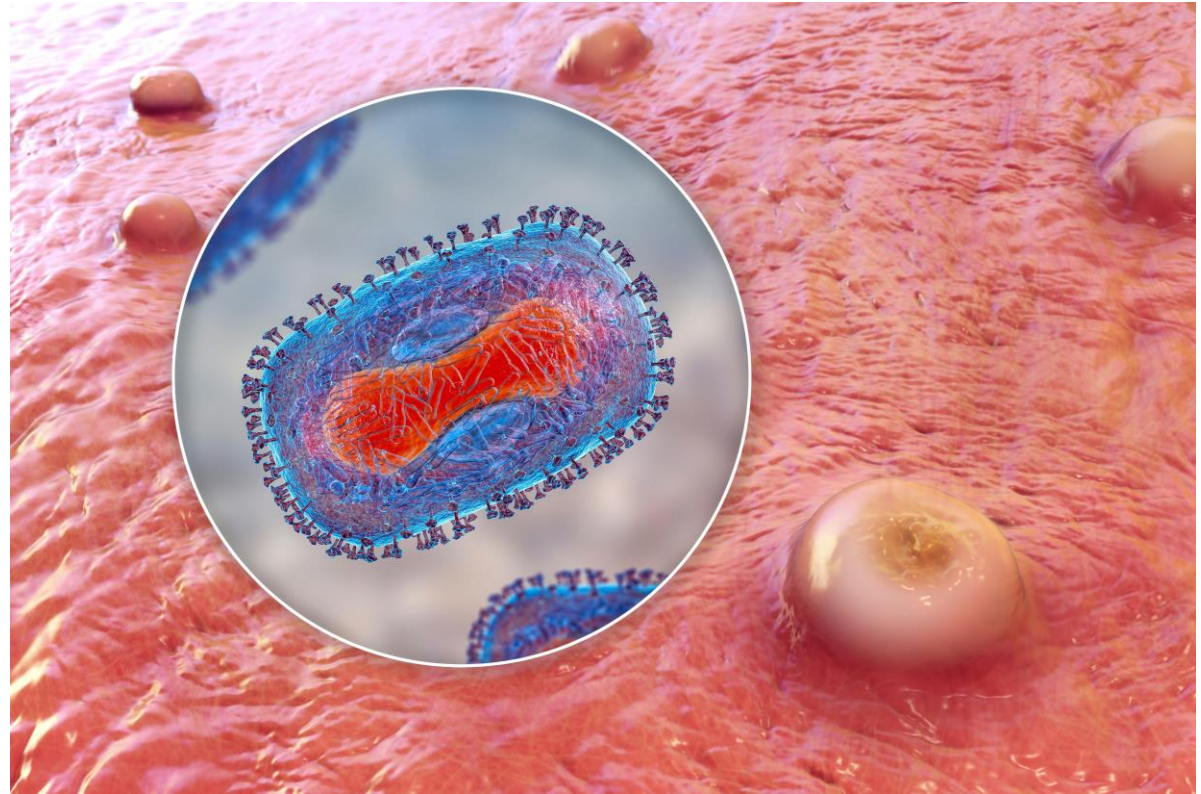


Case recognition and case definition in the multi country outbreak

Prof Chloe Orkin

Queen Mary University of London

I have no disclosures related to monkeypox



Global picture – slide will *definitely* be out of date

Total cumulative number of cases

53K

[VIEW MORE >>](#)

Number of countries reporting at least one case

125

Total cumulative number of cases in WHO European Region

22.9K

Total cumulative number of deaths

18

[VIEW MORE >>](#)

