# Summary of Current Evidence and Information— Mpox in Donor Screening and Transplantation

#### **Table of Contents**

Purpose	1
Methods	1
Discussion	2
Background- Current outbreak in US	
Routes of Transmission	
Infectivity Period	3
Screening Considerations: Deceased Donor Considerations	
Safety of the OPO, Recovery Team and Transplant Programs	
Screening Considerations: Living Donors, Information on Impact on Living Donor S	Safety 4
Testing Considerations: Available testing, how to access, specimen information	•
Bibliography	

# **Purpose**

This document is a summary of evidence and information regarding donor screening for Mpox virus and considerations for organ acceptance from donors with a history of Mpox based on peer-reviewed literature. This resource is subject to revision as new data accumulate and will be reviewed at least annually or as needed for currency. The overarching objective of this document is to compile the latest information known for minimizing the risk of donor-derived Mpox while maximizing donor utilization.

# **Methods**

The OPTN Ad Hoc Disease Transmission Advisory Committee (DTAC) and relevant stakeholders from the Centers for Disease Control and Prevention (CDC), American Society of Transplantation (AST), American Society of Transplant Surgeons (ASTS), Association of Organ Procurement Organizations (AOPO), American Association of Tissue Bank (AATB), US Food and Drug Administration (FDA), and Health Resources & Services Administration (HRSA) reviewed published literature. Specifically, DTAC and

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relevant stakeholders assessed the available evidence as it relates to living and deceased donor screening and testing and recovery of organs from living or deceased donors with a history of Mpox.

#### **Discussion**

#### Background: Current outbreak in US

The US Department of Health and Human Services declared Mpox (formerly monkeypox) to be a public health emergency on August 4, 2022. The CDC, in conjunction with state and local public health partners, continues to confirm cases of Mpox and provide updated <u>case counts by state</u>. As of January 10, 2024, over 31,600 cases have been reported in the US, although the number of daily cases reported has steadily declined since September 2022. In addition to case reporting, the National Wastewater Surveillance System monitors for Mpox virus detection in wastewater as an early warning of Mpox activity.

Among Mpox cases reported to the CDC, and with data available, through October 21, 2022, the median age of infected persons is 34 years (range <1 to 89 years), and 97% of cases have been detected in men. Exclusive male-to-male close intimate or sexual contact is reported in at least 40% of cases. Data also indicate that racial and ethnic minority groups are disproportionately affected.<sup>iii</sup>

While the majority of US Mpox cases have occurred in men who have sex with men, <u>anyone in close</u> <u>personal contact with someone who has Mpox is at risk for infection, regardless of gender identity or sexual orientation.</u>

The CDC is continuing to evaluate and address prevention and treatment protocols. Reduced risk for Mpox has been documented with 1 or 2 doses of the JYNNEOS vaccine, with two doses providing the best protection, compared to unvaccinated persons. The CDC recommends the 2-dose series of JYNNEOS vaccine for any adult at epidemiologic risk for Mpox infection.

#### Routes of Transmission

- There have been no confirmed transmissions of Mpox through organ or tissue transplantation or blood transfusions to date. The risk of transmission of Mpox through organ/tissue/blood donation is currently unknown.
- Mpox has been detected in respiratory secretions, blood, urine, seminal fluid, and tissue abscess fluid of infected persons.<sup>v,vi</sup>
- Mpox can spread in the following ways:
  - o Direct contact with the rash, scabs, or body fluids of a person with Mpox infection.
  - Touching objects, fabrics (clothing bedding or towels), and surfaces that have been used by a person with Mpox.
  - o Contact with respiratory secretions of a person infected with Mpox.vii
- Infected animals can spread Mpox to humans via scratch or animal bite.
  - The natural reservoir of Mpox has not been identified, although rodents are the most likely source. VIII
- Consumption of meat from infected animals may also transmit Mpox.
- Household sampling from a single case of a confirmed infected individual 20 days after symptom onset confirmed Mpox DNA on multiple high-contact surfaces. However, no viable virus was

detected via culture, with normal household cleaning but no specific decontamination performed.<sup>x</sup> Additionally, viral detection of replication competent virus may precede symptom onset by as many as 4 days<sup>xxiv</sup>. Additionally, viral detection of replication competent virus may precede symptom onset by as many as 4 days.

