1% to 83.1%) for Panbio (Table 3). The percentage agreement between the test results from the participant and the study staff was 100% for both kits in Malawi (Table 3).

Self-testing accuracy was 276/335 (82.4%: 95% CI: 77.9 to 86.3) for Standard Q and 300/332 (90.4%: 95% CI: 86.7 to 93.3) for Panbio (Table 3). The percentage agreement between the test results from the participant and the study staff was 99.7% (95% CI: 98.3–100%) for Standard Q with only one false negative self-test self-read result.

Up to 95% of the critical steps were performed correctly on either test kit for both self-sampling and self-testing (Table 4).

The odds of self-sampling accuracy increased 2-fold for participants from QECH compared to participants from Lirangwe primary health centre odds ratio (OR) 2.15 (95% CI 1.44 to

Variable	Characteristic	Overall	Standard Q	Panbio	p-value <sup>a</sup>
Number of participants	n	664	331	333	
Sex	Male	357 (53.8)	176 (53.2)	181 (54.4)	0.820
	Female	307 (46.2)	155 (46.8)	152 (45.6)	
Age (years)	mean (SD)	31.4 (9.8)	31.8 (10.3)	31.0 (9.3)	0.313
Ever tested for COVID-19?	No	568 (87.0)	303 (91.8)	265 (82.0)	< 0.001
	Yes	85 (13.0)	27 (8.2)	58 (18.0)	
Marital status	Divorced	37 (5.7)	20 (6.1)	17 (5.3)	0.512
	Separated	33 (5.1)	15 (4.5)	18 (5.6)	
	Widowed	14 (2.1)	8 (2.4)	6 (1.9)	
	Never married	180 (27.6)	82 (24.8)	98 (30.3)	
	Married	389 (59.6)	205 (62.1)	184 (57.0)	
Money earned per month (MWK)	mean (SD)	68191 (96060)	57471 (90035)	79111 (100803)	0.004
Able to read a newspaper?	No	42 (6.4)	29 (8.8)	13 (4.0)	0.020
	Yes	610 (93.6)	300 (91.2)	310 (96.0)	
Highest level of formal schooling	Never been to school	25 (3.9)	18 (5.5)	7 (2.2)	< 0.001
	Primary	165 (25.4)	101 (31.0)	64 (19.8)	
	Secondary no MSCE	216 (33.3)	118 (36.2)	98 (30.3)	
	Secondary with MSCE	131 (20.2)	54 (16.6)	77 (23.8)	
	Tertiary	112 (17.3)	35 (10.7)	77 (23.8)	
Number of people in household	mean (SD)	4.2 (1.8)	4.2 (1.8)	4.2 (1.8)	0.903
Number of rooms in household	mean (SD)	2.5 (1.0)	2.4 (1.0)	2.6 (1.1)	0.032
Number of households per dwelling	mean (SD)	1.5 (1.3)	1.3 (0.8)	1.7 (1.7)	< 0.001
Enough food / essentials for 14 days?	No	412 (63.1)	215 (65.2)	197 (61.0)	0.307
	Yes	241 (36.9)	115 (34.8)	126 (39.0)	
Recruitment site	QECH	331 (49.8)	165 (49.8)	166 (49.8)	1.000
	Lirangwe	333 (50.2)	166 (50.2)	167 (50.2)	

#### Table 1. Baseline characteristics: Self-sampling.

<sup>a</sup>Chisquare test for categorical variables; t-test for continuous variables SD: standard deviation; QECH: Queen Elizabeth Central Hospital

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3.23, P < 0.001 (Table 5). There appeared to be a linear trend towards increased odds of attaining self-testing accuracy with increasing levels of education, *P* for trend 0.01.

# Other analyses

All in-depth interview participants reported that self-testing was highly acceptable because it was convenient, empowering and private.

Most participants had no problems interpreting contrived panel results with 99% correctly interpreting positive and negative results correctly although 96% correctly interpreted invalid results on either test kit (<u>S1 Table</u>). Up to 90.7% Standard Q and 96.1% Panbio participants found instructions "not at all hard" when asked on exit interviews (<u>S2 Table</u>).

