Leveraging HIV Program and Civil Society to Accelerate COVID-19 Vaccine Uptake, Zambia

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To accelerate COVID-19 vaccination delivery, Zambia integrated COVID-19 vaccination into HIV treatment centers and used World AIDS Day 2021 to launch a national vaccination campaign. This campaign was associated with significantly increased vaccinations, demonstrating that HIV programs can be leveraged to increase COVID-19 vaccine uptake.

OVID-19 vaccine scale-up in Africa, the continent with the lowest vaccine coverage, is a current regional and global priority. As of May 1, 2022, only 17% of persons in Africa had been fully vaccinated (1). Initial vaccination campaigns in Africa were hampered by lower-than-forecasted vaccine donations (2). However, through efforts from multiple stakeholders, the vaccine supply to countries in Africa increased in the latter half of 2021. However, with increasing vaccine availability, new challenges became apparent, including the difficulty for under-resourced health systems with relatively low healthcare worker-to-population ratios to implement COVID-19 vaccination services, as well as difficulties reaching populations unaccustomed to adult immunization programs and vaccine misperceptions and misinformation. Facing these challenges, in August 2021, the government of Zambia worked with stakeholders to leverage its national HIV program (which has been supported by >\$5 billion in funding in the previous 20 years) to enhance its COVID-19 vaccine campaign.

Zambia integrated COVID-19 vaccination into its existing HIV treatment centers with the goal of offering patients and family members vaccination services, thereby rapidly expanding static vaccination site num-

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DOI: https://doi.org/10.3201/eid2813.220743

bers in the country. Successful strategies for engaging HIV treatment centers included using existing human resources by adequately preparing HIV healthcare workers to offer vaccination and encouraging them to get vaccinated themselves, developing targeted promotional materials for persons living with HIV who are at increased risk for severe illness (3), and rapidly adapting and implementing similar models across the country. After this preparatory work, Zambia used the annual World AIDS Day event to launch its December Campaign to help reach African Union targets (4), focusing on engaging civil society leaders to endorse vaccination and using a mixed service delivery model that added community-delivered vaccination based on successful community HIV programs to existing static service delivery (Table). Some strategies were adapted from Zambia's robust childhood vaccination program (5).

To evaluate whether the December Campaign accelerated COVID-19 vaccination in Zambia, we conducted time-series analyses by using publicly available data (Appendix, https://wwwnc.cdc.gov/EID/ article/28/13/22-0743-App1.pdf) (1). All participants entered in the Our World in Data (https://ourworldindata.org/ dataset by February 21, 2022, for Zambia and 55 African Union member states were eligible for the analysis. We conducted 3 statistical analyses. First, in a single-group interrupted time-series analysis in Zambia only, we compared the number of persons reaching full vaccination status per day before the December 1, 2021, campaign start versus after the campaign start. Second, in a multigroup interrupted time-series analysis, we assessed whether Zambia's acceleration in COVID-19 vaccination coverage (i.e., acceleration in the percentage of total population reaching full vaccination status per day) after the December Campaign intervention was statistically superior to 2 control groups: 2 neighboring countries with similar pre-intervention vaccination coverage trajectories and similar vaccine availability, and the average for all 55 Africa Union member states. Third, we implemented 2 sensitivity analyses for each of the above 2 analytic approaches by varying the approach to managing missing data (i.e., most recent value carried forward approach vs. interpolation approach) and comparing varied time periods for the analysis to determine the duration of December Campaign effect.

During December 2021, a total of 585,677 persons in Zambia were reached for vaccination, compared with approximately 1,071,682 million during April-November 2021. Daily COVID-19 vaccinations increased from 3,713/day before December 2021 to 17,783/day after December 1, 2021 (p<0.001) (Figure, panel A; Appendix Table 3).