#### Viral Detection and Infectivity

- An infected person is infectious via skin-to-skin contact for 2-4 weeks beginning from the start of symptoms until the skin rash has scabbed off and a fresh layer of intact skin has formed.xi
- Because Mpox has been detected in blood, tissue, and body fluids, it could be potentially transmitted by organ and tissue transplantation and blood transfusion if the donor is actively infectious at the time of organ donation.
- There is limited information on the duration of Mpox viremia.
- One study described the detection of Mpox DNA shedding in upper respiratory tract and blood for at least 3 weeks after initial infection. The clinical implications of these findings in the context of transplantation are not yet known.
- In animal models, Mpox DNA has been detected in ocular, oral, nasal, fecal and blood samples starting at day 3 and up to 28-days post-inoculation. Viable (i.e., infectious) virus was detected from ocular, oral, nasal, and fecal secretions up to 21-days post-inoculation. The same study identified a peak of blood Mpox DNA between days 6-15 post-inoculation, but virus viability was not assessed.
- A retrospective study included seven patients in the UK with confirmed Mpox, admitted between August 2018 and September 2021. Mpox DNA was detected up to 31, 45, 23 days from blood, upper respiratory and urine samples, respectively. There was no testing to determine if the virus was viable.xiii
- A recent <u>study</u> assessed Mpox viral clearance with serial sample collection across multiple body sites (skin, oropharynx, blood, rectum, semen, vagina). Modeling suggested that 90% of patients will clear PCR positivity by 41 (skin) and 39 (semen) days. However, most samples with positive viral cultures were collected before day 15 of illness.
- In summary, Mpox DNA can be identified from multiple sites within the body but the duration of replication-competent virus based on current evidenceremains under investigation.

## Screening Considerations: Deceased Donors

- Based on the biology of disease, clinical data, and data in animal studies, there may be a risk of disease transmission to recipients of organs from donors with active Mpox.
- Standard physical examination of a potential donor will determine if they present with regional lymph node swelling or a rash, which is a common presentation for Mpox<sup>xiv,xv</sup>
  - o In a donor with suspected Mpox lesions, infectious disease consultation may be appropriate to guide further evaluation and management.
- Current donor medical history screening questions in the universal Donor Risk Assessment Interview (uDRAI) can obtain information on risk factors for Mpox exposure or infection.
  - On November 6, 2023, the <u>American Association of Tissue Banks</u> (AATB) recommended discontinuation of additional donor risk assessment questions used to screen for the risk of Mpox unless there is another significant outbreak.
- Persons infected with Mpox may remain asymptomatic for up to 21 days before showing symptoms.
   The risk of transmission from infected, asymptomatic individuals is unknown.xvi
- For donors with a history of Mpox infection, if the donor is asymptomatic and all of the scabs on the skin lesions are healed the likelihood that they have any replication-competent virus is low.

