



HAL
open science

Intrafamily Transmission of Monkeypox Virus, Central African Republic, 2018

Camille Besombes, Ella Gonofio, Xavier Konamna, Benjamin Selekon, Antoine Gessain, Nicolas Berthet, Jean-Claude Manuguerra, Arnaud Fontanet, Emmanuel Nakouné

► **To cite this version:**

Camille Besombes, Ella Gonofio, Xavier Konamna, Benjamin Selekon, Antoine Gessain, et al.. Intrafamily Transmission of Monkeypox Virus, Central African Republic, 2018. *Emerging Infectious Diseases*, 2019, 25 (8), pp.1602-1604. 10.3201/eid2508.190112 . pasteur-02917509

HAL Id: pasteur-02917509

<https://pasteur.hal.science/pasteur-02917509v1>

Submitted on 19 Aug 2020

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.



Distributed under a Creative Commons CC0 - Public Domain Dedication 4.0 International License

About the Author

Dr. Wieland is a professor of virology at the Institute of Virology of the University of Cologne, Germany. Her research interests include diagnosis and epidemiology of viral diseases.

References

1. Phan TG, Dreno B, da Costa AC, Li L, Orlandi P, Deng X, et al. A new protoparvovirus in human fecal samples and cutaneous T cell lymphomas (mycosis fungoides). *Virology*. 2016;496:299–305. <http://dx.doi.org/10.1016/j.virol.2016.06.013>
2. Väisänen E, Fu Y, Hedman K, Söderlund-Venermo M. Human protoparvoviruses. *Viruses*. 2017;9:E354. <http://dx.doi.org/10.3390/v9110354>
3. Mollerup S, Fridholm H, Vinner L, Kjartansdóttir KR, Friis-Nielsen J, Asplund M, et al. Cutavirus in cutaneous malignant melanoma. *Emerg Infect Dis*. 2017;23:363–5. <http://dx.doi.org/10.3201/eid2302.161564>
4. Wieland U, Silling S, Scola N, Potthoff A, Gambichler T, Brockmeyer NH, et al. Merkel cell polyomavirus infection in HIV-positive men. *Arch Dermatol*. 2011;147:401–6. <http://dx.doi.org/10.1001/archdermatol.2011.42>
5. Väisänen E, Fu Y, Koskenmies S, Fyhrquist N, Wang Y, Keinonen A, et al. Cutavirus DNA in malignant and non-malignant skin of cutaneous T-cell lymphoma and organ transplant patients but not of healthy adults. *Clin Infect Dis*. 2019;68:1904–10. <http://dx.doi.org/10.1093/cid/ciy806>
6. Väisänen E, Mohanraj U, Kinnunen PM, Jokelainen P, Al-Hello H, Barakat AM, et al. Global distribution of human protoparvoviruses. *Emerg Infect Dis*. 2018;24:1292–9. <http://dx.doi.org/10.3201/eid2407.172128>
7. Kreuter A, Nasserani N, Tigges C, Oellig F, Silling S, Akgül B, et al. Cutavirus infection in primary cutaneous B- and T-cell lymphoma. *JAMA Dermatol*. 2018;154:965–7. <http://dx.doi.org/10.1001/jamadermatol.2018.1628>
8. Bergallo M, Daprà V, Fava P, Ponti R, Calvi C, Fierro MT, et al. Lack of detection of cutavirus DNA using PCR real time in cutaneous T-cell lymphomas (CTCL). *G Ital Dermatol Venereol*. 2018;•••. <http://dx.doi.org/10.23736/S0392-0488.18.06161-8>
9. Dickinson A, Xu M, Silén S, Wang Y, Fu Y, Sadeghi M, et al. Newly detected DNA viruses in juvenile nasopharyngeal angiofibroma (JNA) and oral and oropharyngeal squamous cell carcinoma (OSCC/OPSCC). *Eur Arch Otorhinolaryngol*. 2019;276:613–7. <http://dx.doi.org/10.1007/s00405-018-5250-7>
10. Hufbauer M, Akgül B. Molecular mechanisms of human papillomavirus induced skin carcinogenesis. *Viruses*. 2017;9:E187. <http://dx.doi.org/10.3390/v9070187>

Address for correspondence: Ulrike Wieland, University of Cologne Institute of Virology, Fuerst-Pueckler-Str. 56, 50935 Cologne, Germany; email: ulrike.wieland@uni-koeln.de