# Discussion

## Key results

This is one of first studies, to the best of our knowledge, to be conducted on COVID-19 selftesting in low- and middle-income countries and generally indicates that participants in both rural and urban communities in Malawi can self-test correctly for COVID-19. The results of

Variable	Characteristic	Overall	Standard Q	Panbio	p-value <sup>a</sup>	
Number of participants	n	664	336	328		
Sex	Male	292 (44.0)	138 (41.1)	154 (47.0)	0.148	
	Female	372 (56.0)	198 (58.9)	174 (53.0)		
Age (years)	mean (SD)	30.7 (9.6)	30.8 (9.8)	30.52 (9.3)	0.724	
Ever tested for COVID-19?	No	603 (91.5)	303 (91.0)	300 (92.0)	0.659	
	Yes	56 (8.5)	30 (9.0)	26 (8.0)		
Marital status	Divorced	37 (5.6)	15 (4.5)	22 (6.8)	0.208	
	Separated	47 (7.2)	24 (7.2)	23 (7.1)		
	Widowed	25 (3.8)	13 (3.9)	12 (3.7)		
	Never married	195 (29.7)	88 (26.5)	107 (32.9)		
	Married	353 (53.7)	192 (57.8)	161 (49.5)		
Money earned per month (MWK)	mean (SD)	67802 (123668)	70359 (153192)	65190 (83531)	0.591	
Able to read a newspaper?	No	59 (8.9)	29 (8.7)	30 (9.2)	0.922	
I · I ·	Yes	601 (91.1)	305 (91.3)	296 (90.8)		
Highest level of formal schooling	Never been to school	33 (5.0)	19 (5.7)	14 (4.3)	0.209	
	Primary	208 (31.5)	109 (32.6)	99 (30.4)		
	Secondary no MSCE	220 (33.3)	117 (35.0)	103 (31.6)		
	Secondary with MSCE	143 (21.7)	68 (20.4)	75 (23.0)		
	Tertiary	56 (8.5)	21 (6.3)	35 (10.7)		
Number of people in household	mean (SD)	4.2 (1.6)	4.2 (1.7)	4.3 (1.6)	0.548	
Number of rooms in household	mean (SD)	2.5 (1.2)	2.5 (1.0)	2.5 (1.4)	0.559	
Number of households per dwelling	mean (SD)	1.5 (1.3)	1.5 (1.6)	1.5 (0.9)	0.516	
Enough food / essentials for 14 days?	No	351 (53.3)	188 (56.5)	163 (50.0)	0.113	
	Yes	308 (46.7)	145 (43.5)	163 (50.0)		
Recruitment site	QECH	334 (50.8)	169 (51.1)	165 (50.6)	0.971	
	Lirangwe	323 (49.2)	162 (48.9)	161 (49.4)		

Table 2. Baseline characteristics: Self-testing.

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this study show that 82% and 90% of participants were able to self-test for COVID-19 with no supervision following a brief demonstration using Standard Q and Panbio test kits, respectively. Of further note, all self-test results agreed 100% with professionally conducted RDTs for Panbio kit whereas agreement was 99.7% for Standard Q. Similarly, 82% of participants were able to correctly self-sample for COVID-19 using Standard Q compared to 79% using Panbio. COVID-19 self-testing was rated as highly acceptable during in-depth interviews.

#### Table 3. Self-sampling and self-testing accuracy.

	Standard Q					Panbio				
	N	n	%	95%	5 CI	N	n	%	95%	5 CI
Met self-sampling threshold <sup>a</sup>	333	273	82.0	77.4	86.0	331	261	78.8	74	83.1
Met self-testing threshold	335	276	82.4	77.9	86.3	332	300	90.4	86.7	93.3
Agreement with professional test										
Self-sampling	322	322	100	99	100	325	325	100	100	100
Self-testing	333	332	99.7	98	100	330	330	100	100	100

<sup>a</sup>Threshold: participant performing all critical steps correctly CI: confidence interval

