**Feline Panleukopenia Virus in Dogs from Italy and Egypt**

Georgia Diakoudi, Costantina Desario, Gianvito Lanave, Stefania Salucci, Linda A. Ndiana, Aya Attia Koraney Zarea, Ehab Ali Fouad, Alessio Lorusso, Flora Alfano, Alessandra Cavalli, Canio Buonavoglia, Vito Martella, Nicola Decaro

Author affiliations: University of Bari Aldo Moro, Bari, Italy (G. Diakoudi, C. Desario, G. Lanave, L.A. Ndiana, A.A. Koraney Zarea, A. Cavalli, C. Buonavoglia, V. Martella, N. Decaro); Istituto Zooprofilattico Sperimentale dell’Abruzzo e Molise G. Caporale, Teramo, Italy (S. Salucci, A. Lorusso); National Research Centre, Giza, Egypt (E.A. Fouad); Istituto Zooprofilattico Sperimentale del Mezzogiorno, Portici, Italy (F. Alfano)

DOI: https://doi.org/10.3201/eid2809.220388

Canine parvovirus (CPV or CPV-2) and feline panleukopenia virus (FPV) are variants of *Carnivore protoparvovirus 1*. We identified and characterized FPV in dogs from Italy and Egypt using genomic sequencing and phylogenetic analyses. Cost-effective sequencing strategies should be used to monitor interspecies spread, evolution dynamics, and potential host jumping of FPV.

**Canine parvovirus** (CPV or CPV-2) and *feline panleukopenia virus* (FPV) are variants of *Carnivore protoparvovirus 1* and major pathogens of domestic and wild carnivores. The linear, single-stranded DNA genome contains 2 open reading frames that encode 2 nonstructural and 2 capsid proteins (1).

FPV and CPV are closely related antigenically and genetically (≥98% identity at the nucleotide level) but differ in host range and pathogenicity. These biological differences are determined by amino acid mutations in the VP2 capsid protein (2). CPV-2 antigenic variants 2a, 2b, and 2c are able to infect felids and cause FPV-like disease (2). FPV is believed to be incapable of infecting dogs but has been shown to replicate in some canine tissues after experimental oronasal infection (3). Furthermore, studies have reported the presence of FPV in dogs with CPV-like gastroenteritis (4–8).

The Infectious Diseases Unit, Department of Veterinary Medicine, University of Bari, Italy, has performed routine screening and characterization of canine samples for CPV and FPV since the mid-1990s and has combined traditional virological and molecular techniques to differentiate between FPV and CPV types 2a/2b and 2b/2c (9). During 2019–2021, we screened and typed ≤1,000 *Carnivore protoparvovirus 1* strains from cats and dogs. In 2021, FPV was unexpectedly identified in dogs from epidemiologically unrelated cases. We identified FPV in a blood sample obtained from a 1-year-old dog from Giza, Egypt, that had an unexplained fever (case A) and in fecal samples from 3 dead adult dogs in Teramo, Italy (case series B). The dogs from Italy had severe gastrointestinal symptoms that the attending veterinarian initially suspected were from poisoning.

Because finding FPV in dogs is unusual, we analyzed the samples by using 2 PCR primer sets that differentiated between canine and feline cytochrome b genes (10). We confirmed that the FPV samples were of canine origin. Moreover, toxicologic analysis of the 3 dogs in case series B excluded a diagnosis of poisoning, and other enteric canine pathogens were excluded as causes of the gastrointestinal symptoms by using culture and molecular assays. We performed immunofluorescence analyses of tissue from the small intestines of the 3 dogs in case series B and detected parvoviral antigens in epithelial cells (Appendix Figure 1, https://wwwnc.cdc.gov/EID/article/28/9/22-0388-App1.pdf). Attempts to isolate the virus by infecting feline or canine cell lines were unsuccessful, likely because of low viral titers.

To acquire complete viral genome sequences of samples from each case, we performed genomic PCR by using LA Taq polymerase (Takara Bio, http://www.takarabio.com). PCR products were used for library preparation. We performed adapter ligation of genomic DNA by using the Ligation Sequencing Kit (Oxford Nanopore Technologies, https://nanoporetech.com) according to the manufacturer’s guidelines. Sequencing was performed by using the FLO-MIN106D Flow Cell, R9 version, and MinION Mk1C sequencing platform (Oxford Nanopore Technologies).

