Mpox severity and associated hospitalizations among people with HIV and related immunosuppression in Brazil

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Objectives: This study aimed to analyze characteristics of mpox hospitalization in a Brazilian cohort, further exploring the impact of HIV on mpox-related outcomes and hospitalization.

Design: We conducted a descriptive analysis, comparing characteristics of individuals diagnosed with mpox according to hospitalization and HIV status, and described the mpox cases among those living with HIV.

Methods: This was a single-center, prospective cohort study conducted at a major infectious diseases referral center in Rio de Janeiro, Brazil, that enrolled participants older than 18 years of age diagnosed with mpox. Information was collected on standardized forms, including data on sociodemographic, behavioral, clinical and laboratory characteristics. For comparisons, we used chi-squared, Fisher's exact and the Moods median tests whenever appropriate.

Results: From June to December, 2022, we enrolled 418 individuals diagnosed with mpox, of whom 52% were people with HIV (PWH). PWH presented more frequently with fever, anogenital lesions and proctitis. The overall hospitalization rate was 10.5% (n = 43), especially for pain control. Among hospitalized participants, PWH had more proctitis and required invasive support. Mpox severity was related to poor HIV continuum of care outcomes and low CD4⁺ cell counts. All deaths (n = 2) occurred in PWH with CD4⁺ less than 50 cells/µl.

Conclusion: HIV-related immunosuppression likely impacts mpox clinical outcomes. This is of special concern in settings of poor adherence and late presentation to care related to socioeconomic inequalities, such as Brazil. The HIV continuum of care must be taken into account when responding to the mpox outbreak.

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Background

The 2022 multinational mpox outbreak in nonendemic countries led to its declaration as a health emergency in July 2022, reaching the global number of 90 618 cases and 157 deaths by 27 September 2023, with Brazil being the second most affected country [1,2]. Cisgender MSM were disproportionately burdened by this outbreak; 30–50% of individuals diagnosed with mpox were living with HIV [3–11]. Generally, people with HIV (PWH) and mpox were older and more often presented with concomitant sexually transmitted infections (STIs), anogenital lesions and proctitis [4,8,9,12]. Although initial mpox descriptions reported no substantial differences in hospitalizations or clinical severity according to HIV status, these individuals were mostly virologically stable and without severe, immunosuppression [3,4,7,10,13].

Recent studies, however, suggest that uncontrolled HIV might impact on mpox-related outcomes, especially if $CD4^+$ cell counts are below 200 cells/µl, suggesting an association between advanced HIV disease and severe mpox presentation [5,8,9,14-17]. This is of special concern in Latin America - the region accounted for 55% of PWH with mpox and CD4⁺ counts lower than $350 \text{ cells}/\mu$; access to prevention and experimental mpox treatment were quite limited [15,18]. Overall, mpoxrelated hospitalization rates during the 2022 multinational outbreak are up to 10%, mostly related to pain control, secondary bacterial infections, urologic or proctologic complications, odynophagia, keratitis and, rarely, encephalitis, pulmonary involvement or myocarditis [4,5,7,15,17,19–22]. Brazil reported up to 50% of mpox cases and at least three deaths among PWH [4,23]. Despite the country's longstanding commitment to universal access to antiretroviral therapy (ART), structural barriers remain, resulting in a high prevalence of HIV late presentation to care and poor ART adherence. These aspects have worsened after the COVID-19 pandemic and contributed to the high mpox burden [24-26].

This study aimed to characterize mpox-related hospitalizations and further explore the impact of HIV status on mpox outcomes among individuals enrolled at the Evandro Chagas National Institute of Infectious Diseases (INI/Fiocruz), a major referral center for mpox in Rio de Janeiro, Brazil.