- While active lesions are present or incompletely healed, the risk of transmission to potential
  recipients may be increased. DNA has been isolated from different sample types (blood, upper
  respiratory and urine sample types) after lesions have healed, but infectivity has not been
  established.
- The Modified virus Ankara vaccine (MVA) (JYNNEOS) is the primary vaccine being used in the US during this outbreak. It is made from a highly attenuated, nonreplicating vaccinia virus. Thus, donors receiving this vaccine are not at risk of transmitting vaccine-strain Vaccinia virus.
- ACAM2000 is a replication-competent smallpox vaccine. Vaccinia virus can be recovered from the skin at the vaccination site for a mean duration of 7.8 days, with a range of 0 to 18 days. Peak immunity is expected to be reached 4 weeks after administration.xviii
  - O Viremia is unlikely once the immune response is initiated.xviii
  - O Viremia has been more readily detected in people with moderate to severe complications of vaccinia virus infection. These complications include generalized vaccinia, eczema vaccinatum and progressive vaccinia. xviii
- Evidence suggests that the decision to recover organs could include the following:
  - The recipient risk of mortality or further complications while delaying transplantation and remaining on the waitlist.
  - o Current unknown outcomes.
  - Infectious disease experts can offer subject matter expertise when accepting organs from these donors.xvi
- CDC provides <u>recommendations for healthcare worker vaccination for Mpox.</u>
- The FDA has released considerations for tissue donation regarding Mpox.

# Safety of the OPO, Recovery Team and Transplant Programs

- Donors with unrecognized Mpox infection are possible sources of transmission to healthcare personnel, including OPO staff and recovery team members.
- Risk of transmission to healthcare personnel is reduced by standard precautions, personal protective
  equipment (to include an N95 respirator and contact precautions), environmental control tactics,
  and waste management; for more specifics, see <a href="here">here</a>.
- Potential donors with prior Mpox infection are no longer contagious via skin contact once skin lesions are completely re-epithelialized.
- There have been few reports of transmission to healthcare personnel, but currently these are rare.xix,xx
- CDC provides recommendations for post-exposure vaccination for healthcare workers.

# Screening Considerations: Living Donors, Impact on Living Donor Safety

- Although most cases of Mpox have been diagnosed in men who have sex with men, anyone in close contact with a person infected with Mpox is at risk for infection, regardless of gender identity or sexual orientation.
- Screening and educating potential living donors for possible exposure or illness is a strategy to reduce the risk of transmitting Mpox to recipients.
  - o Educate potential living donors regarding increased risk behaviors for contracting Mpox.

- Examination of the genitals and perianal area could identify Mpox lesions.
- The optimal timing of living donation after recovering from Mpox is unknown. Factors to be considered could include:
  - Safety of the living donor. The risk of surgical complications following Mpox is unknown.
  - O Risk of transmission to the recipient. While active lesions are present or incompletely healed, the risk of transmission to potential recipients may be increased. DNA has been isolated from different sample types (blood, upper respiratory and urine sample types) after lesions have healed, but infectivity has not been established.\*\*
  - Waiting list mortality of the intended recipient.
- Individuals infected with Mpox may be asymptomatic for up to 21 days after their exposure.
   Obtaining a careful history to identify <u>contact exposures to Mpox or engagement in activities that have increased risk of infection</u> is crucial.
  - O Potential living donors who have been exposed to Mpox could consider deferring donation until 21 days following their last exposure while monitoring for symptoms. The risk of such wait time should be weighed against the morbidity and mortality risk for the potential recipient.
- Potential living donors who live in the same household as a person infected with Mpox could consider self-quarantining prior to donation to reduce the risk of additional exposure.

# Testing Considerations: Available testing, accessibility, specimen information

- Testing may be coordinated through the <u>Laboratory Response Network</u>, <u>local public health</u> <u>departments</u>, and <u>certain commercial laboratories</u>.
- Nucleic acid testing (NAT) of skin lesions may provide the most definitive results for Mpox.
- There is significant cross-reactivity between orthopoxvirus antibodies, so positive serologic tests do not definitively indicate exposure to Mpox.xxiii
- The CDC has <u>published guidance on preparation and collection of specimens</u>.

