# Mpox in Latin America: the Emergent STI

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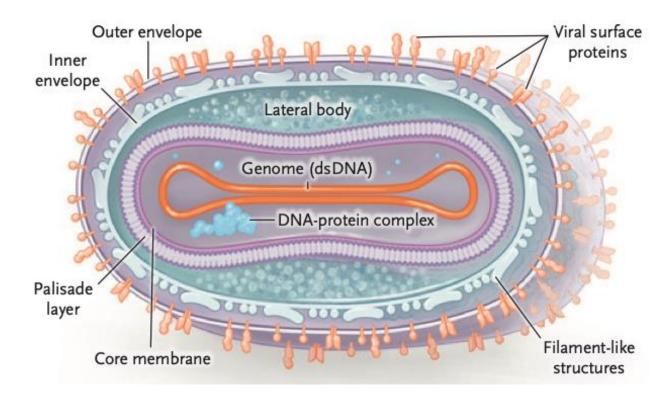






#### Monkeypox Vírus







### Pathogen Clade I Clade IIa Clade IIb



Reservoir

e.g. rodents, monkeys, rabitts



**Transmission** 

Zoonozes Person-person



**Humans** 

First case identified in 1970

#### Monkeypox virus

#### Poxviridae family

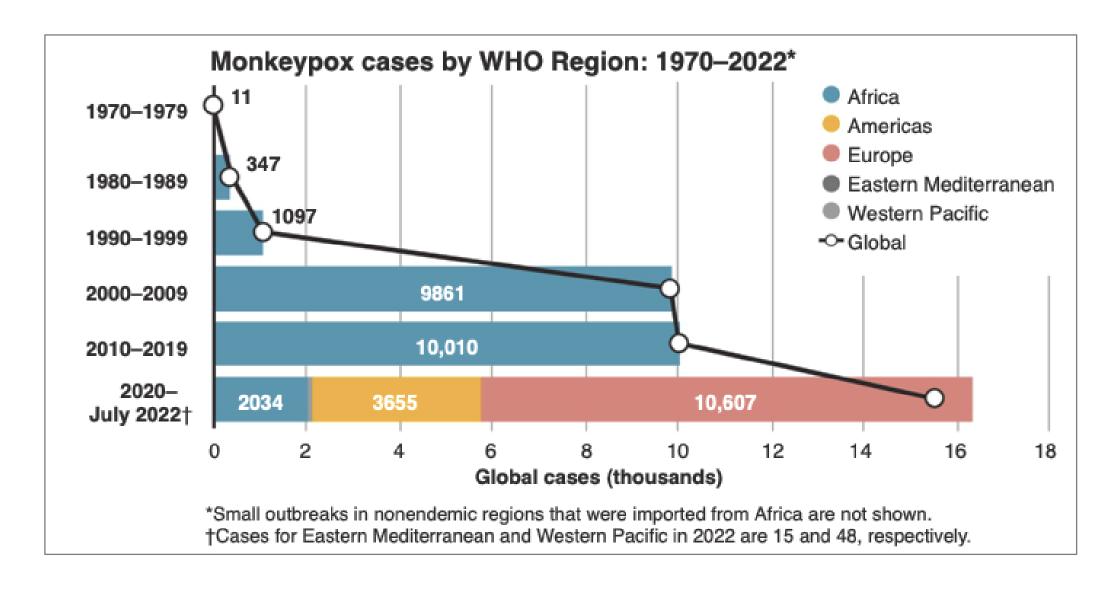
Enveloped, generally oval or brick-shaped viruses, 220–450 nm long

Large single linear molecule of double stranded DNA

Identified in 1958 in monkey in monkey colonies maintained for research in Denmark