Methods

Study design and participants

This was a single-center, prospective cohort study that enrolled participants older than 18 years diagnosed with mpox from 12 June to 31 December 2022, at INI/ Fiocruz. We offered mpox virus quantitative real-time PCR (qPCR) testing to all individuals with suspected infection, which was performed at the Mpox Reference Laboratory at the Oswaldo Cruz Institute/Fiocruz. A positive result on qPCR from any swab specimens led to case confirmation. Follow-up period was 28 days or until resolution of mpox lesions. Individuals requiring hospitalization were followed at the INI-Fiocruz Inpatient Care Unit; if hospitalization occurred elsewhere, data were collected retrospectively. Procedures of INI's mpox cohort have been previously described [4].

Procedures

We prospectively collected sociodemographic, epidemiological, behavior, clinical and laboratory data onto a standardized case report form. Information was gathered on birth date, gender identity and race according to selfreport. In Brazil, smallpox vaccination was compulsory until 1975, so for these analyses we considered those born before 1975 as vaccinated. Systemic signs and symptoms included the presence of fever, asthenia, adenomegaly, myalgia, arthralgia and/or headache. For hospitalized participants, invasive support included mechanical ventilatory support, vasoactive drugs, use of nasoenteric tube or indwelling urinary catheters, need for hemodialysis or either arterial or venous central line. We included data on reasons for hospitalization, length of hospitalization and clinical outcomes. Tecovirimat was requested according to the Ministry of Health's protocol for compassionate use, upon availability, if the individual met criteria of a severe clinical course [27].

At the initial assessment, we offered HIV rapid test (3rd generation) performed according to the Brazilian Ministry of Health algorithm [28]; HIV-RNA viral load; rapid *Treponema pallidum* test for syphilis screening with subsequent confirmation with nontreponemal testing [veneral disease research laboratory (VDRL)]; molecular detection of Chlamydia trachomatis and Neisseria gonorrheae in rectal swabs (Abbott Real Time Platform); hepatitis B surface antigen rapid test; anti-HCV rapid test for hepatitis C. Active syphilis was defined as VDRL titers equal to or higher than 1:8.

For PWH, we assessed CD4⁺ count (cells/µl) and HIV-RNA viral loads records closest to the onset of mpox symptoms, obtained from either clinical charts or the Brazilian Laboratory Control System database (SISCEL). We adapted a cross-sectional cascade of HIV care including the following stages: HIV-diagnosed (previously known HIV infection at mpox diagnosis); linked to HIV care (at least one record of HIV-related care appointment, laboratory examinations or ART prescription after HIV diagnosis and before mpox assessment); retained in HIV care (ART prescription in the last 6 months or at least two records of HIV-RNA viral load or CD4⁺ cell count in the last year); on ART (at least one dispensation of antiretroviral drugs in the last 6 months, obtained from the Brazilian Antiretroviral Logistics database - SICLOM); and virologically suppressed (HIV-RNA viral load ≤ 200 copies/ml) [29].

Statistical analysis

We compared study characteristics of confirmed mpox cases according to HIV status and, among PWH, by hospitalization status (PWH hospitalized vs. PWH not hospitalized). We used the chi-squared test or Fisher's exact test for qualitative variables and the Moods median test for quantitative variables. We performed a subanalysis among PWH to compare clinical outcomes according to $CD4^+$ count strata and to describe HIV continuum of care outcomes by hospitalization status. All analyses were performed in R-Project (4.2.1).

Ethical considerations

This study was approved by the Ethics Review Board at INI-Fiocruz (CAAE #61290422.0.0000.5262). Participants provided written informed consent.

Results

From June 12 to 31 December 2022, 418 participants had a confirmed mpox diagnosis. Overall, median age was 33 years [interquartile range (IQR) 28–40]; 91.6% selfidentified as cisgender men, 5.7% as cisgender women and 2.6% as travesti or transgender women (TGW). The majority were black or pardo (60.5%, n=207/342), 38.6% were white (n=132/342) and 0.9% were indigenous (n=3/342). Most participants had studied until primary education (60.2%, n=213/352), 6% had secondary education (n=119/352). Among the cisgender men, 87% were MSM (n=336/362).