Intrafamily Transmission of Monkeypox Virus, Central African Republic, 2018

Camille Besombes, Ella Gonfio, Xavier Konamna, Benjamin Selekon, Antoine Gessain, Nicolas Berthet, Jean-Claude Manuguerra, Arnaud Fontanet, Emmanuel Nakouné

Author affiliations: Institut Pasteur, Paris, France (C. Besombes, A. Gessain, N. Berthet, J.-C. Manuguerra, A. Fontanet); Institut Pasteur de Bangui, Bangui, Central African Republic (E. Gonfio, X. Konamna, B. Selekon, E. Nakouné); Centre National de la Recherche Scientifique, Paris (A. Gessain, N. Berthet); Unité Pasteur-CNAM Risques Infectieux et Emergents, Conservatoire National des Arts et Métiers, Paris (A. Fontanet)

DOI: <https://doi.org/10.3201/eid2508.190112>

Monkeypox is a rare viral zoonotic disease; primary infections are reported from remote forest areas of Central and West Africa. We report an investigation of a monkeypox outbreak in Lobaye, southwest Central African Republic, in October 2018.

Monkeypox, a zoonotic disease caused by an *Orthopoxvirus*, has clinical signs and symptoms in humans similar to smallpox and a case-fatality rate of 10% (1). The specific reservoir species for monkeypox virus remains, to a large extent, unidentified (2). Spillover events of monkeypox have been reported in remote forest areas of Central and West Africa. After zoonotic infection, the virus can be transmitted from person to person (1).

To date, human monkeypox outbreaks in the Central African Republic (CAR) have been small: ≈10 cases, restricted to a family or village. Primary infection in these outbreaks occurred from contact with wild fauna, with secondary transmission among close contacts in the community (3,4) and limited nosocomial transmission (5). Since 2000, the Virology Laboratory of the Institut Pasteur de Bangui (IP Bangui), a regional reference center for monkeypox, has reported 20 monkeypox outbreaks across several regions of CAR, totaling ≈100 cases, particularly in the region of Lobaye (3,4). In 2018 alone, IP Bangui investigated 6 different outbreaks in CAR, indicating a possible increase in frequency of outbreaks (6,7).

On September 27, 2018, a healthcare worker from Zomea Kaka healthcare center in Lobaye reported to IP Bangui about 3 cases of suspected monkeypox in an Aka Pygmy family. A 25-year-old female sought care at the health center, 10 km from her village, for maculopapular rash and lesions. She was afebrile. Her signs and symptoms indicated

resolving late stage monkeypox infection. She was accompanied by her 2 daughters, 5 months and 4 years of age, both showing typical symptoms of active monkeypox infection, notably maculopapular rash on the palms of their hands and soles of their feet (Appendix Figure, <http://wwwnc.cdc.gov/EID/article/25/8/19-0112-App1.pdf>). Blood or pus samples taken from the 3 patients were confirmed positive for monkeypox infection by PCR on September 29 (8) (Appendix).

On October 5, IP Bangui carried out an investigation among contacts of the index case-patient, in collaboration with the Ministry of Health and the World Health Organization CAR Country Office. The index case-patient reported butchering 3 small mammals known in local Aka language as Yabo (African civet, *Civettictis civetta*), Gbè (Emin's pouched rat, *Cricetomys emini*), and Sende (African rope squirrel, *Funesciurus anerythrus*). She butchered 1 of each in a forested area 2 weeks before the onset of rash.

During October 6–10, two additional family contacts from the village, the index case-patient's 2 sisters, 7 and 16 years of age, reported symptoms consistent with monkeypox infection. Healthcare workers collected blood or pus samples from the patients, and IP Bangui confirmed monkeypox infection by PCR. On October 26, monkeypox infection was confirmed in another family contact, the index case-patient's

33-year-old sister-in-law. The dates of the onset of symptoms suggest 3 waves of intrafamilial transmission (Table) (9).

IP Bangui conducted further investigations by using *Orthopoxvirus* serologic assays (Appendix) on blood samples collected from 2 healthcare worker contacts on October 5 and from 31 village contacts on October 25. Results revealed evidence of *Orthopoxvirus* serologic response in the index case-patient's mother; 2 healthcare workers who had cared for the index case-patient; and the index case-patient's brother, who brought her the wild animals (Table).

