

# Mpox and HIV

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• 14 August 2024: WHO Director-General declared **mpox** a Public Health Emergency of International Concern (PHEIC) to respond to the increase in cases and multi-country spread of mpox from the Democratic Republic of Congo



WHO calls emergency meeting over surge in mpox cases spreading from Congo The committee will meet as soon as possible and will be made up of independent experts from a range of relevant disciplines Detected In Its Neighbouring Countries (Photo: Reuters (Connect with us (□) < □

Sept 2025 WHO: PHEIC rescinded

Africa CDC retained Mpox as Public

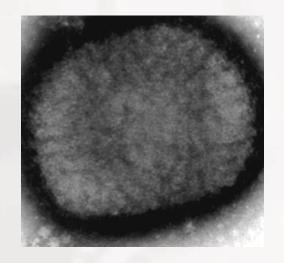
Health Emergency of Continental

Security

### I. What is mpox?

- Mpox is caused by *Orthopoxvirus monkeypox (MPXV)* (ICTV: 2023), a member of the *Orthopoxvirus* genus in the family *Poxviridae*.
- Causes mild to severe rash in people, and may be life threatening in immunocompromised or young children.
- Emerging viral infection first reported in 1970, but causing outbreaks classified as Public Health Emergency of International Concern (PHEIC) since 2022
- Depending on the MPXV variant: zoonotic and/or human-to-human transmission.

Electron micrograph of orthopoxvirus particle
Dr Monica Birkhead, NICD









Images from DermNet: <a href="https://dermnetnz.org/">https://dermnetnz.org/</a>



Zoonotic transmission: X

1970-1986: 245 of 338 cases with zoonotic link, 93 possible human-to-human transmission.

Close contact with infected animals / Bites, scratches, bush meat preparation (in adequately cooked meat; slaughtering; animal derived products), contact with contaminated materials.

True host/reservoir still unknown.

Have been isolated from rope squirrel, mangabey. Also other mammals including tree squirrels, Gambian pouched rats, dormice, other non-human primates.

Person-to-person transmission:
 Close contact (prolonged face-to-face contact, kissing, sexual contact).

Contact with materials contaminated with virus (via scabs, lesion fluid, for example contaminated linen/clothes).

Large droplet transmission is possible. Virus enters the body through broken skin, respiratory tract, or the mucous membranes (eyes, nose, or mouth).

Household outbreaks, up to 7 transmissions in chain<sup>1</sup>

Secondary attack rate, approx. 10% in smallpox unvaccinated individuals.







Source of images: Wikipedia

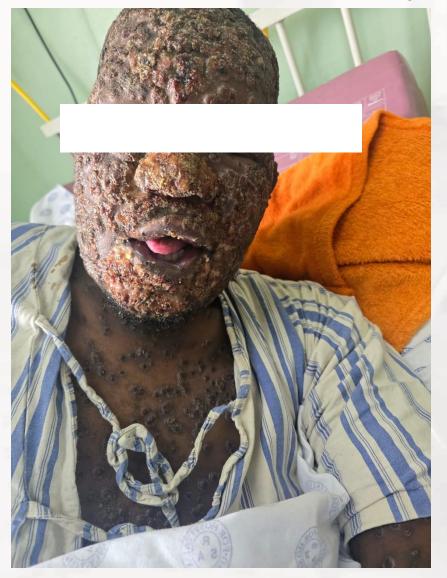
<sup>&</sup>lt;sup>1</sup> **Nolen et al.** Extended human-to-human transmission during a monkeypox outbreak in the Democratic Republic of the Congo. *Emerg Infect Dis.* 2016; **22**: 1014-1021

- 35 years male, MSM
- Fever, sore throat, night sweats, myalgia, fatigue
- 1 week later- painless, pruritic skin lesions 2-15mm on hand, then genital lesion
- 10 days later –extensive painful lesions
- HIV +, Swab lesion : MPOX Clade 11b
- Rx- isolation, contact tracing, analgesia, ARV
- Tecovirimat x 2/52



SAMJ 2024 Permission granted

# Mpox Clade 1b





### **Mpox in South Africa**

 Ongoing reintroductions, with and without transient local transmission noted:

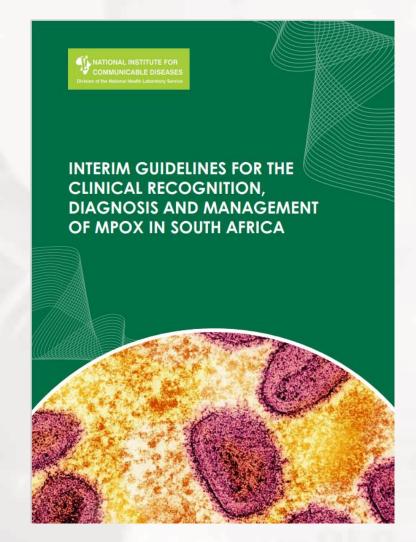
2022: 5 laboratory confirmed cases, no travel histories, ex GP and WC, Clade IIb B.1.7

2023: no cases detected

<u>2024</u>: 25 cases confirmed, 1 with travel history (ex Peru, Clade IIb B.1.6), other no travel history (Clade IIb B.1.20, now F2), after August 2024 no further cases detected, ex KZN/GP/WC (three deaths)

<u>2025</u>: Since Feb, 11 cases laboratory confirmed cases, 9 Clade Ib and 2 Clade Ib (sublineage F2)

- Interim guidelines for mpox in South Africa available from NICD website: <a href="https://www.nicd.ac.za/key-reference-documents/">https://www.nicd.ac.za/key-reference-documents/</a>
- 1<sup>st</sup> vaccination roll out in July 2025, see <a href="https://www.gov.za/news/media-statements/health-rolls-out-mpox-vaccination-new-cases-are-detected-16-jul-2025">https://www.gov.za/news/media-statements/health-rolls-out-mpox-vaccination-new-cases-are-detected-16-jul-2025</a>



	CLADE I		CLADE II		
	Clade Ia	Clade Ib	Clade IIa	Clade IIb (Lineage A)	Clade IIb (Lineage B.1)
Geographic location	Central Africa	DRC and beyond	West Africa and exotic pet trade outbreak in US (2003)	Nigeria	Multi-country outbreak
Transmission dynamics	Approx. 60-75 % zoonotic  Approx. 5-40% human-to-human	100% human-to-human Sexual contact as NB mechanism Close contact	100% zoonotic (no data available from endemic countries, based on US exotic pet trade outbreak)	Transmission mode unknown (61.8%), humanto-human (30%) suspected zoonotic (8.2%)  (2017-2019 from Nigeria)	100% human-to-human Sexual contact as NB mechanism Close contact
Animal reservoir	Unknown	NA	Unknown	Unknown	NA
Demographics	90% in < 15 y	85% in adults in DRC 50% in adults in Burundi	70 % adults	80% adults, 70% males	99% adults, 98% males
Rash characteristics	Mostly centrifugal, often on face (>80%) >100 lesions	Mixed (often with oral, genital or anogenital lesions)	Mostly centrifugal from point of animal contact	Mixed (centrifugal), genital lesions in 60-85%	Many anogenital, localized
Mortality  Adapted from presentation by	5-10% mostly children (less with improved clinical care?)	0.7 %	0%	3-5% in young adults with advanced HIV	0.2%

Adapted from presentation by L Subissi, WHO, September 2024 Full references in notes section

WHO/WHE/IHM/2018.3

Organisation mondiale de la Santé

### Monkeypox

Current status in West and Central Africa



16 MARCH 2018, 936 YEAR / 16 MARS 2018, 93" ANNÉE No 11, 2018, 93, 117-132

### Emergence of monkeypox in West Africa and Central Africa, 1970-2017

http://www.who.int/wer

Kara N. Durski, \*a Andrea M. McCollum, \*b Yoshinori Nakazawa, b Brett W. Petersen, b Mary G. Reynolds, b Sylvie Briand, a Mamoudou H. Djingarey,<sup>c</sup> Victoria Olson,<sup>b</sup> Inger K. Damon,<sup>b</sup> and Asheena Khalakdina

Report of a WHO Informal Consultation Geneva, Switzerland, 3 November 2017



USA, 2003: 72 reported cases (37 lab confirmed, 100% recovery) of mpox diagnosed from different locations in the US Midwest. No human to human transmission noted.