Data on HIV serostatus were available for 409 participants, of whom 52.1% were PWH, mostly with previous HIV diagnosis. Compared with HIV-negative individuals, PWH diagnosed with mpox were older [35 (IQR 30-40) vs. 31 (IQR 26-38), P < 0.01] and most frequently self-reported being cisgender men (95.8 vs. 87.3%, P < 0.01), travesti or TGW (3.3 vs. 2%, P < 0.01) (Table 1). Among those who were negative for HIV, PrEP use was reported by 33% (Table 1).

Compared with HIV-negative participants, more PWH reported being MSM (98.4 vs. 87.3%, P < 0.01); had active syphilis (29.8 vs. 10%, P < 0.01), hepatitis C (9.8 vs. 2.7%, P < 0.1) or any bacterial STI (42.4 vs. 21.7%, P < 0.01) (Table 1); presented more frequently with fever (71.1 vs. 55.7%, P < 0.01), anogenital lesions (82.1 vs. 74.1%, P = 0.05) and proctitis (29.7 vs. 19.1%, P = 0.01) and more often had a positive MPXV qPCR in rectal swabs (80.37 vs. 66.12%, P = 0.01).

Overall, 10.5% (n = 43) of participants were hospitalized because of mpox (39 at INI-Fiocruz Inpatient Care Unit), 55.8% were PWH, with a median age of 31 years old (IQR 28-39); 93% were cisgender men, 50% selfidentified as black/pardo and 53.5% had primary education. The hospitalization rate during follow-up was not different based on HIV status although 34.9% had been admitted at the time of the initial mpox assessment. No participants reported immunosuppression related to drugs or other medical conditions. Median interval between symptoms onset and hospitalization was 9 days (IQR 7–12) and length of hospitalization was 7.5 days (IQR 5–12). The most common reason for hospitalization was pain control (90.7%), often requiring opioids use (67.5%), followed by secondary bacterial infections (55.8%). The majority of hospitalized participants presented with fever (69.1%) and/or anogenital lesions (67.4%). PWH presented more frequently with proctitis [58.33 vs. 26.32%, P=0.04)] and required invasive support (34.8 vs. 0, P=0.01) (Table 2).

When comparing to PWH and mpox who have not required inpatient care (n=189), those hospitalized (n=24) reported less frequent sex contact one month prior to symptoms onset (66.7 vs. 91.5%, P < 0.01) and presented worse outcomes related to the HIV care continuum, with poorer rates of retention (79.2 vs. 94.7%, P=0.01), regular ART use (79.2 vs. 94.1%, P=0.02) and virological suppression (66.7 vs. 87.7%, P=0.01), in addition to showing lower CD4⁺ cell counts [median 448 (IQR 174-850) vs. 630 (IQR 487-889), P=0.02] and less frequently presenting CD4⁺ counts higher than 350 cells/µl (58.3 vs. 91.8%, P < 0.01) (Table 3 and Fig. 1).

Among the 24 PWH hospitalized because of mpox, 11 had detectable HIV-RNA viral load (higher than 40 copies/ml), all of them presented mucosal involvement, frequently requiring intravenous antibiotics (n=8), analysics (n=10) or opioids (n=8). Among them, eight had $CD4^+$ cell counts below 200 cells/µl and most frequently experienced worse mpox-related outcomes, including hospitalization and death (Fig. 2). Concomitant opportunistic infections were identified: tuberculosis (n = 4), histoplasmosis (n = 1), cytomegalovirus infection (n = 1), oropharyngeal candidiasis (n = 1), Kaposi's sarcoma (n = 1). All participants with deep tissue involvement had CD4⁺ counts below 50 cells/ μ l (n = 4). Two participants, both with CD4⁺ counts below 50 cells/ µl, had respiratory complications [pleural effusion (n = 1); pulmonary nodules (n = 1)]. All deaths occurred among PWH (n=2), all of them with poor ART adherence and $CD4^+$ counts below 50 cells/µl. Both developed bowel and urologic obstruction, sepsis and refractory shock, requiring mechanical ventilation and use of vasoactive drugs.