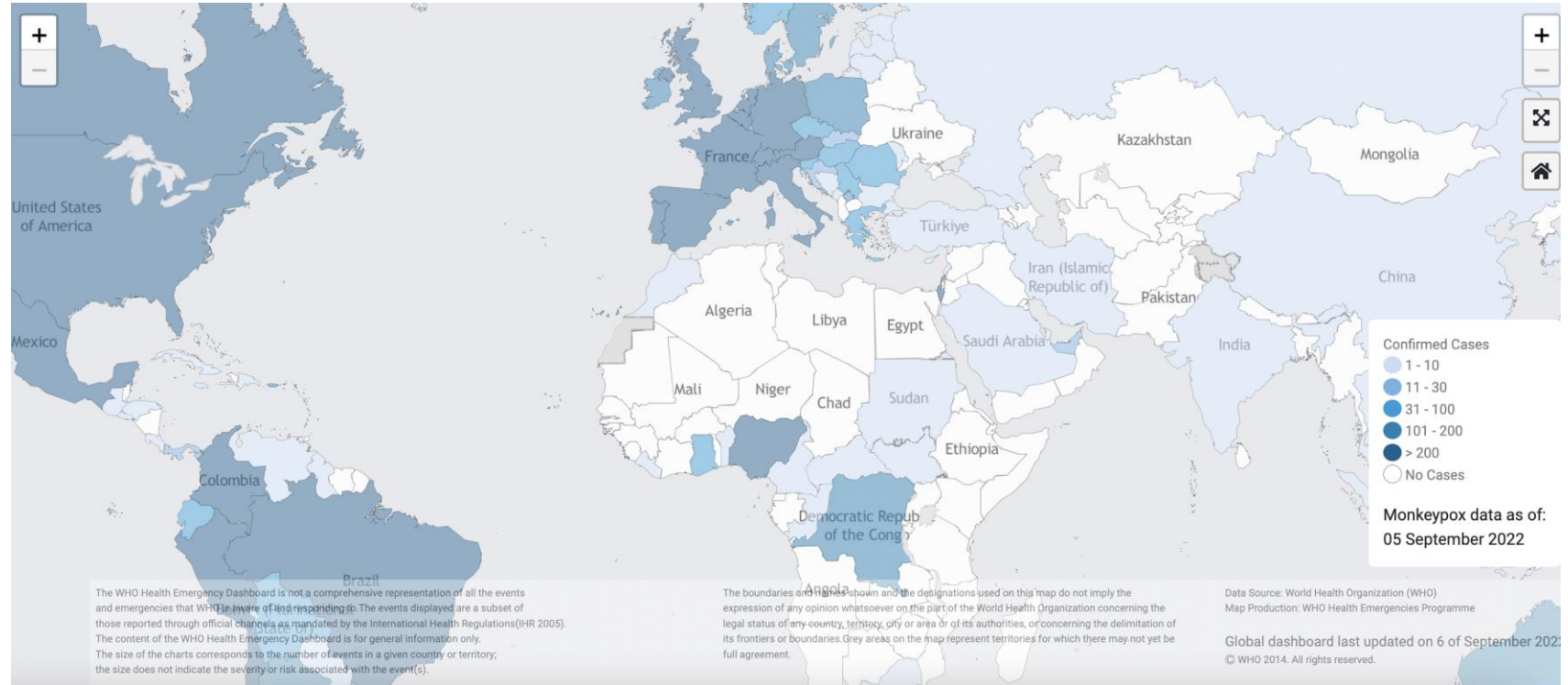
Total cumulative number of cases in WHO Region of the Americas

29.3K

[VIEW MORE >>](#)

Total cumulative number of cases in WHO African Region

521



Key features WHO (Dataset n= 27K)



- 98% male
- 95% GBMSM (1.7% bisexual)
- Median age 36
- 1.8% female
- Most commonly reported setting: party with sexual contact
- Children(<17): 0.6%
- PWH: 45%
- 313 HCW (not all are occupational)

Case profiles

As of September 02 2022

	Reported values ¹		Unknown or Missing Value
	Yes	No	
Men who have sex with men	11923 (95.2%)	606 (4.8%)	33442
HIV-Positive	5576 (44.9%)	6833 (55.1%)	33562
Health worker	313 (4.2%)	7069 (95.8%)	38589
Travel History	1212 (27.9%)	3127 (72.1%)	41632
Sexual Transmission	7822 (91.0%)	777 (9.0%)	37372
Hospitalised ²	1549 (8.4%)	16928 (91.6%)	27494
ICU	9 (0.1%)	8072 (99.9%)	37890

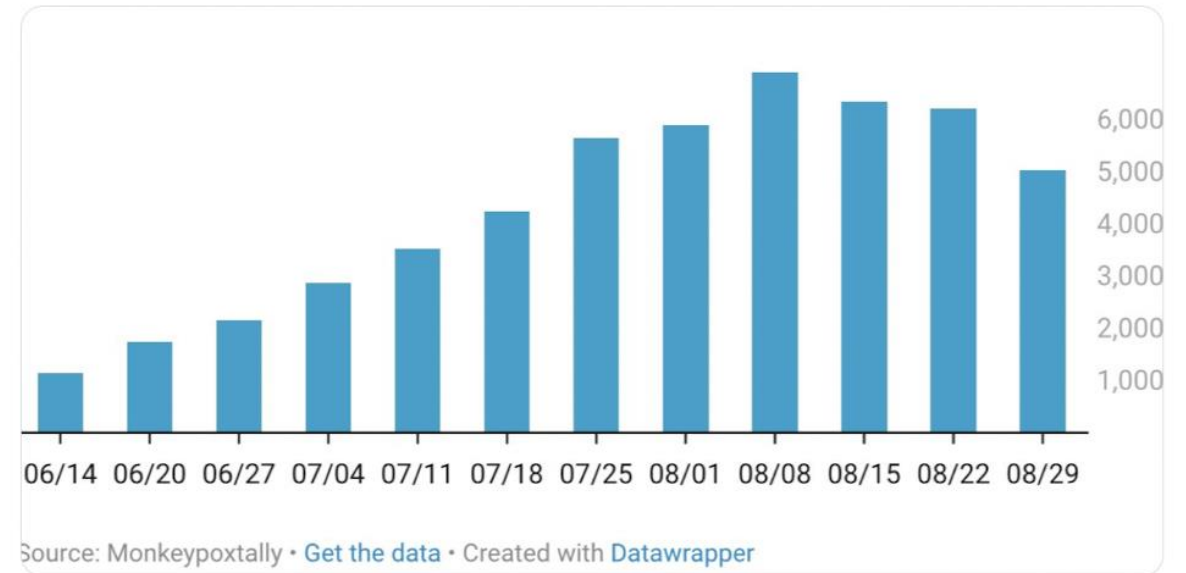
Declining numbers globally

- Overall global decline
- Growing fraction of people with no MSM contact

Monkeypox tally @Monkeypox tally · 2d

This is the third week in a row where we see **monkeypox cases decline** globally and we are seeing a plateauing/ slowing down trend in the last 6 weeks around 5,000-6,000 weekly **cases**.