### First report of human mpox outside of Africa.

Linked to exotic pet trade and involved prairie dogs which was co-housed with several other animals that originated from Ghana, including: rope squirrels (Funiscuirus sp.), tree squirrels (Heliosciurus sp.), Gambian giant rats (Cricetomys sp.), brushtail porcupines (Atherurus sp.), dormice (Graphiurus sp.), and striped mice (Hybomys sp).

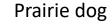
### Monkeypox: Suspected trail of infection

Disease carried into US

by rats imported from

Africa as exotic pets







Disease spreads to prairie dogs captured in Texas for use as pets

#### HUMANS

Contract disease when scratched or bitten by infected prairie dogs

SOURCE: CDC



- Incubation period- 3-17 days Fever/prodrome
  Lymphadenopathy
- Rash blistering, umbilicated, crust
- May be painful, NOT itchy
- Few to 100s of lesions
- Face, limbs, trunk, ano- genital, oral



a) early vesicle,3mm diameter



d) ulcerated lesion, 5mm diameter



b) small pustule, 2mm diameter



e) crusting of a mature



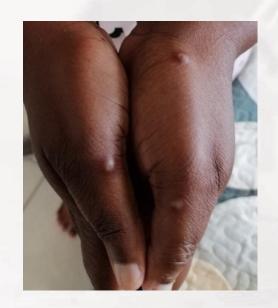
c) umbilicated pustule, 3-4mm diameter

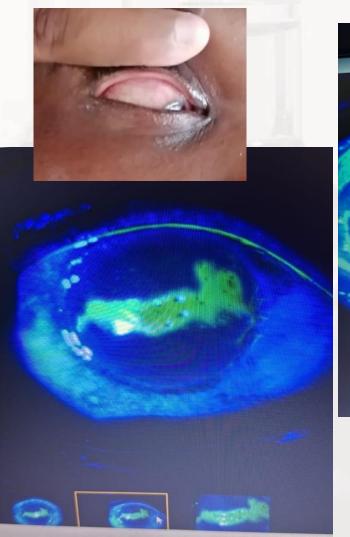


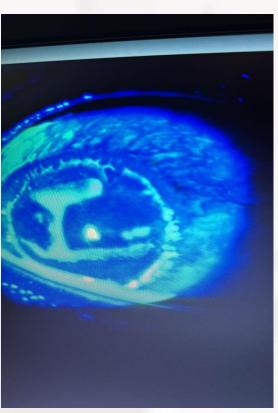
f) partially removed





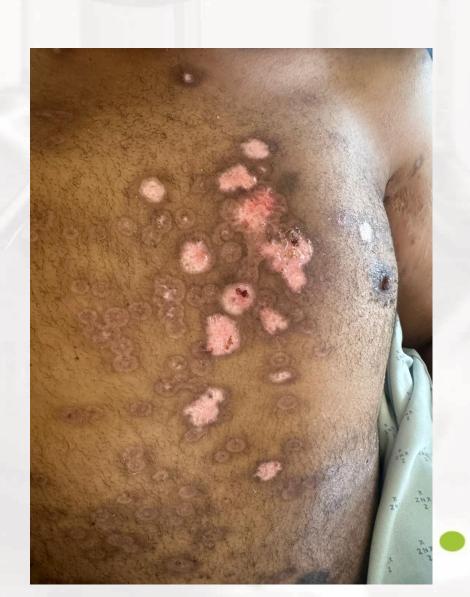






# Post mpox scarring





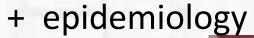
Mpox plus KS



# Fatal mpox case, advanced HIV, misdiagnosed as deep fungal infection



<u>Differential diagnosis</u>: consider rash characteristics





# Varicella zoster: chickenpox





Prodrome of fever and malaise. Itchy papules that become vesicular and sometimes pustular. Lesions at different stages Begins on face usually, then spreads to trunk

Rx with acyclovir in adults – complications in immune and suppressed adults. NB pneumonia

### **Mpox in South Africa**

- Mpox is a category I notifiable medical condition in South Africa via electronic notification system, see www.nicd.ac.za
- Laboratory testing for suspected cases of mpox is available via NICD and private pathology laboratories
- Guidance for sample collection is available from <u>www.nicd.ac.za</u>