Serologic evidence of possible monkeypox infection can indicate prior exposure to the virus or, among persons >38 years of age, immunization against smallpox, and might explain the restricted size of the outbreak in the village. However, smallpox vaccination campaigns with a live-attenuated vaccinia virus ended in 1979 in CAR. Consequently, an increasingly larger proportion of the population is immunologically naive to *Orthopoxvirus* infection.

This investigation identified 5 clinical cases of secondary monkeypox infection spread over 3 waves of intrafamilial infection, originating from an index case-patient with primary infection possibly attributable to contact with wild fauna. The prompt declaration and isolation of suspected cases, as well as possible naturally

Table. Molecular and serologic evidence of index case-patient and contacts with known and possible exposure to monkeypox virus, Central African Republic, 2018*

Patients	Age, y/sex	Symptom onset date	Signs/symptoms	Animal contact	Collection date	Sample type		PCR†		IgG§		Smallpox vaccine¶
						Blood	Pus†	MPXV	CPXV	MPXV	CPXV	
Index case-patient	25/F	2018 Sep 8	Rash, lesions	Y	2018 Sep 27	Y	N	+	–	–	–	N
Contacts												
Daughter	0.4/F	2018 Sep 20	Fever, rash, lesions	N	2018 Sep 27	N	Y	+	–	ND	ND	N
Daughter	4/F	2018 Sep 26	Fever, rash, lesions	N	2018 Sep 27	Y	N	+	–	–	–	N
Sister	16/F	2018 Oct 6	Rash, lesions	N	2018 Oct 8	N	Y	+	–	ND	ND	N
Sister	7/F	2018 Oct 9	Rash, lesions	N	2018 Oct 11	Y	N	+	–	–	–	N
SIL	33/F	2018 Oct 24	Rash, lesions	N	2018 Oct 25	Y	N	+	–	–	–	N
Mother	49/F	NA	None	N	2018 Oct 5	Y	N	ND	ND	+	+	Y
Son	13/M	NA	None	Y	2018 Oct 5	Y	N	ND	ND	–	–	N
Brother	49/M	NA	None	Y	2018 Oct 25	Y	N	ND	ND	+	–	Y
Brother of SIL	8/M	NA	None	NK	2018 Oct 25	Y	N	ND	ND	+	+	N
Nephew of SIL	13/M	NA	None	NK	2018 Oct 25	Y	N	ND	ND	–	–	N
HCW	34/M	NA	None	N	2018 Oct 5	Y	N	ND	ND	+	+	N
HCW	45/F	NA	None	N	2018 Oct 5	Y	N	ND	ND	+	+	Y
Social contact	22/F	NA	None	NK	2018 Oct 25	Y	N	ND	ND	+	–	N

*A total of 33 contacts were tested, 2 HCWs and 31 village contacts. CPXV, cowpox virus; HCW, healthcare worker; MPXV, monkeypox virus; NA, not applicable; ND, not done; NK, not known; SIL, sister-in-law; +, positive; –, negative.

†Samples obtained by HCWs after training on collecting swab samples.

‡Quantitative and conventional PCR were performed by using generic primers G2R-G and Congo Basin primers C3L (8).

§In-house tests were performed by using MPXV antigen isolated from local human cases and CPXV antigen related to Brighton Red strain.

¶History of smallpox vaccination was determined by verbal report and presence of scar.

acquired immunity or persistence of vaccine-derived immunity within the community, likely contributed to the restricted extent of secondary transmission. Further studies are needed to clarify risk factors for primary and secondary monkeypox transmission.

Positive serologic findings in healthcare workers during this investigation also highlight the limited infection prevention and control resources, such as isolation rooms, gowns, gloves, N95 respirators, and goggles, to protect healthcare workers responding to outbreaks in CAR. For communities located in remote forest areas in which zoonotic spillover and secondary transmission are thought to occur regularly, health center capacity and resources need to be strengthened. Health centers urgently need training on case recognition for healthcare workers, access to diagnostic capacities, and appropriate infection prevention and control measures to reduce the possibility of secondary transmission in these areas (10).

Acknowledgments

The authors thank Romain Duda for his assistance with identification of the animal species in Aka language. We also acknowledge Rebecca Grant for her relevant suggestions and her kind participation in the formatting of this research letter.

About the Author

Dr. Besombes is an infectious and tropical disease clinician who works as a researcher in the Emerging Diseases Epidemiology Unit at Institut Pasteur, Paris, France. Her primary research interests include tropical diseases, specifically zoonotic and vectorborne diseases, and hepatitis Delta virus infection.