Compared with the average for 2 neighboring countries with similar vaccination trends before December and vaccine availability, Zambia accelerated its population COVID-19 vaccine coverage rate by an additional 2.73%/month (p<0.001) (Figure, panel B; Appendix Table 4). Compared with Africa as a whole, Zambia vaccine coverage accelerated by 1.87%/month (p<0.001) (Figure, panel C; Appendix Table 5). This accelerated vaccination in Zambia was robust to the sensitivity analysis for which we used an interpolation approach to missing data instead of the approach carrying forward the most recent available data point (Appendix Tables 3, 6). In addition, the average post-December daily vaccination rate dropped only slightly, and the average post-December percentage gain per day in a fully vaccinated population remained relatively stable, indicating a sustained effect for nearly 3 months after the December Campaign launch. If current trends were sustained,

Pillar	Lessons
Planning and coordination	Leverage existing in-country systems/programs/resources for COVID-19 vaccination.
	 Engage national, provincial, and district health bodies from the outset.
	 Develop district-level microplans based on standard tools that are approved at provincial and national levels.
	Use joint planning by Ministry of Health, funding organizations, and provincial representatives. The blick part 1805 the blick part 1805
	 Establish centralized M&E tools for national tracking of progress. Begin with a small pilot in a few sites and rapidly iterate to improve quality, using a continuous
	quality-improvement approach.
	Scale-up successful practices rapidly to quickly enhance effect. Payalan tages to the top have been a simple month of the property of th
	 Develop targets that can be implemented and achieved by lower levels (i.e., district health offices, service delivery teams).
Service delivery	 Adequately capacitate HCWs in HIV, MCH, and other clinics to deliver COVID-19 vaccines.
	 Invest in community mobilization and service delivery to overcome limits of a static service delivery approach and reach the greatest number of eligible persons, which means offering vaccines at publi places (e.g., markets, malls, churches), chiefdoms, workplaces, congregate settings, and others.
	 Use existing community health services for HIV as vaccination points. Anticipate additional human resource needs, and ensure adequate financial resources to support
	Afficipate additional numan resource needs, and ensure adequate infancial resources to support them.
Demand generation	 Ensure adequate HCW training in HIV and other clinics to answer patients' and eligible family members' questions about COVID-19 vaccines.
	 Encourage HCWs themselves to get vaccinated against COVID-19 by creating a safe space for unvaccinated HCWs to have their questions answered.
	 Engage public and private media nationally to address myths and misconceptions about COVID-19 vaccines.
	 Develop promotional materials that emphasize the value of COVID-19 vaccination for persons living with HIV because of the elevated risk for severe illness among members of this group.
	 Engage civil society (community, traditional, religious, and business leaders) to champion COVID-19 vaccination. Listen to and address their concerns about COVID-19 vaccines.
	 Use routine patient reminder call for upcoming visits to share information about vaccine availability i HIV clinics.
M&E	 Harmonize COVID-19 vaccine data collection in HIV and other clinics with the national COVID-19 vaccine M&E system.
	Conduct frequent data analysis to inform site-level performance assessments and guide targeted quality improvement.
	Generate feedback loops, particularly for poorly performing districts.
Logistics	 Push adequate vaccine supplies to each district based on their estimated target populations with the microplan.
	• Take inventory of health facility capacity to adequately store COVID-19 vaccines, and use existing
	infrastructure where possible.
Cofoty	Ensure that HIV clinic vaccine supply is incorporated into the wider health facility request. Provide A FFI training to HCW/s.
Safety	 Provide AEFI training to HCWs. Strengthen AEFI reporting system within HIV clinics.

Zambia could reach its targeted 70% eligible population coverage in November 2023, ahead of other countries in Africa (August 2024) (Appendix Table 7).

For Africa to reach the 2022 Africa Union targets and adequately protect the continent from subsequent COVID-19 waves, substantially accelerated COVID-19 vaccination delivery is needed (4). Moreover, rapidly reaching high vaccination coverage in Africa can help reduce the risk for emergence of new variants that can rapidly spread globally (6,7). These data suggest that strong government leadership can leverage a robust HIV program, civil society, and integrated HIV donor support from the US President's Emergency Plan for