### From a Neglected Disease to a Global Emergency



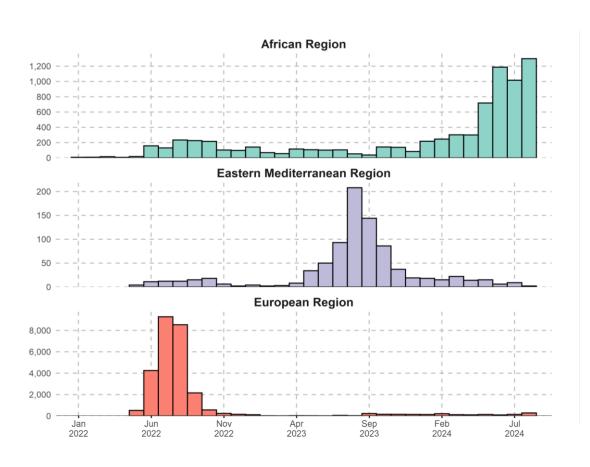


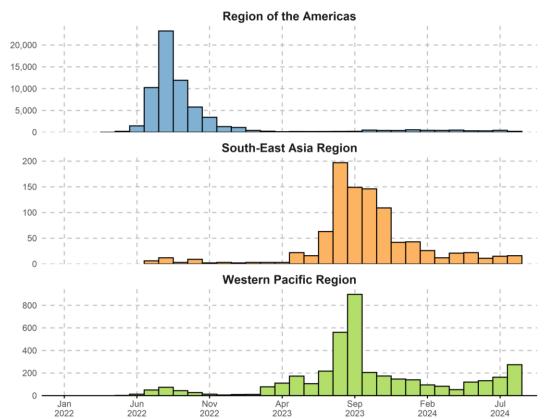
#### **Global Overview of Mpox**



#### 106 310 confirmed cases and 234 deaths reported to WHO

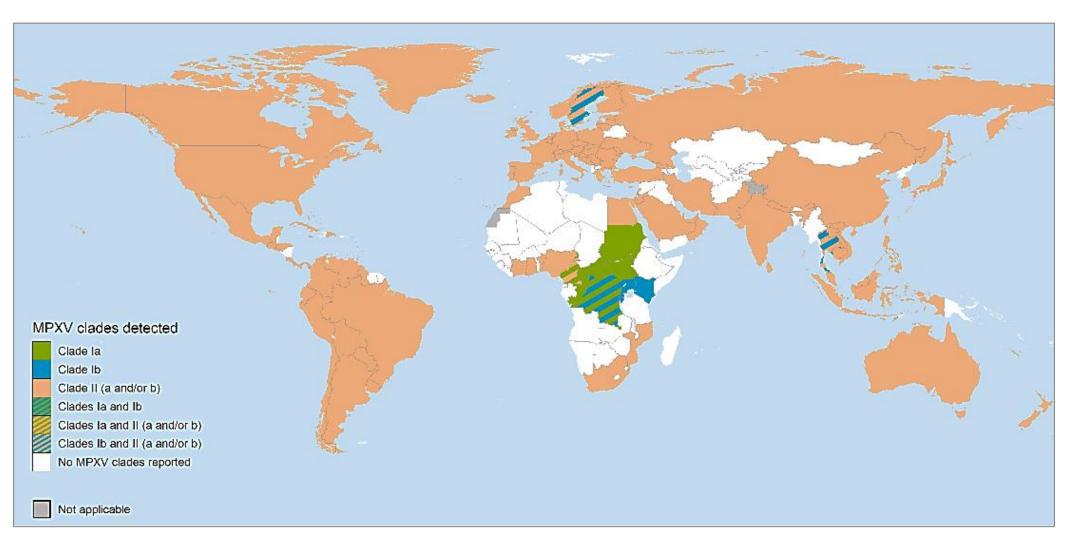
**Americas** are one of the most affected regions





#### **Mpox Outbreak - Distribution by Clade**





#### **Mpox in 2024**





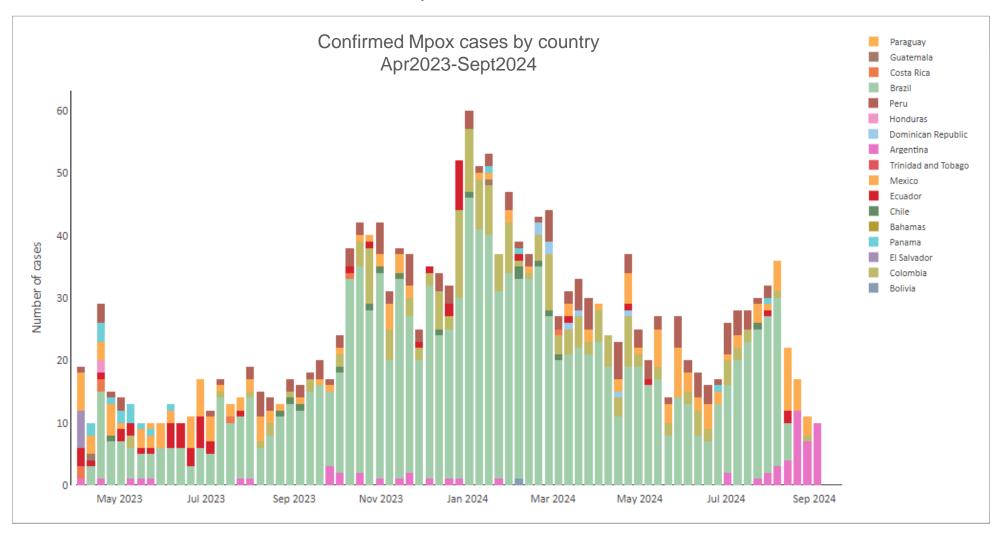


WHO Director-General declares mpox outbreak a public health emergency of international concern