### **Bibliography**

- AATB. "Physicians Council Provides Information on Monkeypox Virus (MPXV)." Monkeypox Announcement 8.15.22. American Association of Tissue Banks, August 15, 2022. https://www.aatb.org/sites/default/files/2022/Monkeypox%20Announcement%208.15.22.pdf.
- AATB. "AATB Physicians Council Recommends Discontinued Use of SARS-CoV-2/COVID-19 and Mpox Virus (MPXV) Addendums." American Association of Tissue Banks, November 6, 2023. https://www.aatb.org/bulletin-23-7
- Adler H, Gould S, Hine P, et al. Clinical features and management of human monkeypox: a retrospective observational study in the UK. *Lancet Infect Dis.* Aug 2022;22(8):1153-1162. doi:10.1016/S1473-3099(22)00228-6.
- Adler, Gould S, Hine P. Clinical features and management of human monkeypox: a retrospective observational study in the UK. Lancet Infect Dis. 2022 Aug;22(8):1153-1162. doi: 10.1016/S1473-3099(22)00228-6.
- Alakunle E, Moens U, Nchinda G, Okeke MI. Monkeypox Virus in Nigeria: Infection Biology, Epidemiology, and Evolution. Viruses. 2020 Nov 5;12(11):1257. doi: 10.3390/v12111257.

- Assistant Secretary for Public Affairs (ASPA). "HHS Expanding Monkeypox Testing Capacity to Five Commercial Laboratory Companies." HHS.gov. U.S. Department of Health and Human Services, August 17, 2022. https://www.hhs.gov/about/news/2022/06/22/hhs-expanding-monkeypox-testing-capacity-five-commercial-laboratory-companies.html.
- Assistant Secretary for Public Affairs (ASPA). "Biden-Harris Administration Bolsters Monkeypox Response; HHS Secretary Becerra Declares Public Health Emergency." HHS.gov. U.S. Department of Health and Human Services, August 17, 2022. https://www.hhs.gov/about/news/2022/08/04/biden-harris-administration-bolsters-monkeypox-response-hhs-secretary-becerra-declares-public-health-emergency.html.
- CDC. "ACAM2000 Vaccine." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 21, 2022. https://www.cdc.gov/poxvirus/monkeypox/interim-considerations/acam2000-vaccine.html.
- CDC. "Clinical Recognition." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, August 23, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/clinical-recognition.html.
- CDC. Clinician FAQs. https://www.cdc.gov/poxvirus/monkeypox/clinicians/faq.html. Last updated August 11, 2022. Last Accessed August 17, 2022.
- CDC. "Guidelines for Collecting and Handling Specimens for Mpox Testing." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, September 20, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/prep-collection-specimens.html.
- CDC. "How It Spreads." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 8, 2022. https://www.cdc.gov/poxvirus/monkeypox/if-sick/transmission.html.
- CDC. "If You Are Sick." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, August 11, 2022. https://www.cdc.gov/poxvirus/monkeypox/if-sick/index.html.
- CDC. "Infection Control: Healthcare Settings." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 31, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/infection-control-healthcare.html.
- CDC. "Information for Laboratory Personnel." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, Updated September 2, 2022. https://www.cdc.gov/poxvirus/monkeypox/lab-personnel/index.html.
- CDC. "Laboratory Procedures." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 6, 2022. https://www.cdc.gov/poxvirus/monkeypox/lab-personnel/lab-procedures.html.
- CDC. "Monitoring and Risk Assessment for Persons Exposed in the Community." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, Updated August 11, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/monitoring.html.
- CDC. "Preparation and Collection of Specimens." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, Updated August 26, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/prep-collection-specimens.html.
- CDC. "Signs and Symptoms." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 9, 2022. https://www.cdc.gov/poxvirus/monkeypox/symptoms/index.html.
- CDC. "Technical Report 4: Multi-National MPOX Outbreak, United States, 2022." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 27, 2022. https://www.cdc.gov/poxvirus/monkeypox/cases-data/technical-report/report-4.html.