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	Standard Q (N = 331)		Pa1 (N =	1bio 331)
	Yes	No	Yes	No
Did the participant place the tube on the kit box tray holder or flat surface correctly?	327 (97.6)	8 (2.4)	329 (99.4)	2 (0.6)
Did participant insert the swab into the left nostril to the correct depth (about 1.5cm or 2cm)?	327 (97.6)	8 (2.4)	332 (100)	0 (0.0)
Did the participant rotate the swab 5 or 10 times in the left nostril?	325 (97.0)	10 (3.0)	327 (98.5)	5 (1.5)
Did participant insert the swab into the right nostril to the correct depth (about 1.5cm or 2cm)?	327 (98.5)	5 (1.5)	329 (99.4)	2 (0.6)
Did the participant rotate the swab 5 or 10 times in the right nostril?	324 (97.0)	10 (3.0)	330 (99.4)	2 (0.6)
Did the participant insert the swab into the solution tube correctly?	331 (98.8)	4 (1.2)	328 (99.4)	2 (0.6)
Did the participant swirl in the fluid 5 or 10 times while pushing against the wall of the tube?	325 (97.3)	9 (2.7)	323 (97.6)	8 (2.4)
Did the participant remove the swab slowly while squeezing the sides of the tube to extract the liquid from the swab?	310 (92.5)	25 (7.5)	316 (95.5)	15 (4.5)
Did the participant press the nozzle cap tightly the tube?	326 (97.9)	7 (2.1)	330 (99.7)	1 (0.3)
Did the participant squeeze 4 or 5 drops of liquid from the tube into the well on the test device?	319 (95.5)	15 (4.5)	330 (99.7)	1 (0.3)
Did the participant read the test result in 15 minutes?	333 (99.7)	1 (0.3)	325 (97.9)	7 (2.1)
Did the participant interpret the test result correctly?	328 (98.5)s	2 (1.5)	329 (99.7)	1 (0.3)

#### Table 4. User errors for Standard Q and Panbio kits.

Standard Q: swab 10 times, depth 2cm, 4 drops

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Current strategies for COVID-19 testing in high income countries are largely dependent on Ag-RDT self-sampling and self-testing [25–27] with over-the-counter self-test kits available for purchase in a wide range of countries [21, 28]. The limited data available in resource-poor settings suggest that, as with HIV self-testing, diagnostic accuracy is not as great with untrained lay users as with trained professionals, mainly affecting sensitivity [20, 25, 29]. Our results on the other hand show that self-testing accuracy improved markedly with a short demonstration supporting previous findings observed with HIV self-testing [14, 30]. However, there is still a place for well-translated and culturally relevant IFUs to support the large-scale implementation of self-testing. Our study investigated self-testing with two kits that were already approved for use in Malawi. However, there are numerous Ag-RDT tests packaged for COVID-19 self-testing that have met performance standards and been approved by Regulatory Authorities such as the FDA, that may yield similar promising results [31, 32].

#### Limitations

There are notable limitations with our study. Firstly, there was a small number of positive selftest results. Although this does not affect the reading of correct results and indeed completing critical steps correctly as assessed here it may be important as it is likely to affect sensitivity [33]. Reassuringly, up to 99% of participants correctly interpreted contrived positive results on either kit. Secondly, there was potential for assessment bias resulting from subjective judgement on the checklist used by research staff for assessing performance of the participant on

Variable Age	Characteristic Yearly increase	Self-sampling (N = 641) Unadjusted				Self-testing (N = 637) Unadjusted			
		0.99	0.97	1.01	0.247	0.99	0.97	1.01	0.299
		Sex	Female	1.00				1.00	
	Male	1.00	0.68	1.48	0.991	0.84	0.54	1.33	0.459
Site	Lirangwe	1.00				1.00			
	QECH	2.15	1.44	3.23	< 0.001	1.47	0.93	2.32	0.097
Literacy	No	1.00				1.00			
	Yes	0.57	0.22	1.49	0.251	1.21	0.57	2.55	0.625
Highest level of education attained?	Primary school	1.00				1.00			
	Never been school	2.75	0.61	12.3	0.186	1.67	0.55	5.02	0.362
	Secondary	0.91	0.58	1.44	0.686	1.88	1.17	3.01	0.009
	Tertiary	2.09	1.03	4.25	0.041	4.05	1.20	13.63	0.024
Marital Status	Divorced/separated/widowed	1.00				1.00			
	Never married	2.10	1.06	4.14	0.033	2.41	1.26	4.61	0.008
	Married	1.07	0.61	1.89	0.820	1.89	1.09	3.28	0.023
Ever tested for COVID-19?	No	1.00				1.00			
	Yes	1.04	0.58	1.86	0.905	1.25	0.52	3.02	0.619

#### Table 5. Factors associated with self-sampling and self-testing accuracy.

OR: odds ratio; CI: confidence interval; QECH: Queen Elizabeth Central Hospital

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each instruction. The impact of this bias could be bi-directional depending on whether the staff was harsh-leading to poor rating, or more lenient resulting in more participants being passed as correctly following instructions.