#### **Mpox in Latin America**



#### 25,273 confirmed cases and 52 deaths reported to WHO



### Cases and Deaths by Country and Year for Those Reporting Cases in 2024



Country \$	Confirmed \$	Confirmed \$	Confirmed 2024	Confirmed †	Deaths 2022	Deaths \$\display\$	Deaths 2024	Deaths Total
Total	57,567	4,091	3,223	64,881	119	24	5	148
United States of America	30,052	1,776	1,986	33,814	52	7	2	61
Brazil	10,639	839	728	12,206	14	2	0	16
Canada	1,402	69	178	1,649	0	0	0	0
Colombia	4,059	89	114	4,262	0	0	0	0
Peru	3,697	162	80	3,939	20	1	2	23
Mexico	3,773	306	76	4,155	25	9	1	35
Argentina	1,025	124	32	1,181	2	0	0	2
Dominican Republic	92	10	8	110	1	0	0	1
Chile	1,400	54	7	1,461	2	1	0	3
Ecuador	466	132	7	605	2	1	0	3
Panama	89	148	4	241	0	1	0	1
Guatemala	302	103	1	406	0	1	0	1
Bolivia	261	4	1	266	0	0	0	0
Costa Rica	102	122	1	225	0	1	0	1



## Temporary adaptations to sexual behaviour during the mpox outbreak in 23 countries in Europe and the Americas: findings from a retrospective cross-sectional online survey

Mateo Prochazka, Pietro Vinti, Ana Hoxha, Andy Seale, Antons Mozalevskis, Rosamund Lewis, Ruben Mayorga Sagastume, Martha Scherzer, Leilia Dore, Meg Doherty



- 16,875 individuals from SGM
- 95% cis men
- 97% non-heterossexual
- Western Europe (48%), Latin America (37%)



- 6.4% self-reported mpox
- 30% at least 1 vaccine dose; 21% two doses ————— Lower rate in Latin America



- ~50% reported changes in sexual behavior after the mpox outbreak
- ~36% continued adaptions by May 2023

► Latin American participants more likely to adapt behavior

### High Knowledge About Mpox Among SGM in Brazil



JMIR PUBLIC HEALTH AND SURVEILLANCE

Torres et al

Original Paper

Evaluation of Mpox Knowledge, Stigma, and Willingness to Vaccinate for Mpox: Cross-Sectional Web-Based Survey Among Sexual and Gender Minorities



- 91% cis men
- 98% non-heterossexual
- Median age: 36 years

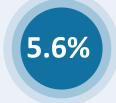


 High mpox knowledge and perception of discrimination and stigma against SGM



 ~50% reported changes in sexual behavior after the mpox outbreak

### 6.236 individuals from sexual and gender minorities



reported mpox diagnosis

- More frequently Black (16%)
- Higher proportion of PrEP users (48%)
- Lower internalized homofobia

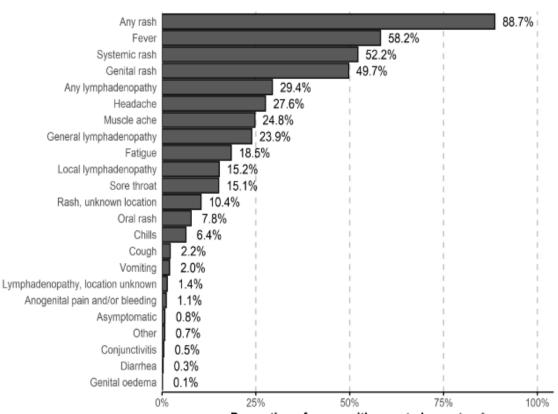
#### Main Characteristics by Mpox Clade



#### SUBSTITUIR PELO SLIDE MEG (SERÁ ENVIADO EM 2OUT)

#### **Symptoms:** n=34,743

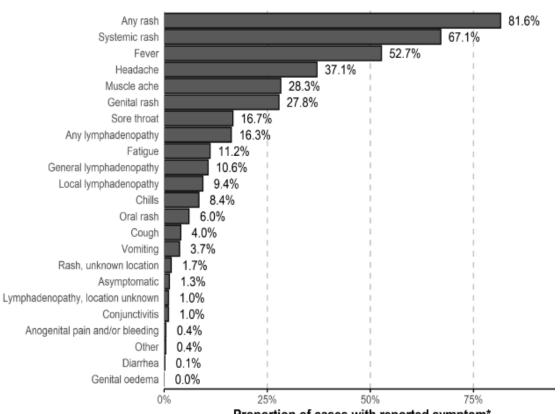
Clade 1?