This week we saw the biggest **decline** in **cases** among the last 3 weeks



Case Definitions May 2022

- Expanded to specify at risk group as sexually active MSM, GBMSM

Clinical symptom/sign case definitions May 2022

	 World Health Organization	 <small>EUROPEAN CENTRE FOR DISEASE PREVENTION AND CONTROL</small>		 UK Health Security Agency
Rash description	An unexplained acute rash or one or more acute skin lesions	An unexplained rash on any part of the body	Deep-seated, well-circumscribed lesion, central umbilication; lesion progression through specific sequential stages—macules, papules, vesicles, pustules, scabs	An unexplained rash on any part of their body
Fever	>38.3°C (101°F)	Fever (usually > 38.5°C)	Not mentioned	>38.5°C
Lymphadenopathy	Lymphadenopathy	Generalised or localised	Not mentioned	Lymphadenopathy
Other	intense headache, back pain, myalgia and intense asthenia	headache, backache, and fatigue	Not mentioned	chills, headache, exhaustion, Myalgia Back pain Asthenia

Clinical symptom/sign case definitions May 2022

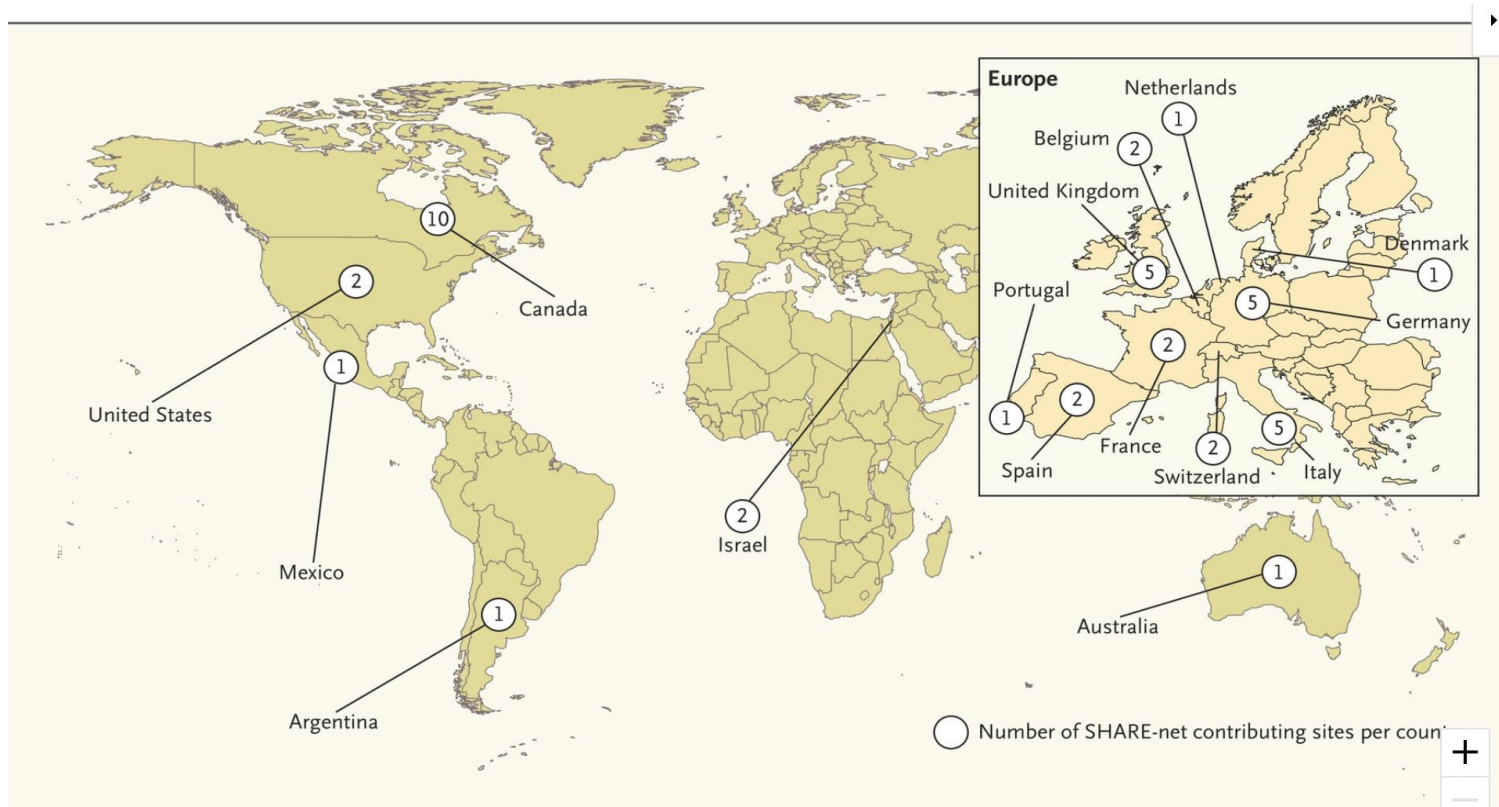
	WHO	ECDC	CDC	UKHSA
Rash description	and acute rash or one or more skin lesions	An unexplained rash on any part of the body	Deep-seated, well-circumscribed lesion, central umbilication, progression through sequential stages, papules, vesicles, scabs	An unexplained rash on any part of their body
Fever	>38.3°C (101°F)	usually >		>38.5°C
Lymphadenopathy	Lymphadenopathy	Characterised by	Not mentioned	Lymphadenopathy
Other	intense headache, myalgia, asthenia	headache, backache, and fatigue	Not mentioned	chills, headache, exhaustion, myalgia Back pain

