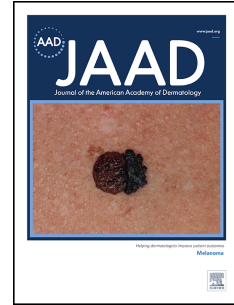


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HIV and mpox: evaluation of clinical course and outcomes from an international dermatologic registry

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PII: S0190-9622(23)03405-9

DOI: <https://doi.org/10.1016/j.jaad.2023.12.032>

Reference: YMJD 18257

To appear in: *Journal of the American Academy of Dermatology*

Received Date: 26 September 2023

Revised Date: 13 December 2023

Accepted Date: 16 December 2023

Please cite this article as: Strahan AG, Galvan Casas C, Prasad S, Fuller LC, Peebles K, Carugno A, Leslie KS, Harp JL, Pumnea T, McMahon DE, Rosenbach M, Lubov JE, Chen G, Pacheco AM, Fox LP, McMillen A, Lim HW, Stratigos AJ, Cronin TA, Kaufmann MD, Hruza GJ, French LE, Freeman EE, HIV and mpox: evaluation of clinical course and outcomes from an international dermatologic registry, *Journal of the American Academy of Dermatology* (2024), doi: <https://doi.org/10.1016/j.jaad.2023.12.032>.

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Article type: Research Letter

Title: HIV and mpox: evaluation of clinical course and outcomes from an international dermatologic registry

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Funding: The AAD/ILDS Dermatology Registry for COVID-19, Monkeypox, and Emerging Infections is supported by a grant from the ILDS and by in-kind support from the AAD.

Conflicts of Interest: Esther Freeman, Klint Peebles, Misha Rosenbach, Terrence Cronin and George Hruza are members of the AAD Ad Hoc Task Force to Create Monkeypox Content. Esther Freeman is the Principal Investigator of the AAD/ILDS Dermatology Registry for COVID-19, Monkeypox, and Emerging Infections, and serves on the WHO Living Monkeypox Atlas and the WHO Monkeypox Guidelines Committee. Cristina Galvan Casas also serves on the WHO Living Monkeypox Atlas committee. Kieron Leslie is a dermatology consultant for the ACTG Study: Tecovirimat For Human Monkeypox Virus (STOMP) and is the Lead Dermatologist on the WHO Living Monkeypox Atlas. Alexander Stratigos is the immediate past President of the EADV. Mark Kaufmann is the President of the AAD. Terrence Cronin is President-elect of the AAD. Lars French is the Past President of the ILDS. Henry W. Lim is the President

of ILDS, and George Hruza and Claire Fuller are Board members of the ILDS. Lindy Fox is a Board member of the AAD. Cristina Galvan Casas serves on the WHO Living Monkeypox Atlas.

IRB approval status: Reviewed by Massachusetts General Hospital Institutional Review Board, deemed not human subjects research. Patient consent not applicable.

Reprint requests: Esther Freeman

Manuscript word count: 500/500

References: 5/5

Figures: 0

Tables: 1

Keywords: general dermatology; medical dermatology; monkeypox; mpox; infectious disease; virus; viral infection; global health; international health; vaccine; immunocompromised; skin lesions

1 In July 2022, the World Health Organization declared mpox a public health emergency. Since
2 outbreak onset, >91,000 cases have been reported.¹ During the 2022 mpox outbreak, most disease
3 courses were mild.² In some cases, however, severe outcomes were reported and included
4 hospitalization and death. People living with HIV/AIDS (PLWH) have been disproportionately affected and
5 all immunocompromised individuals are at higher risk of severe outcomes.^{3,4} Hospitalization in those
6 diagnosed with mpox ranges from 14-59%.⁴ In cases requiring hospitalization, death was reported in 4%
7 of cases but is as high as 25% in PLWH.^{3,4} Reports of dermatologic findings and associated clinical
8 outcomes in PLWH are limited.

9 In August 2022, the AAD (American Academy of Dermatology)/ILDS (International League of
10 Dermatological Societies) Dermatology COVID-19, Mpox, and Emerging Infections Registry began
11 collecting mpox dermatologic manifestations. Cases entered from August 2022 to March 2023 were
12 analyzed in Stata version 16 (StataCorp, LLC). The Chi-square test was used to examine association. This
13 study was deemed not Human Subjects Research by the Massachusetts General Brigham (MGB)
14 Institutional Review Board. Building on our prior work with this registry data, we examine the frequency
15 of severe outcomes, notably hospitalization and death, and evaluate differences in clinical course
16 (clinical signs/symptoms, lesion counts, co-infections, treatment) and outcomes in PLWH compared to
17 those without HIV.⁵

18 Of all 120 registry-reported cases, 23 (19%) required hospitalization. The most common reason
19 for hospitalization was for a “skin-related condition” (8/23, 35%). Other reasons included: isolation in
20 accordance with national guidelines (5/23, 22%), sore throat/ oral lesions (4/23, 17%), and rectal pain
21 (3/23, 13%). Of hospitalized patients, 30% were PLWH (7/23). Nine hospitalized patients received
22 tecovirimat. Co-infection with other sexually transmissible infections (STI) was more common in PLWH
23 than in those without, 57% vs. 38% respectively ($p<0.01$, Table 1, primarily driven chlamydia and
24 syphilis). No differences were noted in total lesion count, use of medical counter measures, time to

25 resolution, or scarring between hospitalized PLWH and those without HIV. One death was reported in a
26 person living with HIV not on anti-retroviral therapy.