Discussion

Our findings contribute to further characterize the mpox outbreak in Latin America, focusing on individuals living with HIV with different levels of immunosuppression.

Table 1.	Sociodemographic,	behavioral and o	linical	characteristics of mpo	x cases according	g to HIV	status (N =	= 409).
					(,		

	HIV- $(N = 196)^{a}$	$HIV + (N = 213)^{a}$	Total $(N = 409)^{a}$	
	[n (%)]	[<i>n</i> (%)]	[n (%)]	P value ^b
Age (median, IQR)	31 (26–38)	35 (30-40)	33 (28-40)	< 0.01
Age range				< 0.01
18–24 years	34/196 (17.4%)	12/213 (5.6%)	46/409 (11.2%)	
25–29 years	49/196 (25%)	33/213 (15.5%)	82/409 (20.1%)	
30–39 years	71/196 (36.2%)	105/213 (49.3%)	176/409 (43%)	
>40 years	42/196 (21.4%)	63/213 (29.6%)	105/409 (25.7%)	
Gender identity			. ,	< 0.01
Cisgender men	171/196 (87.3%)	204/213 (95.8%)	375/409 (91.7%)	
Cisgender women	21/196 (10.7%)	2/213 (0.9%)	23/409 (5.6%)	
Nonbinary	0/196	0/213	0/409	
Transgender men	0/196	0/213	0/409	
Travesti or transgender women	4/196 (2%)	7/213 (3.3%)	11/409 (2.7%)	
Race				0.65
Black	53/166 (31.9%)	52/175 (29.7%)	105/341 (30.8%)	
Pardo or mixed	45/166 (27.1%)	57/175 (32.6%)	102/341 (29.9%)	
White	67/166 (40.4%)	64/175 (36.6%)	131/341 (38.4%)	
Indigenous	1/166 (0.6%)	2/175 (1.1%)	3/341 (0.9%)	
Education				0.19
Primary	110/169 (65.1%)	100/180 (55.6%)	210/349 (60.2%)	
Secondary	9/169 (5.3%)	12/180 (6.6%)	21/349 (6%)	
Post secondary	50/169 (29.6%)	68/180 (37.8%)	118/349 (33.8%)	
MSM	144/165 (87.3%)	190/193 (98.4%)	334/358 (93.3%)	< 0.01
Current sex work	6/178 (3.4%)	6/190 (3.2%)	12/368 (3.3%)	0.91
PrEP use	64/194 (32.9%)	NA	NA	
Reported sex in the last 30 days	175/186 (94.1%)	175/197 (88.8%)	350/383 (91.4%)	0.07
Time from symptoms onset to initial	6 (4, 9)	6.0 (4, 10)	6.0 (4, 9)	0.52
assessment (median, IQR)	0 (1) 0)		010 (1) 5)	0.02
Median number of sex partners in the last 30 days (IQR)	2 (1, 3)	2 (1, 3)	2 (1, 3)	0.54
Reported anal sex in the last 30 days ^c	78/175 (44.6%)	88/175 (50.3%)	166/350 (47.4%)	0.28
Sex contact with potential mpox case ^d	35/171 (20.5%)	35/173 (20.2%)	70/344 (20.3%)	0.58
Home contact with potential mpox cases ^e	20/169 (11.9%)	12/172 (7%)	32/341 (9.4%)	0.12
Probable vaccinated for smallpox	18/196 (9.2%)	19/213 (8.9%)	37/409 (9%)	0.97
Active syphilis	18/180 (10%)	59/198 (29.8%)	77/383 (20.1%)	< 0.01
Anorectal gonorrhea	11/141 (7.8%)	15/164 (9.15%)	26/305 (8.52%)	0.69
Anorectal Chlamydia	12/141 (8.5%)	18/164 (11%)	30/305 (9.8%)	0.47
Any bacterial STI	39/180 (21.7%)	84/198 (42.4%)	123/378 (32.5%)	< 0.01
Hepatitis B	2/181 (1.1%)	3/189 (1.6%)	5/370 (1.4%)	>0.99
Hepatitis C	5/187 (2.7%)	19/194 (9.8%)	23/381 (6%)	< 0.01
Systemic symptoms or signs	157/186 (75.3%)	187/207 (84.1%)	344/393 (79.8%)	0.07
Fever	107/192 (55.7%)	150/211 (71.1%)	257/403 (63.7%)	< 0.01
Anogenital lesions	143/193 (74.1%)	170/207 (82.1%)	313/400 (78.3%)	0.05
Clinical signs of proctitis	37/194 (19.1%)	63/212 (29.7%)	100/406 (24.6%)	0.01
Rectal swab				0.01
PCR MPXV detectable	80/121 (66.1%)	131/163 (80.4%)	211/284 (75.3%)	
PCR MPXV not detectable	41/121 (33.9%)	32/163 (19.6%)	73/284 (24.7%)	
Required hospitalization	19/196 (9.7%)	24/213 (11.3%)	43/409 (10.5%)	0.60
Death	0/196	2/213 (1%)	2/409 (0.5%)	0.50