Mucosal Lesions not mentioned



Monkeypox Virus Infection in Humans across 16 Countries — April–June 2022

John P. Thornhill, M.D., Ph.D., Sapha Barkati, M.D., Sharon Walmsley, M.D., Juergen Rockstroh, M.D., Andrea Antinori, M.D., Luke B. Harrison, M.D., Ph.D., Romain Palich, M.D., Ph.D., Achyuta Nori, M.D., Iain Reeves, M.D., Maximilian S. Habibi, M.D., Ph.D., Vanessa Apea, M.D., M.P.H., Christoph Boesecke, M.D., Linos Vandekerckhove, M.D., Ph.D., Michal Yakubovsky, M.D., Elena Sendagorta, M.D., Ph.D., Jose L. Blanco, M.D., Ph.D., Eric Florence, M.D., Ph.D., Davide Moschese, M.D., Fernando M. Maltez, M.D., Ph.D., Abraham Goorhuis, M.D., Ph.D., Valerie Pourcher, M.D., Ph.D., Pascal Migaud, M.D., Sebastian Noe, M.D., Claire Pintado, M.D., Fabrizio Maggi, M.D., Ph.D., Ann-Brit E. Hansen, M.D., Ph.D., Christian Hoffmann, M.D., Ph.D., Jezer I. Lezama, M.D., Ph.D., Cristina Mussini, M.D., AnnaMaria Cattelan, M.D., Keletso Makofane, M.P.H., Ph.D., Darrell Tan, M.D., Ph.D., Silvia Nozza, M.D., Ph.D., Johannes Nemeth, M.D., Marina B. Klein, M.D., and Chloe M. Orkin, M.D. for the SHARE-net Clinical Group*







4 West London clinics Data covering 14-25/5/22

ARTICLES | ONLINE FIRST

 PDF [2 MB]  Figures  Save  Share  Reprints  Request

Demographic and clinical characteristics of confirmed human monkeypox virus cases in individuals attending a sexual health centre in London, UK: an observational analysis

Nicolò Girometti, MD   Ruth Byrne, MBBS  Margherita Bracchi, MD  Joseph Heskin, MBChB

Alan McOwan, MRCP  Victoria Tittle, MBBS  et al. [Show all authors](#)

Published: July 01, 2022  DOI: [https://doi.org/10.1016/S1473-3099\(22\)00411-X](https://doi.org/10.1016/S1473-3099(22)00411-X)  Check for updates

 PlumX Metrics

Single south London clinic 13 May and 1 July 2022

Research

Clinical features and novel presentations of human monkeypox in a central London centre during the 2022 outbreak: descriptive case series

BMJ 2022 ; 378 doi: <https://doi.org/10.1136/bmj-2022-072410> (Published 28 July 2022)

Cite this as: *BMJ* 2022;378:e072410

Aatish Patel , infectious diseases registrar,



May 11 to June 29, 2022.

Clinical presentation and virological assessment of confirmed human monkeypox virus cases in Spain: a prospective observational cohort study

Eloy José Tarín-Vicente, MD · Andrea Alemany, MD · Manuel Agud-Díos, MD · María Ubals, MD · Clara Suñer, PhD · Andrés Antón, PhD · Maider Arando, PhD · Jorge Arroyo-Andrés, MD · Lorena Calderón-Lozano, MD · Cristina Casañ, PhD · José Miguel Cabrera, MD · Pep Coll, MD · Vicente Descalzo, MD · María Dolores Folgueira, PhD · Jorge N García-Pérez, MD · Elena Gil-Cruz, MD · Borja González-Rodríguez, MD · Christian Gutiérrez-Collar, MD · Águeda Hernández-Rodríguez, PhD · Paula López-Roa, PhD · María de los Ángeles Meléndez, PhD · Julia Montero-Menárguez, MD · Irene Muñoz-Gallego, PhD · Sara Isabel Palencia-Pérez, PhD · Roger Paredes, PhD · Alfredo Pérez-Rivilla, PhD · María Piñana, MD · Nuria Prat, MD · Aída Ramírez, MD · Ángel Rivero, MD · Carmen Alejandra Rubio-Muñiz, MD · Martí Vall, PhD · Kevin Stephen Acosta-Velásquez, MD · An Wang, MD · Cristina Galván-Casas, MD † · Michael Marks, PhD † · Prof Pablo L Ortiz-Romero, PhD † · Oriol Mitjà, PhD †

Show less · Show footnotes

Published: August 08, 2022 · DOI: [https://doi.org/10.1016/S0140-6736\(22\)01436-2](https://doi.org/10.1016/S0140-6736(22)01436-2) · Check for updates

