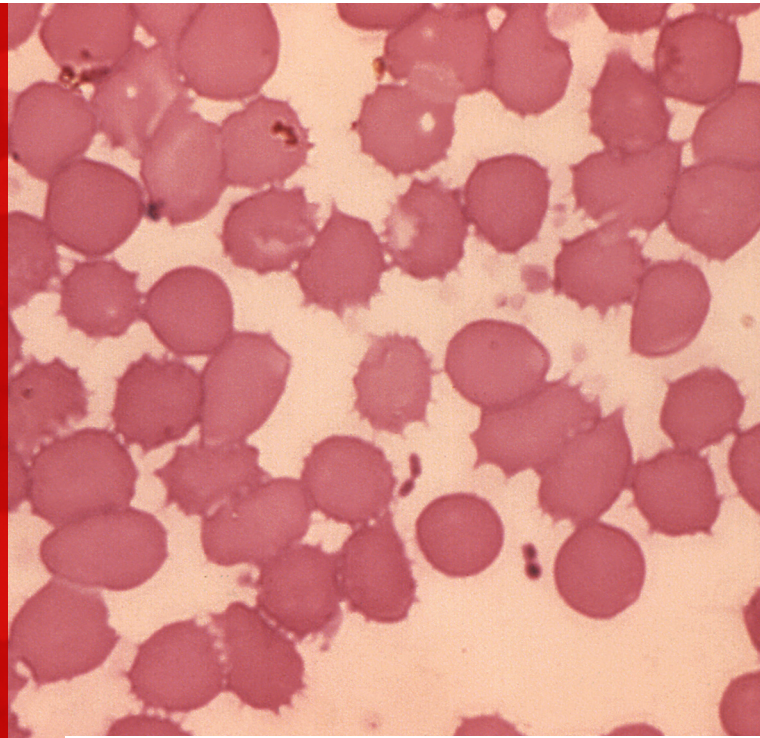


Plague in Cats & Dogs: A Public Health Concern

Radford G. Davis, DVM, MPH, DACVPM
Iowa State University



▲ **FIGURE** Blood smear revealing the presence of gram-negative *Yersinia pestis*. Image courtesy of Centers for Disease Control and Prevention

Background

Plague is a flea-borne zoonotic disease caused by the bacterium *Yersinia pestis*, which has been found on all continents except Australia and Antarctica.¹⁻⁴ Most human cases of plague occur in Africa, particularly the Democratic Republic of Congo and Madagascar, and Peru.¹ At least 17 US states have documented *Y pestis* infection in mammals or fleas.^{3,5} Most human cases in the United States occur in the southwest region (eg, New Mexico, Arizona, Colorado), followed by the Pacific region (eg, California, Oregon, Nevada).^{3,6,7}

Y pestis is maintained in a sylvatic enzootic cycle in rural and semirural areas, primarily through flea transmission among certain wild rodents that experience low mortality rates.^{3,8,9} More than 200 species of mammals are susceptible to infection by *Y pestis*; morbidity and mortality varies by species, among individuals in a population,³ and by geographic location.^{3,10} Several species are more suscepti-

ble to plague and readily experience die-offs. These are referred to as “amplifying hosts.” Prairie dogs (*Cynomys* spp) are highly susceptible, as are California ground squirrels (*Spermophilus beecheyi*) and rock squirrels (*Spermophilus variegatus*), both of which exhibit lower mortality rates than prairie dogs.³ More resistant species include California voles (*Microtus californicus*), although some variation in resistance has been documented.^{3,10} Susceptibility of *Peromyscus* mice varies by species.^{11,12} Large die-offs of chipmunks, wood rats, and various species of ground squirrels have been documented.³ In the sylvatic epizootic cycle, outbreaks occur in more susceptible rodent species with consequential higher mortality rates^{8,9}; these epizootics carry a high risk for infection in dogs, cats, and humans, as fleas search for new hosts after the die-off of their original hosts. This can result in a geographic hotbed of plague that should be avoided.^{8,13,14} Urban outbreaks can occur when *Y pestis* spreads from wild rodents to rats living near human habitation.⁴