IQR, interquartile range; PrEP, pre-exposure prophylaxis.

^an (%); median (IQR).

^bFisher's exact test; Wilcoxon rank sum test; Pearson's chi-squared test.

^cBoth receptive and insertive anal sex.

^dReported sexual contact with someone who was suspect case or diagnosis with mpox or presenting mpox-like lesions in the 30 days previous the study entry.

^eReported living in the same household of suspected/confirmed mpox case in the 30 days previous the study entry.

HIV prevalence was in line with global data [12]. In our study, mpox individuals showed a high frequency of a concomitant STI and hepatitis C virus (HCV). A recent systematic review showed an association between syphilis coinfection and HIV among individuals diagnosed with mpox, despite a significant amount of missing data due to variations in access to STI testing across different settings [12,30]. Our elevated syphilis prevalence contrasts with international data, including other Latin American countries but is compatible with the alarming epidemiological syphilis scenario in Brazil [5,8,10,31]. Such findings might be related to shared sexual networks or higher risk of HIV acquisition in the context of a concomitant STI, and underscores the importance of mpox assessment as an opportunity to expand other STI diagnosis [32,33].

	$HIV - (N = 19)^{a}$	$HIV + (N = 24)^{a}$	Total $(N=43)^a$	P value ^b
Median age (IQR)	30 (25-36)	35 (30-42)	31 (28-40)	0.08
Age range				0.20
18–24 years	3/19 (15.8%)	1/24 (4.2%)	4/43 (9.3%)	
25–29 years	6/19 (31.6%)	3/24 (12.5%)	9/43 (20.9%)	
30–39 years	7/19 (36.8%)	12/24 (50%)	19/43 (44.2%)	
\geq 40 years	3/19 (15.8%)	8/24 (33.3%)	11/43 (25.6%)	
Gender identity				0.08
Cisgender men	16/19 (84.2%)	24/24 (100%)	40/43 (93%)	
Cisgender women	3/19 (15.8%)	0/24	3/43 (7%)	
Race				>0.99
White	7/18 (38.9%)	9/21 (42.9%)	16/39 (41.1%)	
Pardo or mixed	5/18 (27.8%)	5/21 (23.8%)	10/39 (25.6%)	
Black	6/18 (33.3%)	7/21 (33.3%)	13/39 (33.3%)	
Education				0.13
Primary	13/19 (68.4%)	10/24 (41.7%)	23/43 (53.5%)	
Secondary	1/19 (5.3%)	1/24 (4.1%)	2/43 (4.6%)	
Postsecondary	5/19 (26.3%)	13/24 (54.2%)	18/43 (41.9%)	
Fever	14/19 (73.7%)	15/23 (65.2%)	29/42 (69%)	0.55
Anogenital lesions	9/19 (47.4%)	20/24 (83.3%)	29/43 (67.4%)	0.01
Time from symptoms onset to hospitalization (days, IQR)	9 (8-11)	8 (7-14)	9 (7-12)	>0.99
Length of hospitalization (days, IQR)	7 (5-10)	8 (4-14)	7.5 (5-12)	0.56