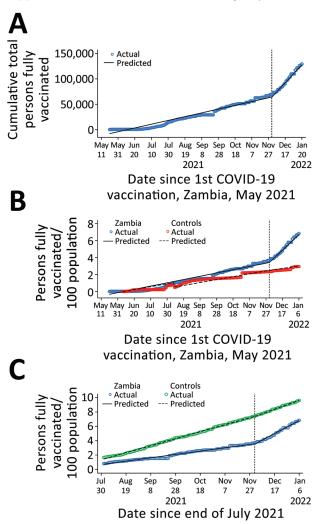


Figure. Time series of COVID-19 vaccination in Zambia, April 2021 to February 2022. A) Before and after the December Campaign. B) Compared with 2 neighboring countries with similar pre-intervention vaccination coverage trajectories and similar vaccine availability. C) Compared with the average for all 55 Africa Union member states. Prais-Winsten and Cochrane-Orcutt regression, lag(1). Vertical dashed line indicates start of Joint HIV Awareness and COVID-19 Vaccination Drive, December 1, 2021.

AIDS Relief and others to rapidly increase COVID-19 vaccine uptake. Zambia's example could hasten similar adaptations in other Africa countries.

Acknowledgment

We thank Francis D. Mwansa, Constance Sakala, Paul Chishimba, David Ngula, Sam Phiri, Liswaniso Liswaniso, Patrick Lungu, Mushota Kabaso, Lackson Kasonka, Ray W. Shiraishi,, Kayt Erdahl, Daniel Sinclair, Penelope Masumbu Kalesha, and Monjur Hossain for their work as part of the Zambia COVID-19 Vaccine Scale-Up Committee.

This work has been supported by the Zambia Ministry of Health and the US President's Emergency Plan for AIDS Relief through Centers for Disease Control and Prevention. The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the funding agencies.

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Appendix

Additional Methods

Study Design

In this interrupted time series analysis (ITSA), we analyzed publicly available data on daily COVID-19 vaccinations available from the Our World in Data (OWID) public data repository for Zambia and all 55 member states of Africa by February 21, 2022.

Data sources and participants

All participants entered into the OWID dataset by February 21, 2022, for Zambia and 55 African Union member states were eligible for the analysis. Eligibility for COVID-19 vaccinations changed over time across the 55 states of the African Union. Details of the changing vaccination policy over time can be found on the OWID Vaccination Policies webpage (https://ourworldindata.org/covid-vaccinations). A summary of the change in policy by month for three focus countries is provided in Appendix Table 1.

Intervention

The intervention of interest was the December Campaign, which started on December 1st in Zambia. Preparation for the December campaign started in August 2021, by training healthcare workers in health facilities to implement vaccinations, equipping the HIV service points at facilities and community HIV service locations, engaging community leaders through the HIV civil society network, preparing the key messages for the December campaign.

Appendix 1 above describes the multiple components of the December Campaign intervention.

Outcomes

The key outcome of interest was a complete vaccination status. OWID defines people with a completed vaccination protocol as those who have received all doses prescribed by their

vaccination regimen (e.g., 2 doses for Pfizer/BioNTech, Moderna, Oxford/AstraZeneca, etc. and 1 dose for Janssen, CanSino, etc.).

Some countries also allow for alternative definitions, such as having been infected with SARS-CoV-2 in the past and having received 1 dose of a two-dose regimen. We currently ignore these alternative definitions to preserve the common definition of a complete protocol, i.e. all doses required in the vaccine regimen. This allows for optimal comparability between countries. The types of vaccines deployed in Zambia, Tanzania and Malawi are provided in Appendix Table 2.

Data management

For Zambia and two neighboring countries, data on vaccination coverage was missing on some days during the study time periods of interest. In a sensitivity analysis, we used two approaches to the missing data: (1) carrying the most recent value for complete vaccinations forward until the next available data point, and (2) using the interpolation function in STATA to cover gaps in missing data points.

Statistical analysis

We implemented the statistical analysis in three phases. First, in a single group ITSA in Zambia only, we compared the number of people reaching full vaccination status per day before the December 1, 2021, campaign start versus after the campaign start (December 1, 2021 onwards). Secondly, in a multigroup ITSA, we assessed whether Zambia's acceleration in COVID-19 vaccination coverage (i.e., acceleration in the percentage of total population reaching full vaccination status per day) following the December Campaign intervention was statistically superior to two different control groups: (a) two neighboring countries with similar preintervention vaccination coverage trajectories and similar vaccine availability, and (b) the average for all 55 Africa Union member states. Thirdly, we implemented two sensitivity analyses for each of the above two analytic approaches by: (a) varying the approach to managming missing data (i.e., most recent value carried forward approach versus interpolation approach), and (b) comparing varying the time periods for the analysis to inform understanding of the duration of December campaign impact (see Appendix 8 below).