- CDC. "2022 Outbreak Cases and Data." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, September 30, 2022. https://www.cdc.gov/poxvirus/monkeypox/response/2022/index.html.
- CDC "U.S. Mpox Wastewater Data." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, January 24, 2024.
- https://www.cdc.gov/nwss/wastewater-surveillance/mpox-data.html
- CDC. "2022 U.S. Map & Case Count." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, Updated August 10, 2022. https://www.cdc.gov/poxvirus/monkeypox/response/2022/us-map.html.
- Center for Biologics Evaluation and Research. "Info for HCT/P Establishments on Monkeypox and HCT/P Donation." U.S. Food and Drug Administration. FDA, August 29, 2022. https://www.fda.gov/vaccines-blood-biologics/safety-availability-biologics/important-information-human-cell-tissue-and-cellular-and-tissue-based-product-hctp-establishments-0.
- CSTE. "Jurisdiction after Hours/Epi-on-Call Phone Number Infectious Disease ..." Kendo UI Grid Export. Council of State and Territorial Epidemiologists. Accessed September 6, 2022. https://cdn.ymaws.com/www.cste.org/resource/resmgr/poc/epioncall\_update1622.pdf.
- Di Giulio, Daniel B., and Paul B. Eckburg. "Human monkeypox: an emerging zoonosis." *The Lancet infectious diseases* 4, no. 1 (2004): 15-25. https://doi.org/10.1016/S1473-3099(03)00856-9
- ECDC. "Factsheet for Health Professionals on Monkeypox." European Centre for Disease Prevention and Control. European Centre for Disease Prevention and Control, August 19, 2022. https://www.ecdc.europa.eu/en/all-topics-z/monkeypox/factsheet-health-professionals.
- Epidemiological, clinical and virological characteristics of four cases of monkeypox support transmission through sexual contact, Italy, May 2022. Antinori, Andrea and Mazzotta, Valentina and Vita, Serena and Carletti, Fabrizio and Tacconi, Danilo and Lapini, Laura Emma and D'Abramo, Alessandra and Cicalini, Stefania and Lapa, Daniele and Pittalis, Silvia and Puro, Vincenzo and Rivano Capparuccia, Marco and Giombini, Emanuela and Gruber, Cesare Ernesto Maria and Garbuglia, Anna Rosa and Marani, Alessandra and Vairo, Francesco and Girardi, Enrico and Vaia, Francesco and Nicastri, Emanuele and the INMI Monkeypox Group, Eurosurveillance, 27, 2200421 (2022), https://doi.org/10.2807/1560-7917.ES.2022.27.22.2200421
- FDA. "Recommendations for Deferral of Donors and Quarantine and Retrieval of Blood and Blood Products in Recent Recipients of Smallpox Vaccine (Vaccinia Virus) and Certain Contacts of Smallpox Vaccine Recipients." Guidance for Industry. Food and Drug Administration. Accessed September 6, 2022. https://www.fda.gov/media/124362/download.
- Fenner F, Henderson DA, Arita I, et al. Smallpox and its Eradication. Geneva: World Health Organization; 1988:307.
- Hutson CL, Lee KN, Abel J. Monkeypox zoonotic associations: insights from laboratory evaluation of animals associated with the multi-state US outbreak. Am J Trop Med Hyg. 2007 Apr;76(4):757-68.
- Hutson CL, Olson VA, Carroll DS, et al. A prairie dog animal model of systemic orthopoxvirus disease using West African and Congo Basin strains of monkeypox virus. J Gen Virol. 2009 Feb;90(Pt 2):323-333. doi: 10.1099/vir.0.005108-0.
- Lapa, D., Carletti, F., Mazzotta, V., Matusali, G., Pinnetti, C., Meschi, S., Gagliardini, R., Colavita, F., Mondi, A., Minosse, C., Scorzolini, L., Cicalini, S., Maffongelli, G., Specchiarello, E., Camici, M., Bettini, A., Baldini, F., Francalancia, M., Mizzoni, K., ... Maggi, F. (2022). Monkeypox virus isolation from a semen sample collected in the early phase of infection in a patient with prolonged seminal viral shedding. *The Lancet Infectious Diseases*. https://doi.org/10.1016/S1473-3099(22)00513-8