# Generalisability

This study demonstrates high acceptability and feasibility of COVID-19 self-testing [11]. The findings are very similar to results reported in other self-testing areas including HIV [34] and hepatitis C virus (HCV) [35]. Thus, we posit that the findings are generalizable to many resource settings and populations including those with limited literacy. However, some support may be useful for specific settings and users–such as older age groups and those with lower literacy. Lessons learned from introduction and scale-up of other self-testing approaches such as HIV and HCV may be appliable here to accelerate adaptation plans and efforts in LMIC.

# Conclusions

This is one the first studies to demonstrate high usability and acceptability of self-testing using SARS-CoV-2 Ag-RDTs among both general and health-care worker populations in LMICs. While most users collected their own samples and self-tested with ease, participants noted demonstrations were helpful and could be important in some settings and populations, such as older age groups and those with low literacy levels. COVID-19 self-testing is an important strategy for further consideration as it may be a promising tool for increasing access to and uptake of COVID-19 testing services as well as strategies to reduce transmission and linkage to further care, treatment and support services. Further research is warranted to provide optimal delivery strategies to reach priority populations in LMICs.

# **Supporting information**

**S1 Text.** Checklists for self-sampling and self-testing-Standard Q and Panbio. (DOCX)

**S1 Table. Interpreting contrived panel results.** (DOCX)

**S2** Table. User views on self-sampling and self-testing. (DOCX)

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#### References

- Chitungo I, Dzobo M, Hlongwa M, Dzinamarira T. COVID-19: Unpacking the low number of cases in Africa. Public Health in Practice. 2020; 1: 100038. https://doi.org/10.1016/j.puhip.2020.100038 PMID: 34173573
- Zoumpourlis V, Goulielmaki M, Rizos E, Baliou S, Spandidos DA. The COVID-19 pandemic as a scientific and social challenge in the 21st century. Mol Med Rep. 2020; 22: 3035. https://doi.org/10.3892/ MMR.2020.11393 PMID: 32945405
- 3. World Health Organization. WHO-convened global study of origins of SARS-CoV-2: China Part. In: WHO [Internet]. 2021 [cited 11 Aug 2022]. Available: https://www.who.int/publications/i/item/who-convened-global-study-of-origins-of-sars-cov-2-china-part
- Zhang SX, Arroyo Marioli F, Gao R, Wang S. A Second Wave? What Do People Mean by COVID Waves?–A Working Definition of Epidemic Waves. Risk Manag Healthc Policy. 2021; 14: 3775. https:// doi.org/10.2147/RMHP.S326051 PMID: 34548826
- Foundation for Innovative New Diagnostics. COVID-19 Test tracker—FIND. In: FIND [Internet]. [cited 1 Jun 2023]. Available: https://www.finddx.org/tools-and-resources/dxconnect/test-directories/covid-19test-tracker/
- World Health Organisation. Contact tracing and quarantine in the context of COVID-19: interim guidance, 6 July 2022. In: WHO [Internet]. 2022 [cited 11 Aug 2022]. Available: <a href="https://www.who.int/publications/i/item/WHO-2019-nCoV-Contact\_tracing\_and\_quarantine-2022.1">https://www.who.int/ publications/i/item/WHO-2019-nCoV-Contact\_tracing\_and\_quarantine-2022.1</a>