Ethical approval

Since the data are publicly available and de-identified, this analysis was considered non-research by authors and independent reviewers at the U.S. Centers for Disease Prevention and Control.

Additional Results

Single-Group ITSA

During December 2021, 585,677 persons in Zambia were reached for vaccination, compared with approximately 1,071,682 million during April to November 2021. Daily COVID-19 vaccinations rate accelerated by 14,070/day from 3,713/day before December to 17,783/day since December (p<0.001) (Figure 1a, Appendix 3). This statistically significant acceleration was robust to the sensitivity analysis where an interpolation approach to missing data was used instead of the approach carrying forward the most recent available data point (Appendix 3). In addition, the average post-December daily vaccination rate only dropped slightly, from 17,783/day for December 1 – January 6, 2022 to 13,911/day for December 1 – February 20, 2022, indicating a sustained impact for nearly 3 months after December Campaign intervention launch (Appendix 3).

Multi-Group ITSA

In the multi-group ITSA, compared with the two neighboring countries, Zambia accelerated its population COVID-19 vaccine coverage rate by an additional 2.73% coverage/month (p<0.001) (Figure 1b, Appendix 4). Compared with the average daily increase in population coverage for Africa, Zambia accelerated 1.87% coverage/month (p<0.001) (Figure 1c, Appendix 5). This statistically significant acceleration was robust to the sensitivity analysis where an interpolation approach to missing data was used instead of the approach carrying forward the most recent available data point (Appendix 6). In addition, the increased average post-December percentage gain per day in population fully vaccinated remained relatively stable through February 20, 2022, indicating a sustained impact for nearly 3 months after December Campaign intervention launch (Appendix 6).

Limitations

This analysis has several strengths and limitations. Strengths include the availability of daily trends in vaccination numbers and coverage, and clear timing of the intervention launch (December 1, 2021), making the interpretation of observed changes in vaccination trends as largely due to December Campaign intervention impact plausible. Limitations include the fact that the analysis relies on routinely collected program data, the analysis design is quasi-experimental which increases risk of confounding, and, given the nature of real-world program implementation, it is possible that some of the observed acceleration in vaccination coverage in Zambia occurred because of increased engagement through non-HIV service delivery points and through processes unrelated to the December Campaign.

Appendix Table 1. Vaccination policies in three focus countries for the study time period of interest*

Period	Zambia	Neighboring country 1	Neighboring country 2
March 1, 2021	No policy	No policy	No policy
April 1, 2021	No policy	All vulnerable groups	No policy
May 1, 2021	One group	Universal	No policy
June 1, 2021	All vulnerable groups	Universal	No policy
July 1, 2021	All vulnerable groups	Universal	No policy
August 1, 2021	All vulnerable groups	Universal	All vulnerable groups
Sept 1, 2021	All vulnerable groups and some others	Universal	All vulnerable groups and some
			others
Oct 1, 2021	All vulnerable groups and some others	Universal	Universal
Nov 1, 2021	All vulnerable groups and some others	Universal	Universal
Dec 1, 2021	All vulnerable groups and some others	Universal	Universal
Jan 1, 2022	All vulnerable groups and some others	Universal	Universal
Feb 1, 2021	All vulnerable groups and some others	Universal	Universal

^{*}Vaccination policies are divided into 6 groups as follows: (1) No availability; (2) Availability for ONE of following: key workers/ clinically vulnerable groups / elderly groups; (3) Availability for TWO of following: key workers/ clinically vulnerable groups / elderly groups; (4) Availability for ALL of following: key workers/ clinically vulnerable groups / elderly groups; (5) Availability for all three plus partial additional availability (select broad groups/ages); (6) Universal availability. Given that the vaccination policy for all 55 member states varied significantly across countries and over time, we have not attempted to summarize the data here, but these data are available at OWID.