We obtained complete sequences of coding regions for the virus strains from Italy (ITA/2021/164-1; GenBank accession no. OM638042) and Egypt (EGY/2021/139-188; GenBank accession no. OM638043). The ITA/2021/164-1 and EGY/2021/139-188 strains were characterized as FPV on the basis of sequence and phylogenetic analyses. We aligned amino acid sequences for VP2 from FPV, CPV, and the dog-associated FPV strains from Italy and Egypt to determine biological differences between the variants (Figure). We identified an I101T aa substitution in substitution in our cases that is likely associated with host range determination (Figure) (2). The FPV strains from Italy and Egypt segregated into different phylogenetic clusters (Appendix Figure 2).
FPV has been recently reported in Pakistan, Vietnam, Thailand, and China in dogs that had gastrointestinalitis (4–8). These viruses were characterized as FPV after partial or complete sequence analysis of the gene encoding VP2. A unique K93N substitution mutation involved in host range control (2) was identified in an FPV strain in Thailand (4), and I101T mutations were found in dog-associated FPV strains from Vietnam (5) and China (6). As noted, the I101T mutation was also found in our dog-associated FPV strains. I101 has been observed in prototypical FPV strains, whereas T101 has been found in recent FPV isolates (5). The I101T substitution has also been observed in CPV-2 and its variant CPV-2a and is believed to be a further adaptation of CPV to the canine host (2).

In conclusion, we identified FPV from unrelated cases in dogs. In case A, fever was the only clinical sign in a young dog, whereas a fatal systemic syndrome with enteric signs occurred in 3 adult dogs in case series B. However, the role of FPV in these cases remains unclear. Adoption of cost-effective sequencing strategies in recent years has demonstrated that residual circulation of FPV or FPV-like viruses occurs in dogs in some settings. Genomic sequencing and further phylogenetic analyses can be used to monitor the spread, evolution, and potential host jumping of Carnivore protoparvovirus 1 variants in domestic and wild carnivores.

This study was supported by grants from the Ministry of Health, Italy: Ricerca Corrente 2019 ‘NGS e diagnostica molecolare in sanità animale’ (A.L., N.D.) and Ricerca Corrente 2018 ‘Nuovi virus gastroenterici di cane e gatto: sviluppo di protocolli NGS per la valutazione del rischio zoonosico’ (F.A., N.D.).

The study was approved by the Ethics Committee of the Department of Veterinary Medicine, University of Bari Aldo Moro, Italy (authorization no. 28/2020) and Medical Research Ethics Committee at the National Research Centre, Egypt (authorization no. 6211022021). All experiments were performed in accordance with relevant guidelines and regulations.

About the Author

Dr. Diakoudi is a research scientist at the University of Bari Aldo Moro, Italy. Her research interests focus on virus discovery in animals, particularly viruses with zoonotic potential.

References

1. Cotmore SF, Agbandje-McKenna M, Canuti M, Chiorini JA, Eis-Hubinger AM, Hughes J, et al. Ictv Report Consortium. ICTV virus taxonomy profile: Parvoviridae. J Gen Virol. 2019;100:367–8. https://doi.org/10.1099/jgv.0.001212

2. Callaway HM, Welsch K, Weichert W, Allison AB, Hafenstein SL, Huang K, et al. Complex and dynamic interactions between parvovirus capsids, transferrin receptors, and antibodies control cell infection and host range. J Virol. 2018;92:e00460-18 PubMed https://doi.org/10.1128/JVI.00460-18

3. Truyen U, Parrish CR. Canine and feline host ranges of canine parvovirus and feline panleukopenia virus: distinct host cell tropisms of each virus in vitro and in vivo. J Virol. 1992;66:5399–408. https://doi.org/10.1128/jvi.66.9.5399-5408.1992

4. Inthong N, Kaewnongkol S, Meekhanon N, Sirinarumit K, Sirinarumit T. Dynamic evolution of canine parvovirus in Thailand. Vet World. 2020;13:245–55. https://doi.org/10.14202/vetworld.2020.245-255