Proportion of cases with reported symptom\*

\*33,430 cases with at least one reported symptom from a country where at least two unique symptoms reported used as denominator

#### Clade 2?



Proportion of cases with reported symptom\*

Source: WHO

\*1,343 cases with at least one reported symptom from a country where at least two unique symptoms reported used as denominator

### Mpox and HIV: a New Opportunistic Infection?



Mpox in people with advanced HIV infection: a global case series



Severe clinical presentation, with fulminant progression and prolonged course, associated with coalescent necrotizing lesions, occasionally involving the lungs

Day 0: genital lesions

Day 19: skin biopsy

Day 81: worsening of lesions

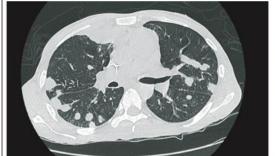
Day 81: necrotising lesions

Day 89: CT larger lung nodules

Day 89: Lung biopsy

Day 89: Lung biopsy

Day 96: CT bowel perforation





382 PWH and Mpox with CD4<350

Lower CD4 counts and higher HIV VL associated to worst outcomes

Immune Reconstitution Inflammatory Syndrome (IRIS)

Coinfections management

Tecovirimat was given to less than 20% of participants: emerging resistance?

Mpox-related mortality among PWH with CD4 <100 (%)

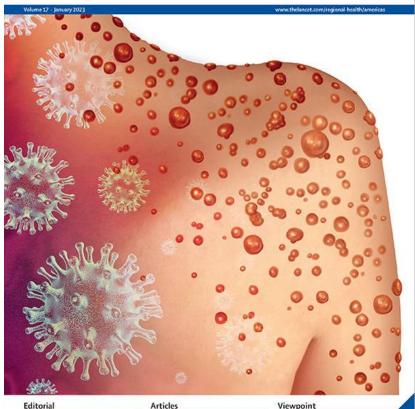
VL HIV <50 **7%** 

VL HIV >4 log 29.7%

#### **Mpox in Brazil**



#### THE LANCET Regional Health Americas



#### Editorial Articles The cycle of neglect: the mpoxemergency Epidemiological and clinical

Epidemiological and clinical characteristics of patients with human mpox infection in Mexico Monkeypox in Brazil between stigma, politics, and structural shortcomings

#### oa open access

#### Ambulatory and hospitalized patients with suspected and confirmed mpox: an observational cohort study from Brazil

Mayara Secco Torres Silva, a Carolina Coutinho, a Thiago Silva Torres, e Eduardo Peixoto, a Ronaldo Ismério, a Flavia Lessa, a Estevão Portela Nunes, a Brenda Hoagland, a Armanda Dolores Echeverria Guevara, a Matheus Oliveira Bastos, a Isabel Cristina Ferreira Tavares, a Maria Pia Diniz Ribeiro, a Maria Roberta Meneguetti Seravalli Ramos, Hugo Boechat Andrade, Ana Paula Lovetro Santana, Marilia Santini-Oliveira, Uliana Barbosa Santos Netto, Paula Reges, Monica Avelar Magalhães, Leonardo Azevedo Silva Rosadas, Sandro Nazer, Luciane Velasque, Sandra Wagner Cardoso, Edson Elias da Silva, Valdilea Gonçalves Veloso, A Mayumi Duarte Wakimoto, And Beatriz Grinsztejn, A on behalf of The INI-Fiocruz Mpox Study Groupi

- First case series report of mpox in Brazil
- Mpox confirmation by PCR





suspected cases

confirmed cases

#### **By September 20, 2024:**

- 1479 suspected cases
- 711 confirmed cases

#### Mpox in Brazil



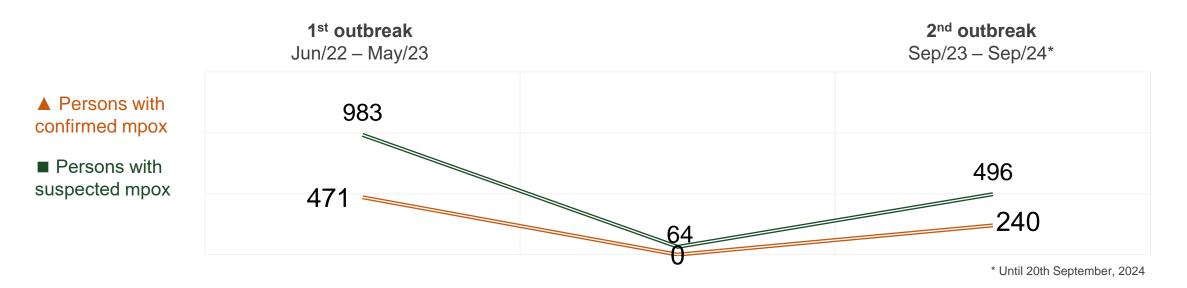
JOURNAL ARTICLE

EDITOR'S CHOICE



Exploring the Resurgence of a Neglected Disease: Lessons From the 2023–2024 Mpox Outbreak in Rio de Janeiro, Brazil 3