Appendix Table 2. Vaccinations deployed in Zambia and two neighboring countries*2

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COVID-19 vaccines			
deployed in focus countries	Zambia	Neighboring country 1	Neighboring country 2
Vaccines	Johnson&Johnson,	Johnson&Johnson,	Johnson&Johnson,
	Oxford/AstraZeneca,	Oxford/AstraZeneca	Pfizer/BioNTech,
	Sinopharm/Beijing		Sinopharm/Beijing

^{*}Data from Our World in Data. Country-specific types of COVID-19 vaccines by February 21, 2022. (https://ourworldindata.org/covid-vaccinations).

Appendix Table 3. Single group interrupted time series analysis for Zambia, comparing pre-December with December 1, 2021-

onward vaccination trends

	Pre-De	cember Interv	ention		Post-December 1 Intervention							
	(daily ทเ	ımber of peop	le fully	Impact on daily numb	Impact on daily number of complete vaccinations				(daily number of people fully			
	vaccinated)*			of Decem	vaccinated)							
			p-	Additional								
Analysis	Trend	95% CI	value	vaccinations/day	95% CI	p-value	Trend	95% CI	p-value			
Primary Ar	alysis†											
Zambia	3,713	(2,995-	<0.001	14,070	(9,654-18,486)	< 0.001	17,783	(13,640-	< 0.001			
		4,431)						21,927)				
Sensitivity	Analysis 1:	‡										
Zambia	3,602	(3,219-	< 0.001	12,807	(9,939-15,676)	< 0.001	16,410	(13,586-	< 0.001			
		3,986)			,			19,234)				
Sensitivity	Analysis 2	§										
Zambia	3,610	(3,221-	<0.001	10,301	(8,532-12,070)	< 0.001	13,911	(12,212-	< 0.001			
		3,999)						15,610)				

^{*}Pre-December time-period: May 20, 2021 - November 30, 2021.

Appendix Table 4. Multi-group interrupted time series analysis, comparing pre-December with December 1, 2021-onward vaccination trends in Zambia with two neighboring countries

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Analysis	Coefficient	95% CI	p-value						
Pre-December trend comparisons to assess suitability of neighboring countries as controls*									
Pre-December mean level difference in cumulative % of population vaccinated	-0.31	(-0.88-0.25)	0.279†						
Pre-December difference in slope (i.e., difference in daily change in % of population vaccinated /day)	0.00	(0-0.01)	0.399†						
December intervention impact on speed of daily vaccine rollout									
Acceleration in % of population vaccinated/day in Zambia compared with average for controls	0.08	(0.04-0.11)	<0.001						
Absolute differences in speed of post-December vaccine rollout between Zambia and Average of controls									
Zambia`s Post-December average % of population vaccinated per day‡	0.09	(0.07-0.12)	< 0.001						
Neighboring Countries (i.e., Controls) Post-December 1, 2021, average % of population vaccinated per day	0.01	(0-0.03)	0.026						
Absolute Difference between Post-December 1, 2021, average % of population vaccinated per day	0.08	(0.05-0.11)	<0.001						

^{*}Pre-December time period: May 20, 2021 - November 30, 2021.

Appendix Table 5. Results of multi-group interrupted time series analysis, comparing pre-December with December 1, 2021onward vaccination trends in Zambia with the average for Africa

Analysis	Coefficient	95% CI	p-value
Pre-December trend comparisons to assess suitability of Africa average as a control*†			
Pre-December mean level difference in cumulative % of population vaccinated	-0.71	(-0.860.57)	<0.001
Pre-December difference in slope (i.e., difference in daily change in % of population	-0.03	(-0.030.02)	< 0.001
vaccinated /day)			
December intervention impact on speed of daily vaccine rollout			
Acceleration in % of population vaccinated/day in Zambia compared with average for	0.05	(0.04-0.07)	<0.001
control group			
Absolute differences in speed of post-December vaccine rollout between Zambia and			
Africa average			
Zambia`s Post-December average % of population vaccinated per day‡	0.09	(0.08-0.1)	<0.001
Africa's Post-December 1, 2021, average % of population vaccinated per day	0.06	(0.06-0.07)	< 0.001
Absolute Difference between Post-December average % of population vaccinated per	0.03	(0.02 - 0.04)	<0.001
day between Zambia and the average for Africa			

^{*}Pre-December time period: July 30, 2021 - November 30, 2021.