PlumX Metrics



SPAIN 

Demographics

	Global² N=528	West London¹ N=54	South London³ N=197	Spain⁴ N=181
Male	100%	100%	100%	97% (3% female)
MSM	98%	100%	99.5%	92%
Age	38	41	38	37
White	75%	48%	N/R	N/R
PWH	41%	24%	36%	40%
PWH undetectable (VL<200c/mL)	97%	85%	79%	N/R
PWH CD4 count	668	500	664	N/R
Prior small pox vaccination	9%	N/R	N/R	18%

Transmission

	Global² N=528	West London¹ N=54	South London³ N=197	Spain⁴ N=181
Suspected sexual transmission	95%	96%	96%	97%
Number of partners (median)	5	➤ >5%	N/R	6.5
Travel history	28%	46%	27%	14%
Sex on site venue	32%	N/R	N/R	N/R
Known sexual contact with monkeypox	26%	2%	26%	26%
Chemsex	20%	N/R	N/R	31%
Household contact	1%	N/R	N/R	3%
STI	29%	25%	31.5%	17%



Where did they attend?

- The health setting of initial presentation included sexual health, HIV clinics, emergency departments, and dermatology clinics , primary care and private clinics.



Incubation period

- Median incubation period was 7 days (range, 3 to 20)



Clinical Presentation

- 95% of the persons presented with a rash
- **Where:**
 - 73% had anogenital lesions, 55% trunk, arms, or legs the face (25%); and the palms and soles (10%).
- **How many lesions?**
 - Most (64%) had < 10 lesions
 - 54 had a single genital lesion (10%).
- **What does rash look like:**
 - A wide spectrum including macular, pustular, vesicular, and crusted lesions, and lesions in multiple phases were present simultaneously.
 - 58% had lesions that were described as vesiculopustular.

Rash Descriptors

	Global² N=528	West London¹ N=54	South London³ N=197	Spain⁴ N=181
Anogenital rash	73%	94%	88%	78%
Description	58% vestibulo-pustular	Multiple stages	Multiple stages	Pustular 90%
Number of lesions	2-10 in 64%	'Multiple' in 89%	2-10 in 51%	92% <20
Single lesion presentations	10%	11%	11%	N/R
Face	25%	20%	36%	N/R
Body (trunk/limbs)	55%	40%	73%	57%
Hands /feet	10%	20%	28%	60%

A) Evolution of cutaneous lesion in an individual with Human Monkeypox infection.

A1-A3 show facial lesions, B1-B3 shows evolution of a penile lesion and C1-2 shows a lesion on the forehead. PCR status is indicated where available.



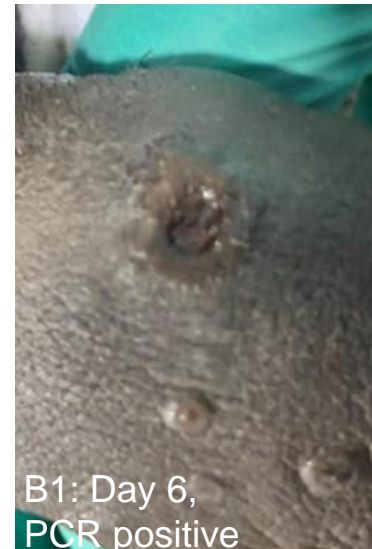
A1: Day 11, PCR positive



A2: Day 21, PCR negative



C1: Day 11



B1: Day 6, PCR positive



B2: Day 21, PCR positive



B3: Day 28, PCR negative



C2: Day 21

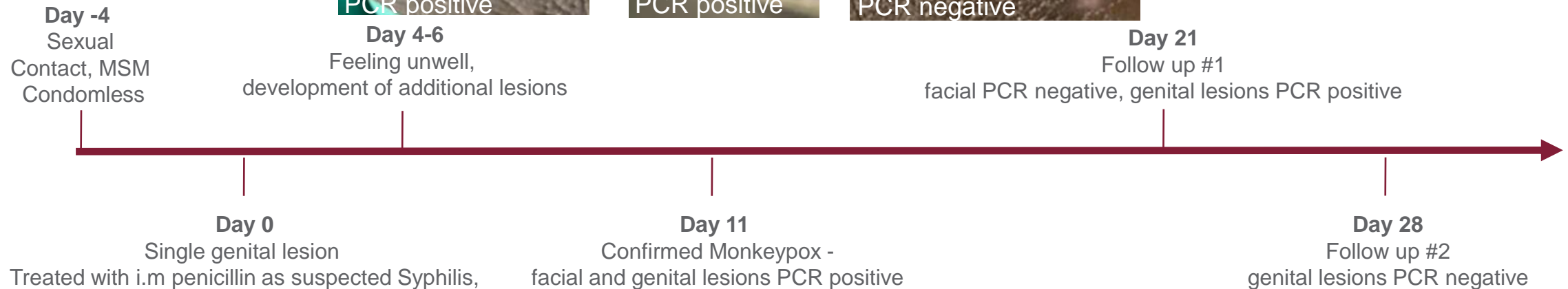


Figure 21: Penile/scrotal lesions in individuals with Human Monkeypox infection.