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Mendoza R, Petras JK, Jenkins P, et al. Monkeypox Virus Infection Resulting from an Occupational Needlestick — Florida, 2022. MMWR Morb Mortal Wkly Rep 2022;71:1348–1349. DOI: http://dx.doi.org/10.15585/mmwr.mm7142e2

- Nuzzo JB, Borio LL, Gostin LO. The WHO Declaration of Monkeypox as a Global Public Health Emergency. JAMA. 2022 Jul 27. doi: 10.1001/jama.2022.12513.
- Payne AB, Ray LC, Cole MM, et al. Reduced Risk for Mpox After Receipt of 1 or 2 Doses of JYNNEOS Vaccine Compared with Risk Among Unvaccinated Persons 43 U.S. Jurisdictions, July 31—October 1, 2022. MMWR Morb Mortal Wkly Rep 2022;71:1560–1564. DOI: http://dx.doi.org/10.15585/mmwr.mm7149a5
- Pfeiffer JA, Collingwood A, Rider LE, et al. High-Contact Object and Surface Contamination in a Household of Persons with Monkeypox Virus Infection Utah, June 2022. MMWR Morb Mortal Wkly Rep 2022;71:1092-1094. DOI: http://dx.doi.org/10.15585/mmwr.mm7134e1.
- Rao AK, Petersen BW, Whitehill F, et al. Use of JYNNEOS (Smallpox and Monkeypox Vaccine, Live, Nonreplicating) for Preexposure Vaccination of Persons at Risk for Occupational Exposure to Orthopoxviruses: Recommendations of the Advisory Committee on Immunization Practices United States, 2022. MMWR Morb Mortal Wkly Rep 2022;71:734–742. DOI: http://dx.doi.org/10.15585/mmwr.mm7122e1
- Suñer C, Ubals M, Tarín-Vicente EJ, Mendoza A, Alemany A, Hernández-Rodríguez Á, Casañ C, Descalzo V, Ouchi D, Marc A, Rivero À, Coll P, Oller X, Miguel Cabrera J, Vall-Mayans M, Dolores Folgueira M, Ángeles Melendez M, Agud-Dios M, Gil-Cruz E, Paris de Leon A, Ramírez Marinero A, Buhiichyk V, Galván-Casas C, Paredes R, Prat N, Sala Farre MR, Bonet-Simó JM, Farré M, Ortiz-Romero PL, Clotet B, García-Patos V, Casabona J, Guedj J, Cardona PJ, Blanco I; Movie Group; Marks M, Mitjà O. Viral dynamics in patients with monkeypox infection: a prospective cohort study in Spain. Lancet Infect Dis. 2022 Dec 12:S1473-3099(22)00794-0. doi: 10.1016/S1473-3099(22)00794-0. Epub ahead of print. PMID: 36521505.
- Vaughan A, Aarons E, Astbury J, Brooks T, Chand M, Flegg P, Hardman A, Harper N, Jarvis R, Mawdsley S, McGivern M, Morgan D, Morris G, Nixon G, O'Connor C, Palmer R, Phin N, Price DA, Russell K, Said B, Schmid ML, Vivancos R, Walsh A, Welfare W, Wilburn J, Dunning J. Human-to-Human Transmission of Monkeypox Virus, United Kingdom, October 2018. Emerg Infect Dis. 2020 Apr;26(4):782-785. doi: 10.3201/eid2604.191164. Epub 2020 Apr 17. PMID: 32023204; PMCID: PMC7101111.
- WHO. "Monkeypox." World Health Organization. World Health Organization, May 19, 2022. https://www.who.int/news-room/fact-sheets/detail/monkeypox.
- WHO. "Monkeypox Outbreak Toolbox." World Health Organization. World Health Organization. Accessed September 6, 2022. https://www.who.int/emergencies/outbreak-toolkit/disease-outbreak-toolboxes/monkeypox-outbreak-toolbox.