27 In 120 cases reported to an international registry, clinical course and outcomes were similar in
28 PLWH and those without HIV, with the exception of frequency of co-infection with other STIs. We did
29 not assess CD4+ count, an important variable noted to influence mpox clinical course and outcomes.³
30 Concurrent testing for both HIV and other STIs remains important in suspected mpox cases. Notably,
31 skin-related conditions were the most frequently reported reason for hospitalization, a finding not
32 previously reported. One notable limitation is the self-identified race of reported cases; 58% were in
33 those who identified as White, yet it is worth noting that mpox disproportionately affects ethnic and racial
34 minorities, groups which also encounter the highest barriers to mpox and HIV care. Registry-reported
35 data and potential for preferential reporting of mpox cases with cutaneous manifestations are additional
36 limitations. These findings underscore the need for prompt recognition and treatment to reduce
37 hospitalization and improve patient outcomes, as well as the importance of dermatologist involvement
38 in the inpatient setting.

Table 1. Demographic and Clinical Features of Mpox PLWH vs. non-PLWH (n,%)

| | PLWH (n=44) | Non-PLWH (n=76) |
|---|-------------|-----------------|
| Age | | |
| 18-30 years | 5 (11) | 26 (34) |
| 31-49 years | 33 (75) | 41 (54) |
| Above 50 years | 6 (14) | 9 (12) |
| Sex | | |
| Female | 1 (2) | 4 (5) |
| Male | 43 (98) | 72 (95) |
| Race/Ethnicity* | | |
| White | 21 (48) | 49 (64) |
| Hispanic/Latino | 18 (41) | 13 (17) |
| Black/African American | 2 (5) | 3 (4) |
| Asian | 0 | 3 (4) |
| Other/Unknown | 1 (2) | 4 (5) |
| Sexual Behavior† | | |
| Same-sex behavior | 40 (91) | 63 (83) |
| Group sex behavior | 14 (32) | 21 (28) |
| Initial Clinical Sign/Symptom | | |
| Skin Lesion | 20 (45) | 44 (58) |
| Fever | 7 (16) | 13 (17) |
| Lymphadenopathy | 2 (5) | 1 (1) |
| Intra-oral/throat lesions | 0 | 1 (1) |
| General Malaise | 7 (16) | 6 (8) |
| Sore Throat | 5 (11) | 5 (7) |
| Rectal Pain | 3 (7) | 4 (5) |
| Ocular/Ophthalmic Involvement | 0 | 1 (1) |
| Edema | 0 | 1 (1) |
| Total Number of Lesions | | |
| 1 | 2 (5) | 8 (11) |
| 2-5 | 11 (25) | 18 (24) |
| 6-20 | 22 (50) | 32 (42) |
| 21-50 | 8 (18) | 10 (13) |
| 51-100 | 1 (2) | 3 (4) |
| 200+ | 0 | 1 (1) |
| Unknown | 0 | 4 (5) |
| Co-Infections | | |
| Gonorrhea | 11 (25) | 7 (9) |
| Chlamydia | 4 (9) | 3 (4) |
| Syphilis | 10 (23) | 6 (8) |
| Varicella Zoster | 0 | 1 (1) |
| Herpes Simplex | 5 (11) | 3 (4) |
| S. aureus | 1 (2) | 1 (1) |
| Hospitalized | | |
| Yes | 7 (16) | 16 (21) |
| No | 37 (84) | 60 (79) |
| Death | 1 (2) | 0 |
| Treatment‡ | | |
| Tecovirimat | 16 (36) | 10 (23) |
| Cidofovir/Brincidofovir | 2 (5) | 2 (5) |
| Oral/IV antibiotics | 3 (7) | 7 (16) |
| Oral/IV pain medication | 4 (9) | 3 (7) |
| Topical antiseptics, antibiotics, or analgesics | 5 (11) | 8 (18) |
| Vaccinia immune globulin | 0 | 0 |
| Other | 1 (2) | 2 (5) |

*Optional response

† Group sex behavior is defined as engagement in a sexual encounter that involves more than one partner

‡Other treatments included: PLWH- bictegrovir, emtricitabine & tenofovir alafenamide (Biktarvy), valacyclovir; non-PLWH- acyclovir

- 39 1. CDC. 2022 Monkeypox Outbreak Global Map. Updated Nov 4 2022. Accessed Nov 6, 2022.
40 <https://www.cdc.gov/poxvirus/monkeypox/response/2022/world-map.html>
- 41 2. Yon H, Shin H, Shin JI, et al. Clinical manifestations of human Mpox infection: A systematic
42 review and meta-analysis. *Reviews in Medical Virology*. 2023;33(4):e2446.
43 doi:<https://doi.org/10.1002/rmv.2446>
- 44 3. Mitjà O, Alemany A, Marks M, et al. Mpox in people with advanced HIV infection: a global case
45 series. *Lancet*. Mar 18 2023;401(10380):939-949. doi:10.1016/s0140-6736(23)00273-85.
- 46 4. Benites-Zapata VA, Ulloque-Badaracco JR, Alarcon-Braga EA, et al. Clinical features,
47 hospitalisation and deaths associated with monkeypox: a systematic review and meta-analysis. *Ann Clin*
48 *Microbiol Antimicrob*. Aug 10 2022;21(1):36. doi:10.1186/s12941-022-00527-1
- 49 5. Prasad S, Casas CG, Strahan AG, et al. A dermatologic assessment of 101 mpox (monkeypox)
50 cases from 13 countries during the 2022 outbreak: skin lesion morphology, clinical course, and scarring.
51 *Journal of the American Academy of Dermatology*. doi:10.1016/j.jaad.2022.12.035