****New****
mucosal
presentations

- 41% had mucosal lesions
- Anorectal mucosa reported as the **presenting symptom** in **61 persons**
- Oropharyngeal symptoms were reported as the initial symptoms in **26 persons**
- In **3 persons, conjunctival mucosa**
- Penile oedema and urethritis reported by other groups (BMJ Patel group)

Mucosal lesions

	Global² N=528	West London² N=54	South London³ N=197	Spain⁴ N=181
Mucosal lesions	41%	??	??	35%
Oral lesions	5% (as 1 st symptom)	7%	16.8%	25%
Anal lesions	12% (as 1 st symptom)	??	36%	25%
Conjunctival lesions	0.2%	N/R	N/R	N/R
Urethral lesions/ Penile oedema	N/R	N/R	15.7% 31 penile oedema	8%



Noteworthy Oral Presentations

B) Oral and perioral lesions in individuals with Human Monkeypox infection.



23

A. perioral umbilicated lesions. B. Perioral vesicular lesion, day 8, PCR positive. C. Ulcer left corner of the mouth, day 7, PCR positive. D. Tongue ulcer. E. Tongue lesion, day 5, PCR positive. F, G, H. Pharyngeal lesions, day 0, 3 and 21 respectively. PCR positive on day 0 and 3 and negative on day 21.



Noteworthy Ano-rectal Presentations

C) Perianal, anal and rectal lesions in individuals with Human Monkeypox infection.



A. Anal and perianal lesions, day 6. **B, C.** rectal and anal lesions in a single individual **D.** Perianal ulcers. **E.** Anal lesions, PCR positive
F. Umbilicated perianal lesion, day 3. PCR positive. **G.** Umbilicated perianal lesions, day 3. PCR positive. **H.** Perianal ulcer, day 2. PCR positive

Systemic Symptoms

26

	Global² N=528	West London¹ N=154	South London³ N=197	Spain⁴ N=181
Fever	62%	67%	62%	72%
Headache	27%	N/R	25%	53%
Myalgia	31%	30%	32%	N/R
Lymphadenopathy	56%	56%	58%	85%
Arthralgia	N/R	N/R	10%	N/R
Fatigue	41%	67%	23%	N/R
Low Mood	10%	N/R	N/R	N/R

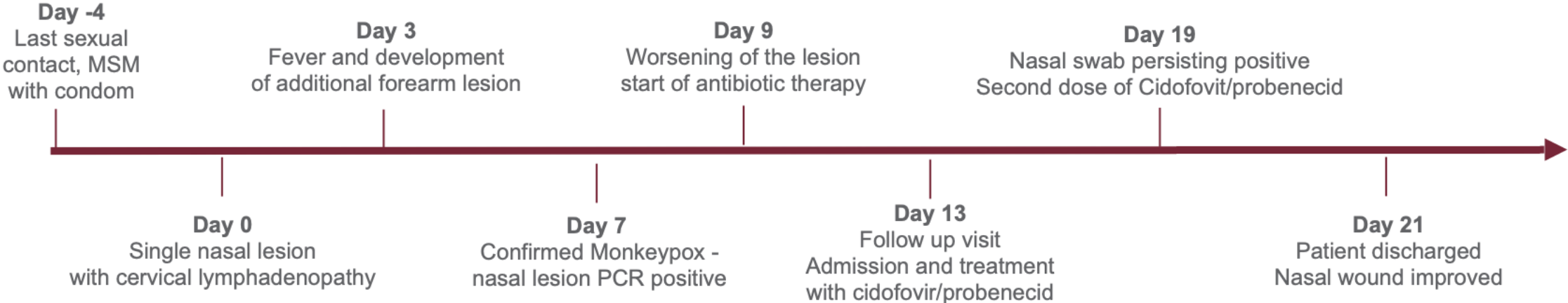
Admissions / complications

	Global ² N=528	West London¹ N=54	South London³ N=197	Spain⁴ N=181
Admissions for medical reasons	57 (11%)	9%	20 (10.%)	2 (<2%) (bacterial cellulitis)
Infections	18	6 (11%)	3 groin 2 tonsillar	6 (3%)
Rectal complications	21	? 5 rectal cellulitis	8	No admissions (25% proctitis)
Oral lesion complications	5	N/R	2	Oral ulcers and tonisllitis (no admissions)
Penile swelling	N/R	7%	5	8% (15)
Other	Epiglottitis Myocarditis x 2 AKI 2	Ano-genital cellulitis Disseminated infection	Urinary retention Paraphimosis Conjunctival lesion 2 Rectal perforation 1 Disseminated lesions 1	Paraphimosis 15(8%) Bacterial skin Abscess 6 3% Exanthem 4%
Received treatment	5%	2% (n=1)	Not mentioned	6
Deaths	0	0	0	0



Noteworthy complicated or
unusual presentations

Figure 15: Evolution of nasal lesions in an individual with Human Monkeypox infection first presented with a single nasal lesion. A1-A5 show evolution of a nasal lesion, B shows a single forearm lesion. PCR status is indicated where available.





Noteworthy bodily
presentations

Figure 23: Bodily Skin lesions in individuals with Human Monkeypox infection.