Specimen type	Collection materials	Comments
Skin lesion material: Swabs of lesion exudate Roofs Lesion crust	Dacron or polyester flocked swabs with VTM or dry swab	Required for all investigations
Throat swab	Dacron or polyester flocked swabs with VTM or dry swab	Optional
Rectal and or genital swabs (if lesions present)	Dacron or polyester flocked swabs with VTM or dry swab	Optional
Semen	Urine specimen jar	Optional
Plasma	EDTA collection tube (purple top)	Optional
Serum	Serum separator tubes or clotted blood	Optional







- Package of care in LMIC: analgesia, nutritional support, fluids, treatment of secondary infections, skin care, home/hospital isolation
- Link diagnosis, treatment and prevention to HIV programmes: manage HIV -related opportunistic infections, STI's
- Infection prevention control
- Psychosocial considerations: stigma, contact tracing challenges
- Public health response

- Established an mpox hotline for review of all suspected cases- 24/7
- ID specialists, dermatologists, medical officers



HIV AND MPOX

### Worse if CD4 <350.

- Case series (Lancet 2023; 939-49): 382 cases
- Median CD4 211; ½ with suppressed viral load.
- Severe complications more common if CD4 <100 vs CD4 >300:
  - Necrotising skin lesions (54% vs 7%), lung involvement (28% vs 0%), secondary infections/sepsis (44% vs 9%)
  - ¼ hospitalised, of whom ¼ died (all deaths had CD4 <200, and more likely if high VL)

# Always check HIV in anyone who you think has mpox!

# HIV and timing of ART

- No good direct evidence on the timing.
- **But** immunological control is the best way of treating mpox.

- Therefore, start ARVs as soon as possible (certainly with 7 days).
  - WHO guidance

 IRIS risk not well established, but benefits of early ART initiation likely outweigh any risks.

Severe mpox, newly diagnosed HIV on ART compliance issues
Good response to tecovirimat –
'self discharge'



Readmission: new active mpox lesions

NOT IRIS, rather progressive mpox needing longer tecovirimat therapy





# Other options in HIV patients



**Chickenpox**: itchy papules that become vesicular and sometimes pustular. Begins on face usually, then spreads centripetally. Prodrome of fever and malaise.

**Zoster**: same evolution, but pain more prominent and itching less so. Dermatomal distribution.





# Other options



**Molluscum contagiosum**: smaller (2-5mm), usually infects children, though also associated with HIV too (in which case they're sometimes bigger).

Firm papules (not vesicles).

Umbilicated, skin-coloured/pinkish.

Resolve spontaneously in weeks/months in nonimmunocompromised individuals (much slower evolution)

### **Scabies**

- Intense itching, with onset of pustular rash
- Typical areas affected include the wrist, elbow, armpit, webbing between the fingers, nipple, penis, waist, belt-line, and buttocks –
- Small raised lines (burrows) may be visible on the skin, caused by the female scabies mite tunnelling beneath the skin surface
- Involvement of the palms, soles, head, face, and neck may be seen in infants and young children



### Vesicles & Bullae

### Infectious

### Viral:

- Herpesvirus HSV, VZV
- Enterovirus (echovirus, coxsackie virus)
- Poxvirus variola, vaccinia

### Bacterial:

- Staph aureus TSS, SSSS
- Streptococcus group A (TSS)
- Rickettsiae R. africae
- V. vulnificus
- M. pneumoniae

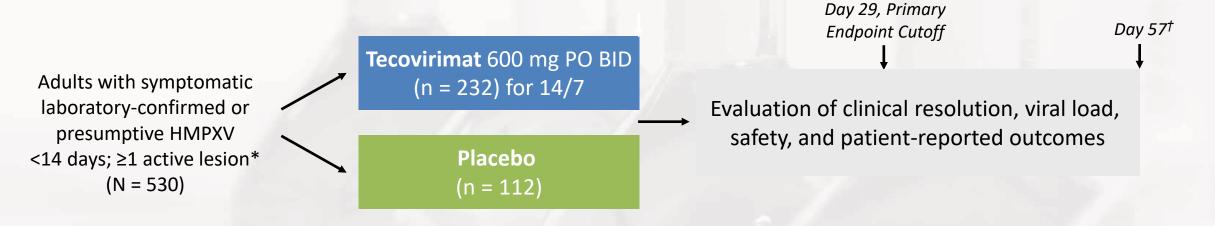
### Non-infectious

- Acute eczema
- Erythema mutiforme
- TEN
- Thermal burn / frostbite
- Bullous pemphigoid



### **STOMP** trial

Mild-moderate disease, without severe immunosuppression



<sup>†</sup>Study was stopped early based on an interim futility analysis requested by the independent DSMB.