- Boum Y, Eyangoh S, Okomo MC. Beyond COVID-19—will self-sampling and testing become the norm? Lancet Infect Dis. 2021; 21: 1194. https://doi.org/10.1016/S1473-3099(21)00197-3 PMID: 33857408
- Drain PK. Rapid Diagnostic Testing for SARS-CoV-2. New England Journal of Medicine. 2022; 386: 264–272. https://doi.org/10.1056/NEJMcp2117115 PMID: 34995029
- Kobia F, Gitaka J. Open Peer Review COVID-19: Are Africa's diagnostic challenges blunting response effectiveness? [version 1; peer review: 2 approved]. 2020 [cited 24 May 2023]. https://doi.org/10.12688/ aasopenres.13061.1 PMID: 32399515
- Shey M, Okeibunor JC, Yahaya AA, Herring BL, Tomori O, Coulibaly SO, et al. Genome sequencing and the diagnosis of novel coronavirus (SARS-COV-2) in Africa: how far are we? Pan Afr Med J. 2020; 36: 1–10. https://doi.org/10.11604/PAMJ.2020.36.80.23723 PMID: 32774639
- World Health Organisation. Use of SARS-CoV-2 antigen-detection rapid diagnostic tests for COVID-19 self-testing. In: WHO [Internet]. 2022 [cited 11 Aug 2022]. Available: <u>https://www.who.int/publications/i/</u> item/WHO-2019-nCoV-Ag-RDTs-Self\_testing-2022.1
- 12. World Health Organisation. COVID-19 RAPID ANTIGEN SELF-TESTING FEBRUARY 2022 2 COVID-19 RAPID ANTIGEN SELF-TESTING. 2022.
- Indravudh PP, Choko AT, Corbett EL. Scaling up HIV self-testing in sub-Saharan Africa: a review of technology, policy and evidence. <u>https://doi.org/10.1097/QCO.0000000000426</u>
- 14. Talumba Choko A, Desmond N, Webb EL, Chavula K, Napierala-Mavedzenge S, Gaydos CA, et al. The Uptake and Accuracy of Oral Kits for HIV Self-Testing in High HIV Prevalence Setting: A Cross-Sectional Feasibility Study in Blantyre, Malawi. https://doi.org/10.1371/journal.pmed.1001102
- 15. Johnson CC, Kennedy C, Fonner V, Siegfried N, Figueroa C, Dalal S, et al. Examining the effects of HIV self-testing compared to standard HIV testing services: a systematic review and meta-analysis. J Int AIDS Soc. 2017;20. https://doi.org/10.7448/IAS.20.1.21594 PMID: 28530049
- 16. Kennedy CE, Yeh PT, Gholbzouri K, Narasimhan M. Self-testing for pregnancy: a systematic review and meta-analysis. Reproductive Health and Research. 2022;12. <u>https://doi.org/10.1136/bmjopen-2021-054120 PMID: 35228285</u>
- Stohr JJJM Zwart VF, Goderski G, Meijer A, Nagel-Imming CRS, Kluytmans-van den Bergh MFQ, et al. Self-testing for the detection of SARS-CoV-2 infection with rapid antigen tests for people with suspected COVID-19 in the community. Clinical Microbiology and Infection. 2022; 28: 695. https://doi.org/10.1016/ j.cmi.2021.07.039 PMID: 34363945
- Herbert C, Broach J, Heetderks W, Qashu F, Gibson L, Pretz C, et al. Feasibility of At-Home Serial Testing Using Over-the-Counter SARS-CoV-2 Tests With a Digital Smartphone App for Assistance: Longitudinal Cohort Study. JMIR Form Res 2022; 6(10):e35426 https://formative.jmir.org/2022/10/ e35426. 2022;6: e35426. https://doi.org/10.2196/35426 PMID: 36041004
- Hirst JA, Logan M, Fanshawe TR, Mwandigha L, Wanat M, Vicary C, et al. Feasibility and Acceptability of Community Coronavirus Disease 2019 Testing Strategies (FACTS) in a University Setting. Open Forum Infect Dis. 2021;8. https://doi.org/10.1093/OFID/OFAB495 PMID: 34904117