Reason of hospitalization				
Pain control	18/19 (94.7%)	21/24 (87.5%)	39/43 (90.7%)	0.62
Urological complication	7/19 (36.8%)	10/24 (41.7%)	16/43 (37.21%)	0.75
Proctitis	5/19 (26.3%)	14/24 (58.3%)	18/43 (41.9%)	0.04
Superbacterial infection	11/19 (57.9%)	13/24 (54.2%)	24/43 (55.8%)	0.81
Ophthalmologic complications	2/19 (10.5%)	1/24 (4.2%)	3/43 (7%)	0.58
Neuropsychiatric symptoms	4/19 (21.1%)	5/24 (20.8%)	9/43 (20.9%)	>0.99
Odynophagia	6/19 (31.6%)	5/24 (20.8%)	11/43 (25.6%)	0.49
Need for invasive support ^c	0/19	8/23 (34.8%)	8/42 (19.1%)	0.01
Urological or bowel obstruction	1/19 (5.3%)	3/23 (13%)	4/42 (9.5%)	0.61
Use of opioids for pain control	14/17 (82.4%)	13/23 (56.5%)	27/40 (67.5%)	0.09
Death	0/19	2/24 (8.3%)	2/43 (4.7%)	0.50

Fable 2. Sociodemographic, laboratoria	l, and clinical information on mp	ox-related hospitalized	participants according t	o HIV status ($N = 43$)
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IQR, interquartile range.

^an (%); median (IQR).

^bFisher's exact test; Wilcoxon rank sum test; Pearson's chi-squared test.

^cInvasive support included need for mechanical ventilation, vasoactive drugs, hemodialysis, venous or arterial lines and/or vesical catheterization.

Despite presenting more often with fever, anogenital lesions and proctitis, PWH in our cohort showed similar proportions of hospitalization as individuals not living with HIV, which is still controversial in available global data [5,9,10,14]. Reasons for admission were quite similar to other cohorts, being mainly associated with pain control, bacterial superinfection, or management of urologic and proctological complications [17,34]. Evidence shows that, among PWH, those with CD4⁺ counts lower than 200 cells/µl or unsuppressed HIV-RNA viral load are more prone to developing mpox-related complications, including hospitalization, intensive care support and death [15-17]. Mpox-related complications in individuals with CD4⁺ cell counts below 200 cells/µl can lead to fulminant clinical course coalescing necrotizing skin and soft tissue lesions, as well as pulmonary involvement [15]. There is still a gap regarding the best practices related to time for ART initiation or resumption, given the risk of Immune Reconstitution Inflammatory Syndrome, as well as the duration of mpoxspecific antiviral treatment [15,16]. Special attention must be given to potential concurrent opportunistic infections [15].