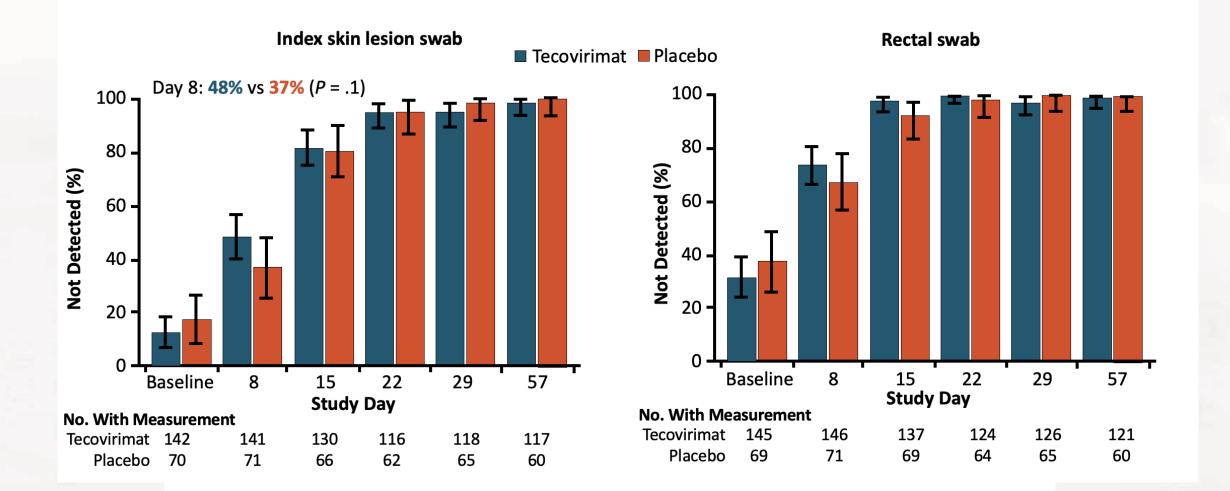
**Primary endpoint:** time to clinical resolution (defined as all skin lesions scabbed or epithelialized and all visible mucosal lesions healed) by Day 29

Secondary endpoints: daily pain score, mpox detection in various compartments, and patient-reported outcomes