- Lindner AK, Nikolai O, Rohardt C, Kausch F, Wintel M, Gertler M, et al. Diagnostic accuracy and feasibility of patient self-testing with a SARS-CoV-2 antigen-detecting rapid test. Journal of Clinical Virology. 2021; 141: 104874. https://doi.org/10.1016/j.jcv.2021.104874 PMID: 34144452
- NHS UK. How to do a rapid lateral flow test for coronavirus (COVID-19)—NHS. In: NHS [Internet]. 2022 [cited 11 Aug 2022]. Available: https://www.nhs.uk/conditions/coronavirus-covid-19/testing/howto-do-a-test-at-home-or-at-a-test-site/how-to-do-a-rapid-lateral-flow-test/
- Lindner AK, Nikolai O, Rohardt C, Burock S, Hülso C, Bölke A, et al. Head-to-head comparison of SARS-CoV-2 antigen-detecting rapid test with professional-collected nasal versus nasopharyngeal swab. European Respiratory Journal. 2021;57. https://doi.org/10.1183/13993003.04430–2020
- Simwinga M, Kumwenda MK, Dacombe RJ, Kayira L, Muzumara A, Johnson CC, et al. Ability to understand and correctly follow HIV self-test kit instructions for use: applying the cognitive interview technique in Malawi and Zambia. J Int AIDS Soc. 2019; 22 Suppl 1. <u>https://doi.org/10.1002/jia2.25253</u> PMID: 30907496
- R Core Team. R: The R Project for Statistical Computing. 2022 [cited 7 Jun 2023]. Available: <a href="https://www.r-project.org/">https://www.r-project.org/</a>
- 25. Wachinger J, Olaru ID, Horner S, Schnitzler P, Heeg K, Denkinger CM. The potential of SARS-CoV-2 antigen-detection tests in the screening of asymptomatic persons. Clinical Microbiology and Infection. 2021; 27: 1700.e1–1700.e3. https://doi.org/10.1016/j.cmi.2021.07.020 PMID: 34325065
- 26. Young BC, Eyre DW, Kendrick S, White C, Smith S, Beveridge G, et al. Daily testing for contacts of individuals with SARS-CoV-2 infection and attendance and SARS-CoV-2 transmission in English secondary schools and colleges: an open-label, cluster-randomised trial. The Lancet. 2021; 398: 1217–1229. https://doi.org/10.1016/S0140-6736(21)01908-5 PMID: 34534517
- 27. European Centre for Disease Prevention and Control. Considerations on the use of self-tests for COVID-19 in the EU/EEA. 2021 [cited 7 Jun 2023]. Available: https://www.ecdc.europa.eu/en/publications-data/considerations-use-self-tests-covid-19-eueea
- 28. Government UK. Order COVID-19 rapid lateral flow tests—GOV.UK. 2022 [cited 7 Jun 2023]. Available: https://www.gov.uk/order-coronavirus-rapid-lateral-flow-tests
- Lindner AK, Nikolai O, Rohardt C, Kausch F, Wintel M, Gertler M, et al. SARS-CoV-2 patient self-testing with an antigen-detecting rapid test: a head-to-head comparison with professional testing. medRxiv. 2021; 2021.01.06.20249009. https://doi.org/10.1101/2021.01.06.20249009
- 30. Choko AT, MacPherson P, Webb EL, Willey BA, Feasy H, Sambakunsi R, et al. Uptake, Accuracy, Safety, and Linkage into Care over Two Years of Promoting Annual Self-Testing for HIV in Blantyre, Malawi: A Community-Based Prospective Study. PLoS Med. 2015; 12: e1001873. <u>https://doi.org/10. 1371/journal.pmed.1001873 PMID: 26348035</u>
- Peeling RW, Olliaro PL, Boeras DI, Fongwen N. Scaling up COVID-19 rapid antigen tests: promises and challenges. Lancet Infect Dis. 2021; 21: e290. <u>https://doi.org/10.1016/S1473-3099(21)00048-7</u> PMID: 33636148
- 32. Foundation for Innovative New Diagnostics. Test directory—FIND. In: FIND [Internet]. 2022 [cited 12 Aug 2022]. Available: https://www.finddx.org/covid-19/test-directory/
- Altman DG, Bland JM. Diagnostic tests. 1: Sensitivity and specificity. BMJ: British Medical Journal. 1994; 308: 1552. https://doi.org/10.1136/bmj.308.6943.1552 PMID: 8019315
- Pant Pai N, Sharma J, Shivkumar S, Pillay S, Vadnais C, Joseph L, et al. Supervised and Unsupervised Self-Testing for HIV in High- and Low-Risk Populations: A Systematic Review. PLoS Med. 2013; 10: e1001414. https://doi.org/10.1371/journal.pmed.1001414 PMID: 23565066
- 35. Reipold EI, Farahat A, Elbeeh A, Soliman R, Aza EB, Jamil MS, et al. Usability and acceptability of self-testing for hepatitis C virus infection among the general population in the Nile Delta region of Egypt. BMC Public Health. 2021; 21: 1–11. https://doi.org/10.1186/S12889-021-11169-X/TABLES/4