[†]Primary analysis: Missing data approach: Most recent value carried forward. Post-intervention time-period: December 1- January 6, 2022.

[‡]Sensitivity analysis 1: Missing data approach: Interpolation. Post-intervention time-period: December 1- January 6, 2022.

[§]Sensitivity analysis 2: Missing data approach: Interpolation. Post-intervention time-period: December 1- February 20, 2022.

[†]Because difference in mean baseline levels of vaccination coverage and speed or slope of percentage vaccinated per day, control group is appropriate statistically.

[‡]Post-December time period: December 1, 2021 – January 6, 2022.

[†]Baseline levels and slopes are statistically different pre-December (i.e., Africa average is faster than Zambia), making the Africa average a less suitable control group for Zambia. However, because the pre-intervention vaccination coverage slope is higher in Africa overall, this makes it less likely that the acceleration in Zambia would be observed to be statistically significant (i.e., the approach taken is conservative). The Africa average is taken directly from Our World in Data and includes Zambia, making the comparison with Zambia more conservative (i.e., less likely to detect a significant difference).

[‡]Post-December time period: December 1, 2021 – January 6, 2022

Appendix Table 6. Multi-group sensitivity analysis to assess robustness of primary estimates of accelerated full vaccination coverage depending on (1) approach to missing data management and (2) time period included in the analysis after December 1, 2021 vaccination campaign launched

-	Prir	nary Analysis	*	Sensi	tivity Analysis	; 1†	Sensitivity Analysis 2‡		
Robustness	Coefficient	95% CI	p-value	Coefficient	95% CI	p-value	Coefficient	95% CI	p-value
Acceleration in % of population vaccinated/day in Zambia compared with average for controls (two neighboring countries)	0.08	(0.04-0.11)	<0.001	0.06	(0.04-0.07)	<0.001	0.05	(0.04-0.06)	<0.001
Acceleration in % of population vaccinated/day in Zambia compared with average for control group (Average for Africa)	0.05	(0.04-0.07)	<0.001	0.05	(0.04-0.06)	<0.001	0.04	(0.03-0.05)	<0.001

^{*}Primary analysis: Missing data approach: Most recent value carried forward. Post-intervention time-period: December 1- January 6, 2022. †Sensitivity analysis 1: Missing data approach: Interpolation. Post-intervention time-period: December 1- January 6, 2022. ††Sensitivity analysis 2: Missing data approach: Interpolation. Post-intervention time-period: December 1- February 20, 2022.

Appendix Table 7. Estimated time taken to reach 70% population coverage with COVID-19 vaccination

	Additional % of										
	population fully	Additional %	Baseline								
	vaccinated per	of population	coverage	Target		Currently		Numbers fully	Days	Months	
	Day post	fully	population	coverage		vaccinated	Target (i.e.	vaccinated per	needed to	needed to	Date 70%
	December 1,	vaccinated	at January	of	Population	(calculated	70% of total	day at current	reach 70%	reach 70%	target is
Country	2021*	per Month†	6, 2022‡	population	size§	estimate)¶	population)	rates#	target**	target	reached
Zambia	0.09	2.73	6.84%	70%	18,384,000	1,257,466	12,868,800	16,734	694	23.13	30-Nov-23
Two	0.01	0.42	2.94%	70%	78,864,000	2,314,658	55,204,800	11,124	4,754	158.48	12-Jan-35
neighboring											
countries											
(average)											
Africa	0.06	1.87	9.62%	70%	1,370,700,000	131,861,340	959,490,000	856,470	966	32.21	29-Aug-24

^{*}Taken from appendices 3 and 4 above.

[†]Additional % of population fully vaccinated per Month = Additional % of population fully vaccinated per day*30

[‡]From our world in data: https://ourworldindata.org/covid-vaccinations. §Population sizes for Zambia and Africa for 2021 from https://www.statista.com/statistics/1224168/total-population-of-africa.

[#]Calculated as baseline coverage population at January 6, 2022 * population size.

#Calculated as post-December additional % of population vaccinated per day multiplied by population size.

**Calculated as the total number of people that still need to be reached with full vaccination services divided by the number vaccinated per day post-December.