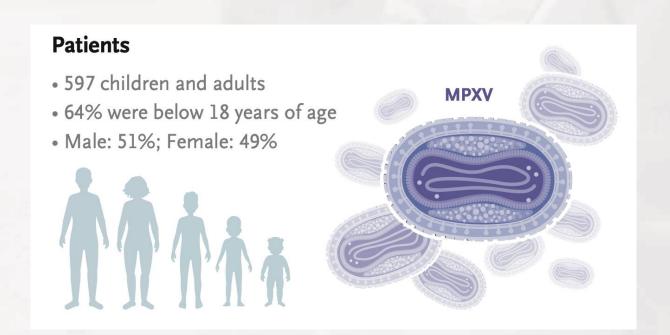
## Results

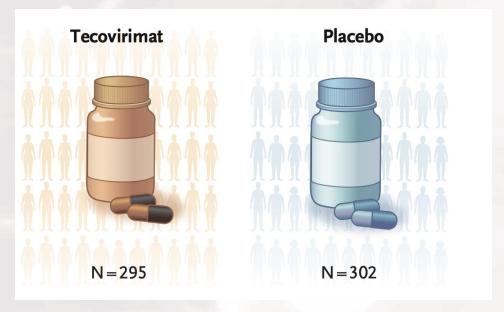
### **STOMP: Viral Clearance**



### PALM007 trial

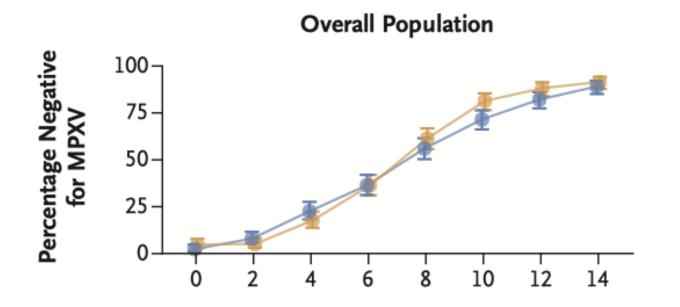
- Who? Children & adults with confirmed mpox (clade I) & ≥1 skin lesion.
- Where? DRC
- What? Double-blind, placebo-based RCT
- Outcome? Time to lesion resolution (all lesions scabbed/epithelialised)





#### PALM007 - Results

#### C Skin-Lesion Specimens



Also no difference whether started early (≤7 days) or late (> 7 days)

#### Percentage with Lesion Resolution

Tecovirimat 0 0.7 13.2 33.9 59.7 80.3 87.1 90.8 Placebo 0 3.0 16.9 34.1 55.0 69.9 81.5 88.4

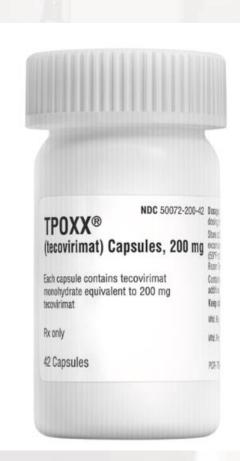
## Treatment - summary

	STOMP	PALM007	
Population	Mostly men	Men, women, children	
Severity	Mild-moderate	All levels	
Clade	II	I	
Clinical effect?	No	No	
Virological effect?	No	No	
HIV	34%	0.7%	
Severely immunosuppressed	No	Not known	

Is there still a role for tecovirimat in patients with advanced HIV and similarly immunosuppressed patients?

#### In South Africa

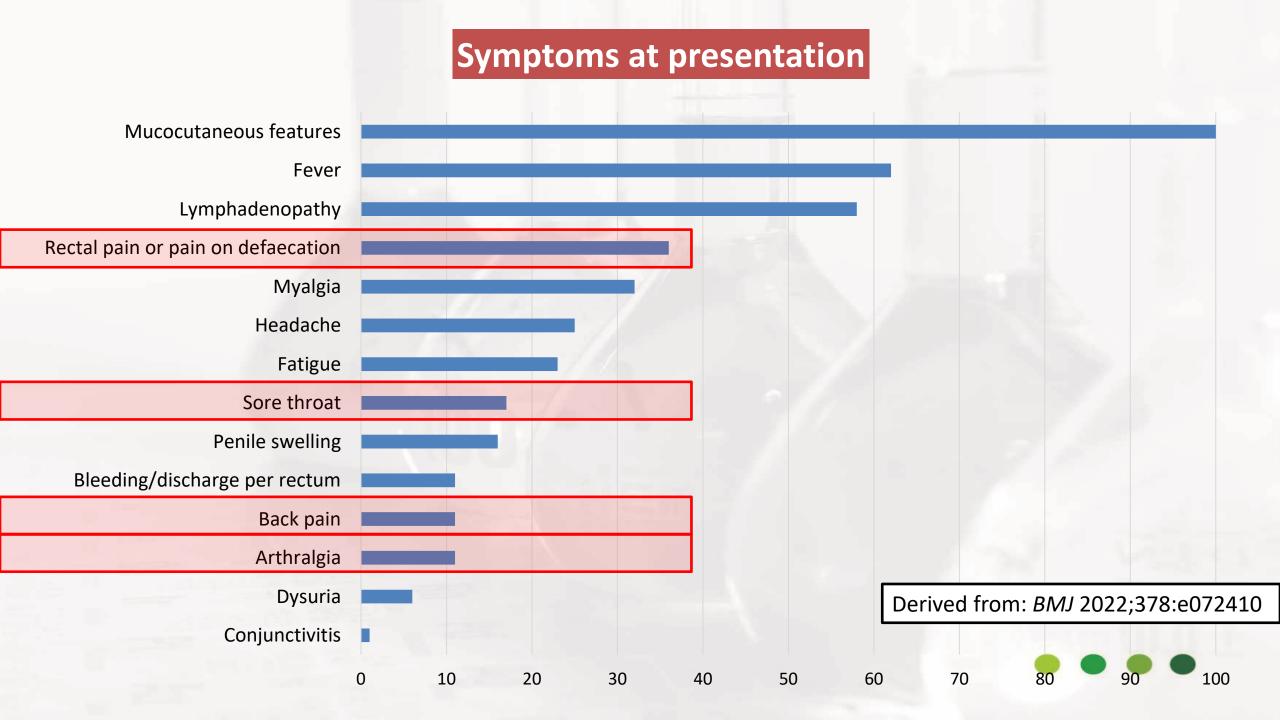
- Eligible if confirmed mpox AND symptomatic AND severe/complicated disease
- Contraindications: severe renal/hepatic impairment, pregnancy
- Section 21 application (fast-tracked)
- Stockpile centrally stored in Pretoria for both state and private sectors
- Treating physician completes section 21 progress report and WHO reporting tool