5. Hoang M, Wu CN, Lin CF, Nguyen HTT, Le VP, Chiuo MT, et al. Genetic characterization of feline panleukopenia virus from dogs in Vietnam reveals a unique Thr101 mutation in VP2. PeerJ. 2020;8:e9752. https://doi.org/10.7717/peerj.9752
6. Chen B, Zhang X, Zhu J, Liao L, Bao E. Molecular epidemiological survey of canine parvovirus circulating in China from 2014 to 2019. Pathogens. 2021;10:588. https://doi.org/10.3390/pathogens10050588
7. Ahmed N, Riaz A, Zubair Z, Saqib M, Ijaz S, Nawaz-Ul-Rehman MS, et al. Molecular analysis of partial VP-2 gene amplified from rectal swab samples of diarrheic dogs in Pakistan confirms the circulation of canine parvovirus genetic variant CPV-2a and detects sequences of feline panleukopenia virus (FPV). Virol J. 2018;15:45. https://doi.org/10.1186/s12985-018-0958-y
8. Charoenkul K, Tangwangvivat R, Janetanakit T, Boonyapisitsopa S, Bunpapong N, Chaityawong S, et al. Emergence of canine parvovirus type 2c in domestic dogs and cats from Thailand. Transbound Emerg Dis. 2019;66:1518–28. https://doi.org/10.1111/tbed.13177
9. Ndiana LA, Lanave G, Desario C, Berjaoui S, Alfano F, Puglia I, et al. Circulation of diverse protoparvoviruses in wild carnivores, Italy. Transbound Emerg Dis. 2021;68:2489–502. https://doi.org/10.1111/tbed.13917
10. Abdel-Rahman SM, El-Saadani MA, Ashry KM, Haggag AG. Detection of adulteration and identification of cat’s, dog’s, donkey’s and horse’s meat using species-specific PCR and PCR-RFLP techniques. Aust J Basic & Appl Sci. 2009;3:1716–9. http://www.ajbasweb.com/old/ajbas2009/1716-1719

Address for correspondence: Nicola Decaro, Department of Veterinary Medicine, University of Bari Aldo Moro, Strada Prov. per Casamassima Km 3, 70010 Valenzano, Bari, Italy; email: nicola.decaro@uniba.it

Trichodysplasia Spinulosa Polyomavirus Endothelial Infection, California, USA

Lauren Lawrence, Aihui Wang, Gregory Charville, Angus Toland, Benjamin Pinsky, Yasodha Natkunam, Sheren Younes, Henning Stehr, Dita Gratzinger

Author affiliations: Stanford Healthcare, Palo Alto, California, USA (L. Lawrence, A. Toland); Stanford University School of Medicine, Stanford, California, USA (A. Wang, G. Charville, B. Pinsky, Y. Natkunam, S. Younes, H. Stehr, D. Gratzinger)

DOI: https://doi.org/10.3201/eid2809.220856

We describe 3 patients in California, USA, with trichodysplasia spinulosa polyomavirus (TSPyV) infection of endothelium after steroid administration. We detected TSPyV RNA in tissue specimens by in situ hybridization, which revealed localization to endothelial cells. These cases suggest that diseases associated with endothelial inflammation could be associated with TSPyV infection.

Trichodysplasia spinulosa polyomavirus (TSPyV) is an alphapolyomavirus whose primary clinical manifestation in a posttransplant setting is folliculocentric papular cutaneous eruptions, typically involving the face (1). Identification of TSPyV nucleic acids in tonsillar tissue has led to the speculation that lymphoid tissue might be a latency site (2); however, some disagreement exists in the literature as to whether the clinical diagnosis of trichodysplasia spinulosa reflects primary infection or reactivation of latent virus (3,4). Although cutaneous disease is the primary clinical manifestation of infection, TSPyV has been identified in blood, urine, cerebrospinal fluid, tonsils, and respiratory specimens by various methods, including nucleic acid detection, immunohistochemistry, and electron microscopy (2,4–6). TSPyV DNA loads can be high, especially in blood (up to 10^9 viral copies/mL), months before the appearance of typical trichodysplasia spinulosa skin lesions (4).

This case study was part of a larger project approved by the Stanford Institutional Review Board (approval no. 58311) designed to explore oncogenesis by alphapolyomaviruses. We identified rare cases on our next generation sequencing panel of solid tumors with off-target, high quality reads that aligned to the TSPyV genome. We hypothesized that some of these cases might represent TSPyV-mediated neoplasms. Our cases comprised 1 patient with metastatic lung adenocarcinoma involving the brain (case 1), 1 patient with meningioma (case 2), and 1 patient with a metastatic perivascular epithelial cell tumor involving the liver (Table). It is unclear whether the mental status changes observed in cases 1 and 2 were attributable to viral infection or were secondary to the tumors (Table). All 3 patients received steroids immediately preceding resection. We performed in situ hybridization using a custom RNAscope probe that targeted the complete TSPyV viral genome (GenBank accession no. NC_014361.1) and RNAscope 2.5 HD Reagent Kit-RED (Advanced Cell Diagnostics, https://acdbio.com) to detect TSPyV RNA in cutaneous biopsy specimens. One case of cutaneous trichodysplasia spinulosa was