Number of persons with suspected and confirmed mpox according to time of diagnosis



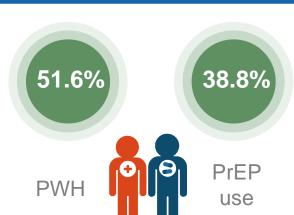
#### **INI Mpox Cohort**





confirmed cases





A	ge	
<18y	0.4%	
18-24y	12.0%	
25-29y	21.0%	
30-39y	41.0%	
40+y	25.6%	

Signs & Simptoms						
Disseminated exantema	73.1%					
Local exantema	26.9%					
Systemic symptoms	84.0%					
Genital lesions	80.4%					
Clinical features of proctitis	24.0%					
Hospitalization	10.0%					
Death	0.4%					

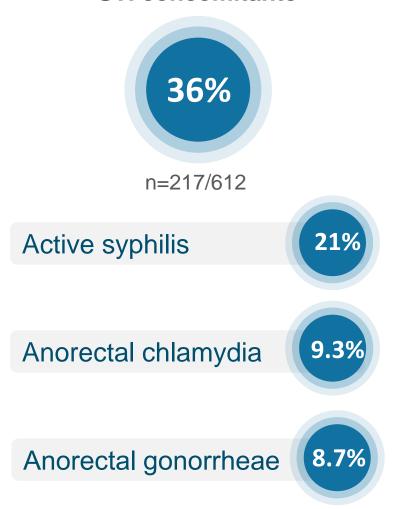
Concomitant STIs					
	Gonorrhea	9.3%			
	HBV	1.2%			
	HCV	6.4%			