PAIN MANAGEMENT

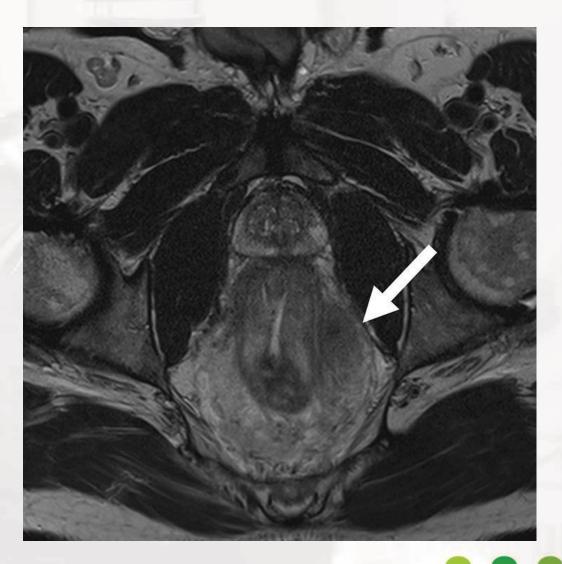


## Pain as the dominant symptom

- Skin lesions not typically that painful unless secondarily infected.
- But mucosal involvement often described as "10/10" pain.
- Pain is the most common reason for hospitalisation.
- Rectal pain → tenesmus or constipation.
- Penile pain → urinary retention
- Oral/pharyngeal pain → inability to eat or drink

## T2 weighted magnetic resonance imaging scan of pelvis showing a 3.5 cm cavity in left mesorectum, adjacent to the rectal wall representing an area of localised perforation (arrow)

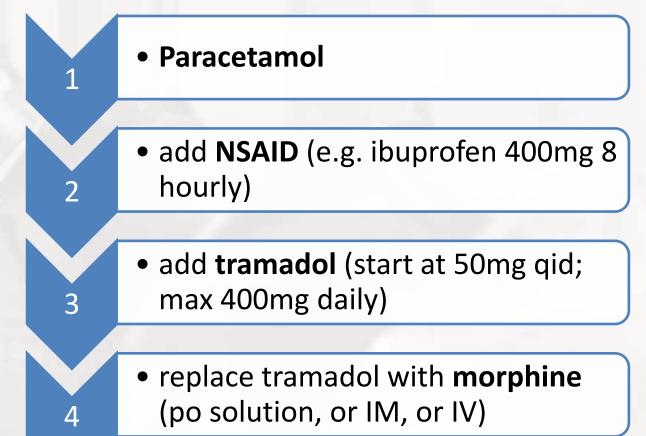






## Management of pain

Rectal > penile >> skin



# Pain – tips and tricks

#### Mouth: try to prevent secondary infection:

- Rinse with clean salt water 4x daily
- Topical lidocaine gel

#### Genital/anorectal lesions:

- Sitz baths
- Topical lidocaine gel
- Constipation: lactulose, bisacodyl, senna
- Tenesmus/spasms: hyoscine butylbromide



PREVENTION

## Infectious period

- Infectious until scabs have all fallen off and there is a fresh layer of healthy skin underneath.
  - Usually takes 2-4 weeks

- Infectious period starts around the time when symptoms do.
  - Some evidence for infectious period preceding symptoms by a few days in rare cases.