References

- Durski KN, McCollum AM, Nakazawa Y, Petersen BW, Reynolds MG, Briand S, et al. Emergence of monkeypox—West and Central Africa, 1970–2017. *MMWR Morb Mortal Wkly Rep*. 2018;67:306–10. <http://dx.doi.org/10.15585/mmwr.mm6710a5>
- Sklenovská N, Van Ranst M. Emergence of monkeypox as the most important orthopoxvirus infection in humans. *Front Public Health*. 2018;6:241. <http://dx.doi.org/10.3389/fpubh.2018.00241>
- Berthet N, Nakouné E, Whist E, Selekon B, Burguière AM, Manuguerra JC, et al. Maculopapular lesions in the Central African Republic. *Lancet*. 2011;378:1354. [http://dx.doi.org/10.1016/S0140-6736\(11\)61142-2](http://dx.doi.org/10.1016/S0140-6736(11)61142-2)
- Kalthan E, Tenguere J, Ndjapou SG, Koyazengbe TA, Mbomba J, Marada RM, et al. Investigation of an outbreak of monkeypox in an area occupied by armed groups, Central African Republic. *Med Mal Infect*. 2018;48:263–8. <http://dx.doi.org/10.1016/j.medmal.2018.02.010>
- Nakoune E, Lampaert E, Ndjapou SG, Janssens C, Zuniga I, Van Herp M, et al. A nosocomial outbreak of human monkeypox in the Central African Republic. *Open Forum Infect Dis*. 2017;4:ofx168. <http://dx.doi.org/10.1093/ofid/ofx168>
- ProMED-mail. Monkeypox—Africa (12): Central African Republic. 2018 Jul 30 [cited 2018 Oct 24]. <https://www.promedmail.org/archive/no.20180730.5936829>
- ProMED-mail. Monkeypox—Africa (05): Central African Republic (Haute-Kotto). 2018 Apr 4 [cited 2019 Jan 7]. <https://www.promedmail.org/archive/no.20180403.5726159>
- Li Y, Zhao H, Wilkins K, Hughes C, Damon IK. Real-time PCR assays for the specific detection of monkeypox virus West African and Congo Basin strain DNA. *J Virol Methods*. 2010;169:223–7. <http://dx.doi.org/10.1016/j.jviromet.2010.07.012>
- Di Giulio DB, Eckburg PB. Human monkeypox: an emerging zoonosis. *Lancet Infect Dis*. 2004;4:15–25. [http://dx.doi.org/10.1016/S1473-3099\(03\)00856-9](http://dx.doi.org/10.1016/S1473-3099(03)00856-9)
- Munster VJ, Bausch DG, de Wit E, Fischer R, Kobinger G, Muñoz-Fontela C, et al. Outbreaks in a rapidly changing Central Africa—lessons from Ebola. *N Engl J Med*. 2018;379:1198–201. <http://dx.doi.org/10.1056/NEJMp1807691>

Address for correspondence: Emmanuel Nakouné, Arboviruses, Viral Haemorrhagic Viruses, Emerging Viruses and Zoonosis Unit, Institut Pasteur de Bangui, Bangui, Central African Republic; email: emmanuel.nakoune@pasteur-bangui.org

Intact *Mycobacterium leprae* Isolated from Placenta of a Pregnant Woman, China

Zhiming Chen,¹ Yanfei Kuang,¹ Haiqin Jiang, Wenyue Zhang, Ying Shi, Santosh Chokkaku, Huan Chen, Junhua Li, Hongsheng Wang

Author affiliations: Chinese Academy of Medical Sciences Institute of Dermatology, Nanjing, China (Z. Chen, H. Jiang, W. Zhang, Y. Shi, S. Chokkaku, H. Wang); Hunan Provincial Center for Disease Control and Prevention, Changsha, China (Y. Kuang, H. Chen, J. Li); Jiangsu Key Laboratory of Molecular Biology for Skin Diseases and STIs, Nanjing (H. Wang); Nanjing Medical University Center for Global Health, Nanjing (H. Wang)

DOI: <https://doi.org/10.3201/eid2508.190114>

Whether *Mycobacterium leprae* transmits from placenta to fetus remains unknown. We describe the case of a pregnant woman with untreated histoid leproma. Although her newborn was healthy, laboratory examination revealed intact *M. leprae* present in the placenta, suggesting that the placental barrier might prevent vertical dissemination of *M. leprae*.

¹These authors contributed equally to and are co-first authors for this article.