<sup>&</sup>lt;sup>1</sup> Assistant Secretary for Public Affairs (ASPA). "Biden-Harris Administration Bolsters Monkeypox Response; HHS Secretary Becerra Declares Public Health Emergency." HHS.gov. U.S. Department of Health and Human Services, August 17, 2022. https://www.hhs.gov/about/news/2022/08/04/biden-harris-administration-bolsters-monkeypox-response-hhs-secretary-becerradeclares-public-health-emergency.html.

<sup>&</sup>quot;CDC. "2022-2023 Outbreak Cases and Data." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, January 16, 2024. https://www.cdc.gov/poxvirus/monkeypox/response/2022/index.html.

iii CDC. "Technical Report 4: Multi-National MPOX Outbreak, United States, 2022." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 27, 2022. https://www.cdc.gov/poxvirus/monkeypox/cases-data/technical-report/report-4.html.

<sup>&</sup>lt;sup>iv</sup> Payne AB, Ray LC, Cole MM, et al. Reduced Risk for Mpox After Receipt of 1 or 2 Doses of JYNNEOS Vaccine Compared with Risk Among Unvaccinated Persons — 43 U.S. Jurisdictions, July 31–October 1, 2022. MMWR Morb Mortal Wkly Rep 2022;71:1560–1564. DOI: http://dx.doi.org/10.15585/mmwr.mm7149a5

<sup>&</sup>lt;sup>v</sup> Lapa, D., Carletti, F., Mazzotta, V., Matusali, G., Pinnetti, C., Meschi, S., Gagliardini, R., Colavita, F., Mondi, A., Minosse, C., Scorzolini, L., Cicalini, S., Maffongelli, G., Specchiarello, E., Camici, M., Bettini, A., Baldini, F., Francalancia, M., Mizzoni, K., ... Maggi, F. (2022). Monkeypox virus isolation from a semen sample collected in the early phase of infection in a patient with prolonged seminal viral shedding. The Lancet Infectious Diseases. https://doi.org/10.1016/S1473-3099(22)00513-8

vi Antinori Andrea, Mazzotta Valentina, Vita Serena, Carletti Fabrizio, Tacconi Danilo, Lapini Laura Emma, D'Abramo Alessandra, Cicalini Stefania, Lapa Daniele, Pittalis Silvia, Puro Vincenzo, Rivano Capparuccia Marco, Giombini Emanuela, Gruber Cesare Ernesto

Maria, Garbuglia Anna Rosa, Marani Alessandra, Vairo Francesco, Girardi Enrico, Vaia Francesco, Nicastri Emanuele, the INMI Monkeypox Group. Epidemiological, clinical and virological characteristics of four cases of monkeypox support transmission through sexual contact, Italy, May 2022. Euro Surveill. 2022;27(22):pii=2200421. https://doi.org/10.2807/1560-7917.ES.2022.27.22.2200421

vii CDC. "How It Spreads." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 8, 2022. https://www.cdc.gov/poxvirus/monkeypox/if-sick/transmission.html.

viii Di Giulio, Daniel B., and Paul B. Eckburg. "Human monkeypox: an emerging zoonosis." The Lancet infectious diseases 4, no. 1 (2004): 15-25. https://doi.org/10.1016/S1473-3099(03)00856-9

<sup>ix</sup> WHO. "Monkeypox." World Health Organization. World Health Organization, May 19, 2022. https://www.who.int/newsroom/fact-sheets/detail/monkeypox.