HIV

- 41% (218) were PWH



HIV

- Median CD4 680 cells/MM³
- 97% VL <200 c/mL and on ART
- No differences in natural history or outcomes observed across entire cohort (see table in supplement)
- We reviewed site of presentation, nature, site and type of rash, extent of rash, mucosal presentations, systemic symptoms, admissions, complications
- However, of the three most significant complications (epiglottitis (1), myocarditis (2), 2 were PWH



Semen

- Found in 29/32 of samples tested

HIV: 40 years of activism drove progress

THE NEW YORK TIMES,
FRIDAY, JULY 3, 1981
A20 L

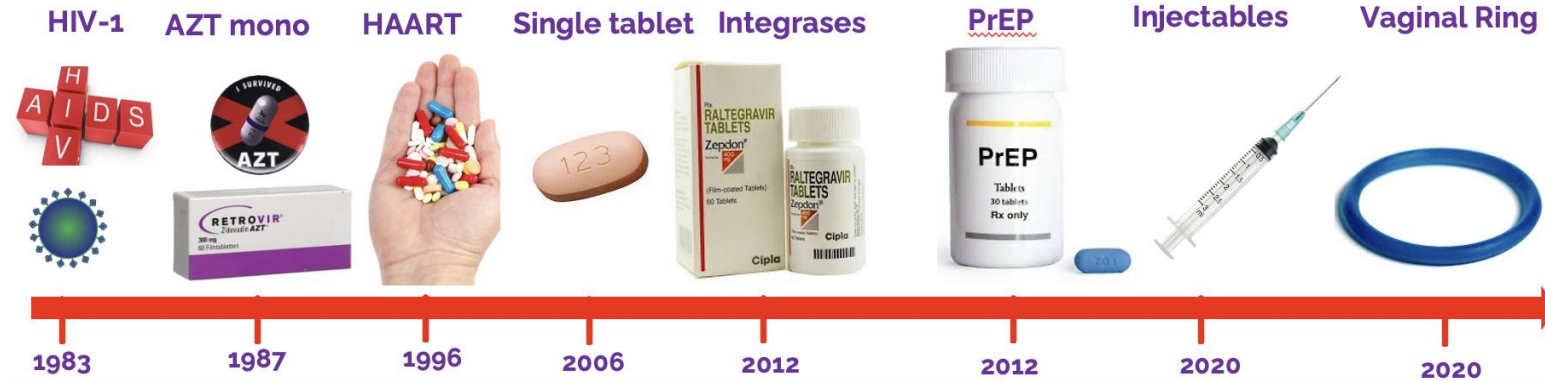
RARE CANCER SEEN IN 41 HOMOSEXUALS

Outbreak Occurs Among Men
in New York and California
— 8 Died Inside 2 Years

By LAWRENCE K. ALTMAN
Doctors in New York and California
have diagnosed among homosexual men
41 cases of a rare and often rapidly fatal
form of cancer. Eight of the victims died
less than 24 months after the diagnosis
was made.
The cause of the outbreak is unknown.



U=U
UNDETECTABLE EQUALS UNTRANSMITTABLE



New monkeypox activists

Harun Tulunay @HarunTulunay · 27/08/2022 ...
I was lucky to have support and understanding from @Positively_UK but government need to support people who are isolating with #monkeypox financially and legally.

islingtontribune.co.uk/article/monkey...

Calls for extra support for those with virus

Friday, 26th August – By [Anna Lamche](#)



pinknews.co.uk
Man who shared monkeypox experience flooded with disgusting, homophobic abuse

A podcast cover for 'WHAT THE POX?' featuring Harun Tulunay as the guest. The cover has a yellow and pink background. The text 'WHAT THE POX?' is in large yellow letters. Below it, it says 'SEASON ONE GUEST: HARUN TULUNAY'. There is a 'Listen now' button with a microphone icon and a search icon. At the bottom, it says 'A QUEER AF PRODUCTION' and 'GIFacast Monkeypox stories and resources without shame or stigma'.

'When will you include include mucosal and single lesions CDC , WHO, ECDC?'

Activism to change case definitions



AIDS2022 Conference , Monkeypox session, July, Montreal Canada

Chloe Orkin @profchloeorkin · 6d

1/ new @WHO case definitions are out!

I worked on the wording with @WHO

They now specify: single & multiple anogenital lesions and mucosal presentations: oral, anal, conjunctival, urethral, penile, vaginal, proctitis.

@QMUL

Identify mucosal lesions to include genital presentations to include oral to prevent further...
groups for the purpose of case definition, to prevent further health systems and to allow effective control and prevention measures.

- In the context of the current outbreak, the inclusion of oral lesions, lesions of the oral cavity, including those also likely to be observed in the development of MPV infection, is a key addition to the case definition for the purpose of case definition, to prevent further health systems and to allow effective control and prevention measures.
- Other mucosal presentations, such as conjunctival, urethral, penile, vaginal, and anal, are also included in the case definition to ensure that all presentations are captured.
- The inclusion of oral lesions, including those also likely to be observed in the development of MPV infection, is a key addition to the case definition for the purpose of case definition, to prevent further health systems and to allow effective control and prevention measures.
- The inclusion of oral lesions, including those also likely to be observed in the development of MPV infection, is a key addition to the case definition for the purpose of case definition, to prevent further health systems and to allow effective control and prevention measures.
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- The inclusion of oral lesions, including those also likely to be observed in the development of MPV infection, is a key addition to the case definition for the purpose of case definition, to prevent further health systems and to allow effective control and prevention measures.

who.int
Surveillance, case investigation and contact tracing for Monkeypox: Interim guidance

3 40 68

'When will you include include mucosal and single lesions CDC?'