## **Precautions: Hospital**

- Isolate in single-person room with dedicated bathroom.
- Keep door closed, but special air handling not required. (CDC)
- Limit patient movement outside of their room
  - If unavoidable, cover any exposed lesions and get patient to wear a medical mask.
- Perform aerosol-generating procedures like intubation/extubation in airborne infection isolation room.
- Standard disinfection and cleaning procedures & soiled laundry practices.

# PPE: healthcare workers

- Gloves
- Gown
- Medical mask
- ± eye protection



### Advice for home: while infectious

- Avoid close contact with others:
  - Own room & bathroom if possible.
  - Abstain from sex
  - Cover up lesions and wear mask when in close contact with others.
- Try to cover & manage own lesions

WHO suggests that persons with mild, uncomplicated mpox infection cared for at home are not required to isolate provided their lesions are covered and they wear a well-fitting medical mask when in close proximity with others until all lesions are healed. (Conditional recommendation, low certainty evidence)

## Contact monitoring

Monitor close contacts for 21 days after last exposure.

• Individuals exposed to monkeypox virus can continue their routine daily activities (e.g., go to work or school) as long as they do not have signs or symptoms consistent with mpox.



VACCINATION

## Vaccines

All originally developed as smallpox vaccines

	MVA-BN (Imvanex /Jynneos)	ACAM2000	LC16m8
Company	Bavarian Nordic	Emergency BioSolutions	KM Biologics
Туре	Live, non-replicating modified vaccinia Ankara (MVA)	Live, replicating vaccinia virus	Live, attenuated vaccinia virus
Major approvals	FDA, EMA, WHO	FDA for smallpox	Japan (& WHO EUL)
Dosing	2 doses ≥ 28 days apart	1 dose	1 dose
Efficacy	66-90% effective (35-80% with 1 dose)	Extrapolated from smallpox data	Extrapolated from smallpox data
Side effects	Injection site reactions mostly	Injection site reactions mostly. Myocarditis. Can't be used in IMC.	Injection site reactions mostly. Caution in IMC.

#### Vaccine rollout – Africa

- WHO prequalification for MVA-BN in Sept 2024
  - enabled organisations to more readily procure it for LMIC
- Access and Allocation Mechanism (AAM) allocated 900,000 doses to 9 African countries: DRC (85%), Kenya, Nigeria, South Africa, Rwanda, Uganda, Côte d'Ivoire, Liberia, and CAR
  - From Canada, Gavi, Vaccine Alliance, European Union, USA
- Japan donated 3 million LC16m8 doses to DRC via bilateral agreements.
- Current status (mid-2025):
  - 3 million doses distributed (~half MVA-BN, ~half LC16m8), but only 900,000 administered (69% in DRC). Target was 10 million within 6 months.
- Reasons: fragile local healthcare systems, funding shortages, poor supply chain management, vaccine shortages, delays in local approvals, wastage, weak local surveillance, stigma/mistrust

#### In South Africa

Section 21 approval for 10,000 doses

#### Eligibility:

- Gay, bisexual and MSM
- Sex workers & anyone participating in high-risk sexual behaviours
- Healthcare and lab workers at risk of exposure to patients/samples with mpox
- At-risk travelers to West and Central Africa
- Consider: PLWH in general too
- Contacts of persons with mpox sexual, household, healthcare
- Provincial vaccination sites in Gauteng, KZN, and Western Cape
- Screening tool & informed consent required
- Delayed-dose strategy (2<sup>nd</sup> dose once more stock available)

# Vaccine-escape variant – hopefully unlikely

- DNA virus low mutation rate
- Breadth of vaccine protection is wide:
  - At least 100-200 conserved T-cell epitopes across orthopoxviruses (MPXV shares ≥71% with vaccinia)
  - 6-10 major proteins targeted for humoral response
  - Includes peptides from multiple viral lifecycle stages (attachment, entry, replication)
- No evidence of significantly reduced efficacy with new variants yet
- New vaccines in the pipeline too