<sup>X</sup> Pfeiffer JA, Collingwood A, Rider LE, et al. High-Contact Object and Surface Contamination in a Household of Persons with Monkeypox Virus Infection — Utah, June 2022. MMWR Morb Mortal Wkly Rep 2022;71:1092-1094. DOI:

http://dx.doi.org/10.15585/mmwr.mm7134e1.<sup>Xi</sup> CDC. Clinician FAQs. Centers for Disease Control and Prevention. Centers for Disease Control and Prevention. https://www.cdc.gov/poxvirus/monkeypox/clinicians/faq.html, Updated August 11, 2022.

xii Hutson CL, Lee KN, Abel J. Monkeypox zoonotic associations: insights from laboratory evaluation of animals associated with the multi-state US outbreak. Am J Trop Med Hyg. 2007 Apr;76(4):757-68.

xiii Adler, Gould S, Hine P. Clinical features and management of human monkeypox: a retrospective observational study in the UK. Lancet Infect Dis. 2022 Aug;22(8):1153-1162. doi: 10.1016/S1473-3099(22)00228-6.

xiv CDC. "Signs and Symptoms." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 9, 2022. https://www.cdc.gov/poxvirus/monkeypox/symptoms/index.html.

xV Alakunle E, Moens U, Nchinda G, Okeke MI. Monkeypox Virus in Nigeria: Infection Biology, Epidemiology, and Evolution. Viruses. 2020 Nov 5;12(11):1257. doi: 10.3390/v12111257.

xvi CDC. "Infection Control: Healthcare Settings." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 31, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinicians/infection-control-healthcare.html.

xvii Rao AK, Petersen BW, Whitehill F, et al. Use of JYNNEOS (Smallpox and Monkeypox Vaccine, Live, Nonreplicating) for Preexposure Vaccination of Persons at Risk for Occupational Exposure to Orthopoxviruses: Recommendations of the Advisory Committee on Immunization Practices — United States, 2022. MMWR Morb Mortal Wkly Rep 2022;71:734–742. DOI: http://dx.doi.org/10.15585/mmwr.mm7122e1

xviii CDC. "ACAM2000 Vaccine." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, October 21, 2022. https://www.cdc.gov/poxvirus/monkeypox/interim-considerations/acam2000-vaccine.html.

xix Vaughan A, Aarons E, Astbury J, Brooks T, Chand M, Flegg P, Hardman A, Harper N, Jarvis R, MawdsleyS, McGivern M, Morgan D, Morris G, Nixon G, O'Connor C, Palmer R, Phin N, Price DA, Russell K, Said B, Schmid ML, Vivancos R, Walsh A, Welfare W, Wilburn J, Dunning J. Human-to-Human Transmission of Monkeypox Virus, United Kingdom, October 2018. Emerg Infect Dis. 2020 Apr;26(4):782-785. doi: 10.3201/eid2604.191164. Epub 2020 Apr 17. PMID: 32023204; PMCID: PMC7101111.

XX Mendoza R, Petras JK, Jenkins P, et al. Monkeypox Virus Infection Resulting from an Occupational Needlestick — Florida, 2022. MMWR Morb Mortal Wkly Rep 2022;71:1348–1349. DOI: http://dx.doi.org/10.15585/mmwr.mm7142e2

xxi CDC. "Clinical Recognition." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, August 23, 2022. https://www.cdc.gov/poxvirus/monkeypox/clinical-recognition.html.

xxii CDC. "Monitoring and Risk Assessment for Persons Exposed in the Community." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, Updated August 11, 2022.

https://www.cdc.gov/poxvirus/monkeypox/clinicians/monitoring.html.

xxiii CDC. "Laboratory Procedures." Centers for Disease Control and Prevention. Centers for Disease Control and Prevention, December 6, 2022. https://www.cdc.gov/poxvirus/monkeypox/lab-personnel/lab-procedures.html.

xxivBrosius I, Van Dijck C, Coppens J et al. Presymptomatic viral shedding in high-risk mpox contacts: A prospective cohort study. J Med Virol. 2023 May;95(5):e28769. doi: 10.1002/jmv.28769. PMID: 37212312.