New York Times

Dr. Orkin is president of the Medical Women's Federation, past president of the British H.I.V. Association and a governing council member of the International AIDS Society. "I've got a loud voice," she said, "and I'm still finding it difficult to get a response."

Senior members of the W.H.O. responded to Dr. Orkin on Aug. 2, asking to discuss the cases that she and her colleagues had described. The C.D.C. did not reply to Dr. Orkin but added rectal pain and bleeding, along with other new symptoms, to its [guidance to clinicians](#) on Aug. 5.

Dear Dr Walensky... CDC



Chloe Orkin @profchloeorkin · 6d

Thanks for sharing your (great) story again @gOingmad. The @CDCgov case definition still does not specify oral lesions. I remain concerned this will lead to misdiagnosis. People must be tired of me going on but effective activism is incessant advocacy! @dr_demetre @CDCDirector

Madison Muller @gOingmad · 6d

"Case recognition is vital, and we haven't been equipped to actually recognize the disease," @profchloeorkin told me back in July. (Story here: [bloomberg.com/news/articles/...](https://www.bloomberg.com/news/articles/)) Great to see the WHO is now broadening its case definitions. [twit...](#)



Expanded case definition for monkeypox infection

4

14



Chloe Orkin

Saturday, 23 July 2022 at 15:27

To: rwalensky@cdc.gov; Cc: [Carlos del Rio, MD](#)

To protect your privacy, some external images in this message were not downloaded.

[Download external images](#)

Dear Dr Walensky,

I am a clinician in London, past Chair of the British HIV Association and the senior author for the recently published large global case series on monkeypox infection published on Thursday in the NEJM.

<https://www.nejm.org/doi/full/10.1056/>

In our large global case series n=528 from 16 countries, we have described first presentations which are not included in the global case definitions. One example is single genital lesions (~10%). But much more important were primary mucosal presentations with no other skin lesions.

Ano-rectal involvement as the first symptom was reported in 61 people (11.5%) associated with severe anorectal pain, proctitis, tenesmus and/or diarrhoea. Ano-rectal symptoms sometimes led to admission for pain control.

Oropharyngeal symptoms were the initial presentation reported in 26 individuals with pharyngitis, painful swallowing, epiglottitis, and oral or tonsillar lesions. Some required hospitalisation. Conjunctival mucosa and nasal lesions also occurred.

Managing these oral and rectal lesions has necessitated admission and we found these were amongst the most common causes of admission.

We found that people presented to a wide range of places including GPs and emergency departments. These clinicians are less likely to recognise monkeypox than clinicians at sexual health and HIV clinics making the case definition especially important.

In our paper the authors called for an expanded case definition to assist with case recognition to include single lesions, and most importantly, to include mucosal primary presentations without other skin lesions.



Chloe Orkin @profchloeorkin · 1h

The @CDC definition still does not reference oral lesions even after the change on 5/8/22

This is an important omission given the severe tonsillar presentations which can require hospitalisation
[@JohnPThornhill](#) [@Boghuma](#)
[@apoorva_nyc](#) [@dr_demetre](#)




bit.ly/3PGg7js



Case definitions-
changed



Clinical case Definitions August 2022- paraphrased

	 World Health Organization	 ECDC EUROPEAN CENTRE FOR DISEASE PREVENTION AND CONTROL	 CDC	 UK Health Security Agency
Rash description	An explained acute rash, mucosal lesions or lymphadenopathy (swollen lymph nodes). The skin rash may include single or multiple lesions in the ano-genital region or elsewhere on the body.	An unexplained rash on any part of the body	Lesions are firm or rubbery, well-circumscribed, deep-seated, and often develop umbilication (resembles a dot on the top of the lesion). Lesions typically develop simultaneously and evolve together on any given part of the body. The evolution has four stages—macular, papular, vesicular, to pustular—before scabbing over and desquamation.	Unexplained rash on any part of their body Lesions, including but not limited to: genital, ano-genital lesions
Fever/prodrome lymphadenopathy	Acute onset of fever (>38.5°C), headache, myalgia (muscle pain/body aches), back pain, profound weakness or fatigue.	Fever (usually > 38.5°C)	Fever and other prodromal symptoms (e.g., chills, lymphadenopathy, malaise, myalgias, or headache) can occur before rash /after rash/ not occur	Febrile prodrome consists of fever ≥ 38°C, chills, headache, exhaustion, myalgia, arthralgia, backache, lymphadenopathy.
Mucosal lesions	Mucosal lesions may include single or multiple oral, conjunctival, urethral, penile, vaginal, or ano-rectal lesions. Ano-rectal lesions can also manifest as ano-rectal inflammation (proctitis), pain and/or bleeding.	Not specifically mentioned	Rectal symptoms (purulent or bloody stools, rectal pain, or rectal bleeding). Mouth lesions	Oral lesion(s) – for example, ulcers, nodules Proctitis – for example anorectal pain, bleeding

<https://www.gov.uk/guidance/monkeypox-case-definitions> <https://www.cdc.gov/poxvirus/monkeypox/clinicians/case-definition.html>

<https://www.who.int/publications/i/item/WHO-MPX-Surveillance-2022.2> <https://www.canada.ca/en/public-health/services/diseases/monkeypox/health-professionals/national-case-definition.html>

<https://www.ecdc.europa.eu/en/news-events/epidemiological-update-monkeypox-multi-country-outbreak-15-june>



Thank you