The global health response to mpox is marked by inequities related to a focus on Northern countries and scarce access

to preventive and treatment alternatives in Latin America and Africa. Moreover, mpox's description as an HIVrelated opportunistic infection and its emergence as a global threat underscores the concern regarding the poorer HIV cascade of care outcomes [15,35]. Our findings suggest that PWH with mpox requiring hospitalization performed poorly in HIV continuum of care when compared with those followed only at an outpatient unit, with significantly lower ART adherence. Likewise, a recent report on mpox-related deaths in the United States unveiled structural inequities mostly affecting black and people experiencing homelessness [16]; most of whom had a late HIV diagnosis, and no previous ART use, as opposed to our scenario [16]. Notably, in our cohort, the most concerning outcomes were concentrated in the distal end of the HIV continuum of care, suggesting that poor ART adherence plays a greater role than late diagnosis.

In Brazil, access to HIV prevention and treatment is guaranteed by the Unified Health System (SUS), reinforcing its principles of equity, universality and integrality [36], and allowing consistent improvement in HIV-related outcomes. By 2021, Brazil had made steady progress in achieving UNAIDS 95–95–95 goals, with an

	Not hospitalized, $(N = 189)^{a}$	Hospitalized $(N = 24)^a$	Total ^a ($N = 213$)	P value ^b
Median age (IOR)	35 (30-40)	35 (30-42)	35 (30-40)	0.75
Age range				0.83
18–24 years	11/189 (5.8%)	1/24 (4.2%)	12/213 (5.6%)	
25–29 years	30/189 (15.9%)	3/24 (12.5%)	33/213 (15.5%)	
30–39 years	93/189 (49.2%)	12/24 (50%)	105/213 (49.3%)	
>40 years	55/189 (29.1%)	8/24 (33.3%)	63/213 (29.6%)	
Gender identity				>0.99
Cisgender men	180/189 (95.2%)	24/24 (100%)	204/213 (95.8%)	
Cisgender women	2/189 (1.1%)	0/24	2/213 (0.9%)	
Travesti or TGW	7/189 (3.7%)	0/24	7/213 (3.3%)	
Race				0.75
White	55/154 (35.7%)	9/21 (42.9%)	64/175 (36.6%)	
Indigenous	2/154 (1.3%)	0/21	2/175 (1.1%)	
Pardo or mixed	52/154 (33.8%)	5/21 (23.8%)	57/175 (32.6%)	
Black	45/154 (29.2%)	7/21 (33.3%)	52/175 (29.7%)	
Education				0.22
Primary	90/156 (57.7%)	10/24 (41.7%)	100/180 (55%)	
Secondary	11/156 (7.1%)	1/24 (4.2%)	12/180 (7.6%)	
Postsecondary	55/156 (35.3%)	13/24 (54.2%)	68/180 (38%)	
Reported sexual contact 30 days	161/176 (91.5%)	14/21 (66.7%)	175/197 (88.8%)	< 0.01
before symptoms onset				
Probably vaccinated for smallpox	15/189 (7.9%)	4/24 (16.7%)	19/213 (8.9%)	0.12
HIV-RNA viral load				0.04
<40 copies/ml	143/188 (76.1%)	13/24 (54.2%)	156/212 (73.6%)	
41–200 copies/ml	21/188 (11.2%)	3/24 (12.5%)	24/212 (11.3%)	
201-1000 copies/ml	6/188 (3.2%)	2/24 (8.3%)	8/212 (3.8%)	
>1000 copies/ml	18/188 (9.5%)	6/24 (25%)	24/212 (11.3%)	
Median CD4 ⁺ (cells/µl, IQR)	630 (487, 889)	448 (174, 850)	624 (463, 889)	0.02
CD4 ⁺ range (cells/µl)				< 0.01
≤50	1/170 (0.6%)	4/24 (16.7%)	5/194 (2.6%)	
51-100	0/170	1/24 (4.2%)	1/194 (0.5%)	
101-200	2/170 (1.2%)	3/24 (12.5%)	5/194 (2.6%)	
201-350	11/170 (6.5%)	2/24 (8.3%)	13/194 (6.7%)	
>350	156/170 (91.7%)	14/24 (58.3%)	170/194 (87.6%)	
ART status				
Never initiated	5/187 (2.7%)	1/24 (4.2%)	6/211 (2.8%)	0.02
On ART, regular adherence ^c	166/187 (88.8%)	17/24 (70.8%)	183/211 (86.8%)	
On ART, irregular adherence ^c	13/187 (6.9%)	3/24 (12.5%)	16/211 (7.6%)	
No ART in last 6 months ^c	3/187 (1.6%)	3/24 (12.5%)	6/211 (2.8%)	

Table 3. Sociodemographic, behavioral and clinical characteristics of mpox participants coinfected with HIV according to hospitalization during mpox follow-up (N = 213).