Chlamydia – 8,8% Syphilis – 21%

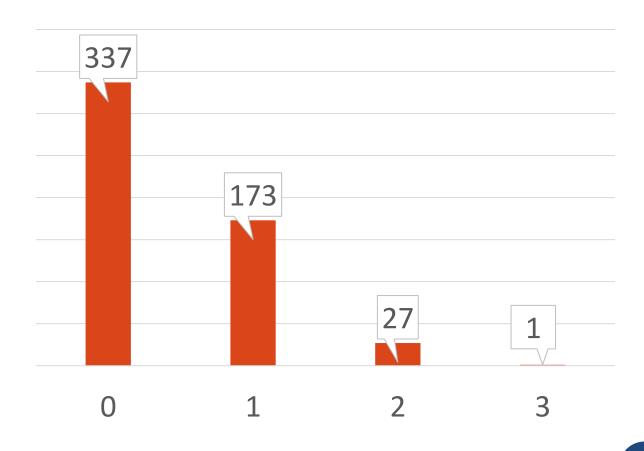
#### **Mpox and Bacterial STIs**



#### Prevalence of any concomitant bacterial STI concomitante



#### N. of concomitant bacterial STIs



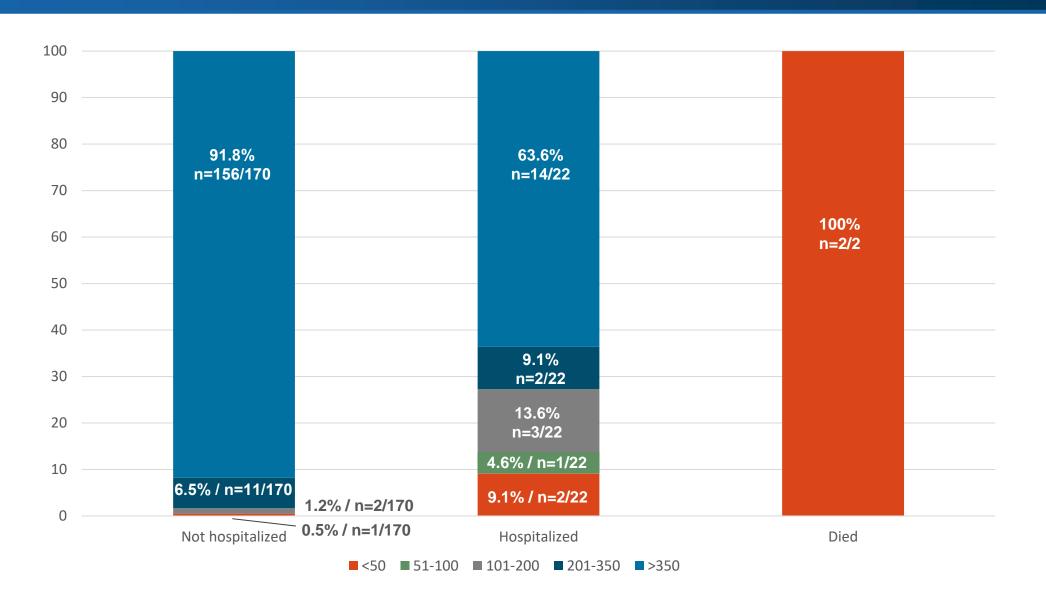
#### **INI Mpox Cohort**



	Overall, N=703 (n, %)	People without HIV, N=340¹ (n, %)	People with HIV, N=363 (n, %)	p-value	
Age (in years) [Median, IQR]	33 (28,40)	31 (26,38)	35 (30,41)	<0.01	
Gender identity: Cisgender men	641 (91.2)	290 (85.3)	351 (96.7)		
Gender identity: Cisgender women	44 (6.3)	41 (12.1)	3 (0.8)	<0.01	
Gender identity: TGW or Travesti	17 (2.4)	8 (2.4)	9 (2.5)	_	
Men who have sex with men	556 (86.7)	240 (82.8)	316 (90)	<0.01	
Fever	411 (59.2)	177 (53)	234 (65)	<0.01	
Genital lesions	559 (80.4)	255 (75.9)	304 (84.7)	<0.01	
Proctitis	193 (27.7)	72 (21.5)	121 (33.5)	<0.01	
Active syphilis <sup>2</sup>	137 (25.6)	34 (14)	103 (35.2)	<0.01	
HCV seroprevalence	43 (6.4)	8 (2.4)	35 (10.2)	<0.01	
Detectable MPXV PCR in Anorectal Swabs	101 (39)	67 (23)	168 (30)	<0.01	

#### **Mpox Hospitalization at INI-FIOCRUZ**

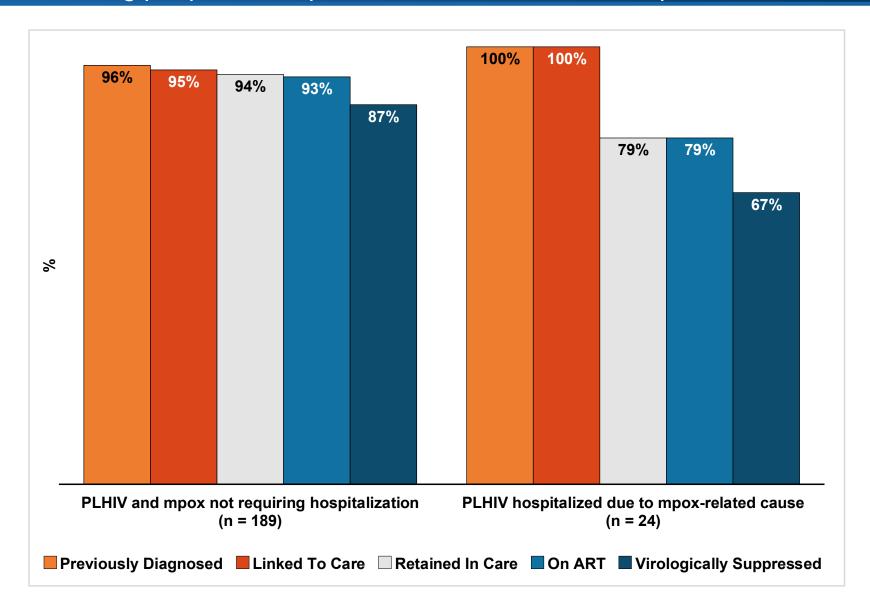




#### **Mpox Hospitalization at INI-FIOCRUZ**



HIV care cascade among people with mpox based on the need of hospitalization



#### Landscape Analysis of Mpox Therapeutics



<b>Candidate</b> Manufacturer	WHO-listed authority approved	WHO EUL	Use in under- 18s	Ongoing trials	Availability	Manufacturing capability	Comments
<b>Tecovirimat* </b> Siga	✓ EMA, US FDA**	×	×	0 Ph I 1 Ph II 6 Ph III 1 Ph IV	South Africa	Easily manufactured at scale	Primary endpoint not met in PALM007 (Clade I in DRC) PK/PD results awaited as well as Clade II results from UNITY and STOMP
Brincidofovir () Emergent BioSolutions	×	×	×	0 Ph I 0 Ph II 0 Ph III 0 Ph IV	Used under EIND for mpox in the USA	N/A	To be tested in the MOSA trial in DRC, Nigeria
VIGIV () Emergent BioSolutions	×	×	×	0 Ph I 1 Ph II 0 Ph III 0 Ph IV	N/A	N/A	Manufacturing/access at scale not currently feasible in LMICs
Cidofovir 🥡 Gilead	X	×	X	0 Ph I 0 Ph II 1 Ph IIII 0 Ph IV	N/A still in trials	N/A	No additional trials planned