ART, antiretroviral therapy; IQR, interquartile range.

^an (%); median (IQR).

^bFisher's exact test; Wilcoxon rank sum test; Pearson's chi-squared test.

^cSelf-reported information on ART adherence referring to the 6-month period prior to mpox first assessment. Regular ART adherence was considered if the participant reports taking at least five pills per week in the recall time.

estimate of 89% PWH diagnosed, of whom 82% were on regular ART and, among those, 95% were virologically suppressed [29]. Nevertheless, the country faces a concentrated HIV epidemic, mirroring structural gender and racial inequities: MSM and TGW are still the most vulnerable populations, and, despite the overall reduction in AIDS-related mortality since 2011, it has increased among Black individuals [37–39]. Moreover, the HIV continuum of care among sexual and gender minorities is significantly worse than among the overall population of PWH [38,40]. This might be related to HIV stigma; lack of social support or access to basic rights, such as housing, food and education; and barriers to access to HIV care, representing ultimately a proxy for further socioeconomic disparities [41–45].

A lack of sustainable improvement of the HIV-continuum of care outcomes may be related to the last government's

conservative policies, which reduced HIV program budget, deepened violence against the LGBTQIA+ community and reinforced HIV stigma [36,46,47]. Furthermore, the COVID-19 pandemic disrupted healthcare services access [24,29,39]. In this sense, socioeconomic disparities and HIV-related immunosuppression synergically contributed to the overlapping of an uncontrolled HIV epidemics in Rio de Janeiro, Brazil, with a high burden of mpox cases [48].

Our study had limitations. It was a single-center study located in the second Brazilian state most affected by the mpox outbreak, thus results might not be generalizable to other contexts. Nevertheless, they are in line with data from international cohorts and case series published so far. By December 2022, mpox vaccination was not available in Brazil in the private or public sectors, remaining mostly unavailable, and no participants with mpox in our cohort



Fig. 1. HIV continuum of care outcomes among mpox-confirmed participants, according to hospitalization status during follow-up.



Fig. 2. Outcomes during follow-up stratified by CD4⁺ cell count.

were vaccinated abroad. Information on previous smallpox vaccination was subject to recall bias, and we opted to use birth date as a proxy, which could have overestimated smallpox vaccination. The low numbers of individuals with advanced HIV disease jeopardizes the possibility of establishing stronger associations, especially in the context of a global decline in number of cases, including in Brazil after a period of sustained community transmission. Most participants who required hospitalization had access to our center's inpatient care unit, allowing consistent and detailed data collection.

In conclusion, despite a usual benign clinical course, mpox-related hospitalizations during the 2022 multinational outbreak were concerning. Our findings suggest that HIV-related immunosuppression may lead to severe complications, especially in the context of ART discontinuation and poor HIV care continuum outcomes. In Brazil, the combination of public health emergencies - mpox and AIDS - unveils a landscape marked by structural inequities that reinforce a cycle of neglected diseases, affecting the most vulnerable population, increasing stigma and discrimination. Tackling social determinants of health is critical to improve ART adherence and HIV-related outcomes. Strategies to revert the negative impact of COVID-19 pandemics on HIV care are urgent. In Brazil, strengthening SUS is key to ensuring adequate responses to epidemics and outbreaks.

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Conflicts of interest

There are no conflicts of interest.

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