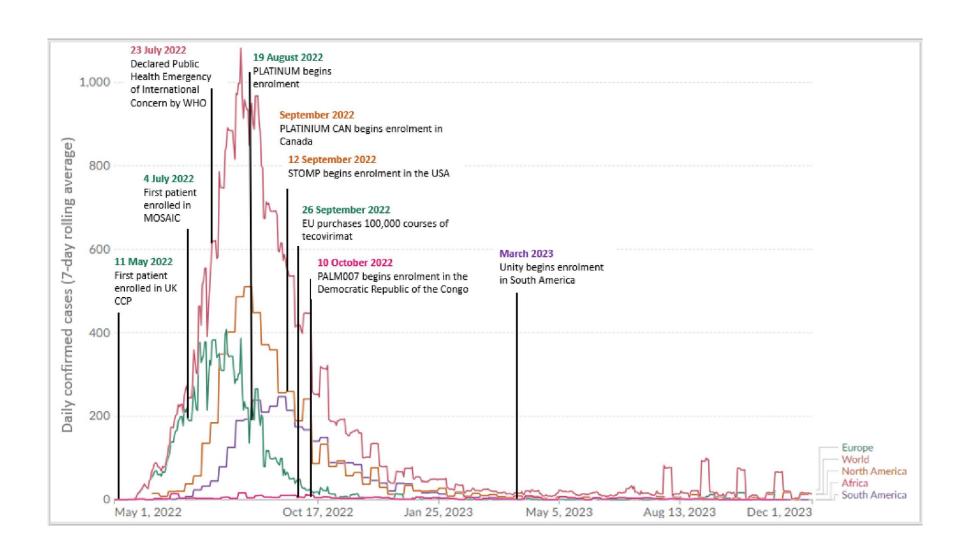
Novel antivirals: 3 novel antiviral candidates for mpox in preclinical development; 1 in phase 1 (ASC10)

Monoclonal antibodies (mAbs): 2 anti-mpox mAbs with ongoing preclinical studies [BFI 753 (Biofactura) and JEPO-CBRND (Just Evotec)]



### Timeline of the Events During the 2022 Multi-country Mpox Outbreak and Daily Confirmed Cases (Seven Day Rolling Average)





#### Trials











**STUDY DESIGN** 

Randomized, placebo-controlled double blinded study to evaluate the safety and efficacy of tecovirimat



STUDY PARTICIPANTS

Symptomatic Mpox <14d Any age Symptomatic Mpox any duration >14 years of age Lab confirmed Mpox any duration >13kg



**OBJECTIVE** 

Time to lesion resolution

Time to lesion healing

Time to lesion resolution



**SECONDARY** 

Pain, viral clearance, complete healing

Viral clearance, lesion resolution

Viral clearance, complete healing

#### International Collaboration Led by PALM



#### **PALM 007**

- Randomized, placebo-controlled, double blinded study to test safety and efficacy of Tecovirimat
- Population
  - Adult and pediatric participants
  - Mpox of any duration
  - Clade I
- Primary outcome: Time to lesion resolution = all lesions are scabbed, desquamated or a new layer of epidermis has formed
- Protocol shared with other mpox trials



#### PALM 007 Topline – Safe But Not Effective



#### **NEWS RELEASES**

Thursday, August 15, 2024

The antiviral tecovirimat is safe but did not improve clade I mpox resolution in Democratic Republic of the Congo

NIH-cosponsored study examined tecovirimat in mpox-endemic country.

Mortality decreased to 1.7% in all participants indicating that better outcomes can be achieved with high quality supportive care

### CDC Expanded Access – Investigational New Drug (EA-IND)



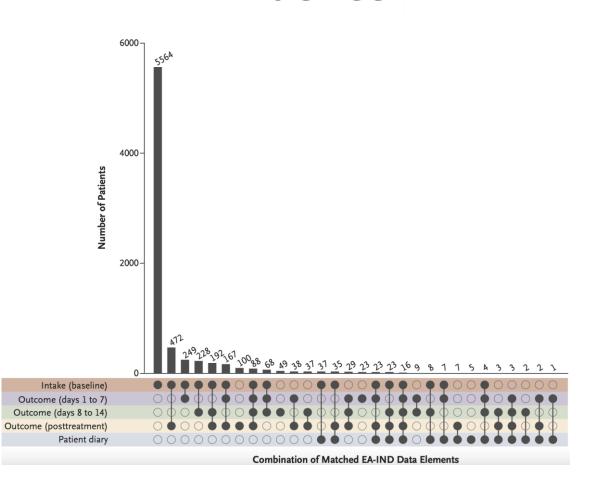
**ORIGINAL ARTICLE** 

#### Tecovirimat Use under Expanded Access to Treat Mpox in the United States, 2022–2023

#### NEJM Evidence

- Clade II mpox outbreak
  - May 29, 2022

     July 10, 2023
- Tecovirimat prescribed for > 7100 patients
  - Median age 35 (IQR 30, 43)
  - 51.7% PLWH
  - 72.4% were outpatient Median time from sx onset to tx 7 days (IQR 4, 10)
  - 92.6% <100 lesions; 40.5% <10
- 223 SAEs and 40 deaths



#### Brincidofovir



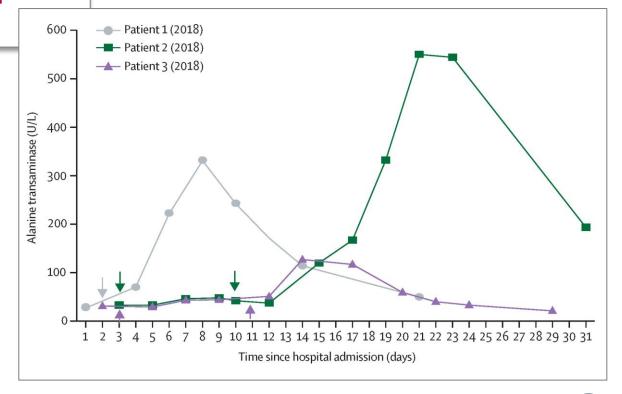
#### Clinical features and management of human monkeypox: a retrospective observational study in the UK



Hugh Adler, Susan Gould, Paul Hine, Luke B Snell, Waison Wong, Catherine F Houlihan, Jane C Osborne, Tommy Rampling, Mike BJ Beadsworth, Christopher JA Duncan, Jake Dunning, Tom E Fletcher, Ewan R Hunter, Michael Jacobs, Saye H Khoo, William Newsholme, David Porter, Robert J Porter, Libuše Ratcliffe, Matthias L Schmid, Malcolm G Semple, Anne J Tunbridge, Tom Wingfield\*, Nicholas M Price\* on behalf of the NHS England High Consequence Infectious Diseases (Airborne) Network†



- First three patients were treated with oral brincidofovir ~7 after onset of rash
- All three patients developed elevated alanine transaminase and none completed the course of treatment
- Evidence of synergy with tecovirimat in animal models of Cowpox



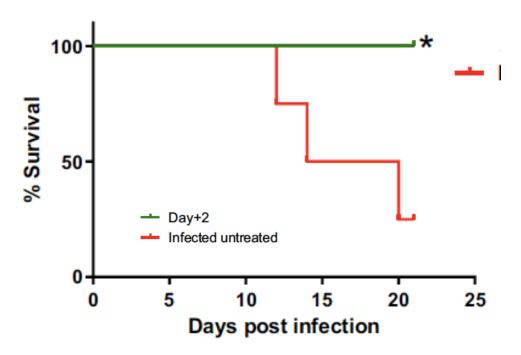
#### **Mpox mAbs – Effective in Animal Models**

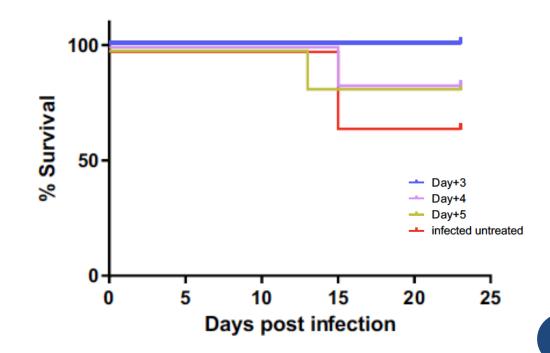




https://doi.org/10.1038/s41467-024-47328-y

Synergistic effect of two human-like monoclonal antibodies confers protection against orthopoxvirus infection





#### **Mpox Vaccination**

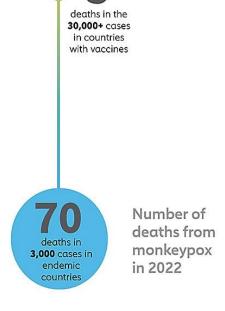


#### Monkeypox Vaccine Access & Mortality, 2022

Countries receiving monkeypox vaccines in 2022



No vaccines have been allocated or purchased by African countries



### Latin America: mpox vaccine access

Mexico: none

Argentina: none

Venezuela: none

Brazil: yes, for vulnerable individuals

Chile: yes, for vulnerable individuals

Dominican Republic: yes, for

vulnerable individuals

Peru: vulnerable populations in 2022

Ecuador: yes, for vulnerable

individuals in 2022

Mostly no actual access

#### **Final Remarks**



Challenges in genomic surveillance and laboratory diagnosis

Mpox and HIV coinfection in the context of HIV late diagnosis and disengagement from care

Equitable distribution of medical countermeasures, tackling North-South disparities

#### Acknowledgements



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# Thank you! Obrigada!

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