Cats and dogs can become infected with *Y pestis* through consumption of infected small mammals and via flea bites.^{13,15,16} Infection in humans occurs primarily from flea bites⁸ but can occur through consumption^{3,17} or handling of infected animal tissue or body fluids^{3,18-21}; direct or indirect contact with infectious exudates²²; inhalation of respiratory droplets from humans, cats, or dogs with pneumonic plague^{5,8,18,23-26}; cat bites^{5,27} and scratches^{5,28}; and dog bites.²¹ Cats are more susceptible to plague than are dogs and pose a greater public health risk to humans.²⁹ From 1977 to 1998, 23 cases of cat-associated plague—some involving veterinary staff members—occurred in 8 US states.⁵ Increased risk to humans comes from close contact with cats and dogs (eg, sharing a bed)^{15,30,31} and from pets bringing plague-infected fleas into the home.^{8,21,31}

Clinical Signs

The primary clinical manifestations of plague include bubonic (ie, enlarged lymph nodes), septicemic, and pneumonic. Cats and humans manifest clinical signs and pathologic changes similarly.^{16,32-34} If plague is left untreated and death does not occur, disease often progresses from bubonic to septicemic to secondary pneumonic plague; however, the bubonic form does not always manifest.^{16,33,35,36} These stages may occur in dogs as well, although dogs are less likely than cats to develop illness.^{22,37} Bubonic plague results from a flea bite³² or, in cats and possibly dogs, from consuming infected prey.¹⁶ Septicemia can lead to secondary pneumonic plague. Cats

with secondary pneumonic plague can shed *Y pestis* in respiratory droplets via breathing, coughing, or sneezing, which can lead to primary pneumonic plague in humans. Pneumonic plague is the most serious stage, as it is the only form that can spread directly from human to human.³²

The incubation period for *Y pestis* in cats has been estimated to be 1 to 4 days.³⁸ Clinical signs include fever, anorexia, lethargy, lymphadenitis, and lymphadenopathy (buboes), with 75% of patients exhibiting submandibular lymphadenopathy.^{29,33} Lymph nodes may be abscessed or show evidence of hemorrhage or necrosis.^{33,35} Abscesses in many other areas of the body have also been reported.³³ Signs of more advanced disease include vomiting, diarrhea, discharge from the mouth or nose, coughing, sneezing, ataxia, dehydration, weak pulse, cold extremities, pale or brick-red mucous membranes, prolonged capillary refill time, disseminated intravascular coagulation, dyspnea, pneumonia, and coma.^{16,33,34} Approximately 10% of feline cases are pneumonic.^{5,29,33} If left untreated, mortality rates can approach 41%.³³

Infected dogs are often subclinically infected or have only mild, self-limiting febrile illness, but severe disease and death have been reported.^{25,39} Clinical signs in dogs include fever, lethargy, anorexia, submandibular lymphadenitis, lymphadenopathy, abscesses, oral cavity lesions, purulent intramandibular lesions, coughing, bloody sputum, ataxia, vomiting, diarrhea, dyspnea, and harsh lung sounds.^{13,39} The mortality rate in dogs has been estimated to be approximately 3%.³⁹

Diagnosis

Clinicians who suspect plague in a patient should immediately notify state health officials, who can assist in facilitating diagnostic testing.^{16,22} Ideally, samples should be collected prior to antibiotic administration,³⁵ but treatment should never be delayed while waiting for laboratory results. *Y pestis* may be identified microscopically by staining

Cats are more susceptible to plague than are dogs and pose a greater public health risk to humans.²⁹

smears of peripheral blood, sputum, or lymph node material.²² Visualizing bipolar-staining, gram-negative bacteria (**Figure**, page 57) with a “safety pin” appearance can provide a quick presumptive diagnosis.²² Antemortem testing should include culture of whole blood, lymph node aspirates, swabs from draining lesions, and oropharyngeal swabs from patients with oral lesions or pneumonia.^{16,38} A slide smear of a lymph node aspirate can also be used for detecting the F1 antigen on *Y pestis*.³⁸ Recent infection can be confirmed by serologic specimens collected 14 to 21 days apart that demonstrate a 4-fold rise in antibody titers.^{16,38} Postmortem specimens for testing should include liver, spleen, lungs, bone marrow, whole blood, and lymph nodes.²²

Management & Public Health Considerations

Animals suspected of having plague should not be immediately released from the clinical setting, as they pose a risk to the owners, and instead should be placed in isolation and treated using strict infection prevention procedures.^{16,38} A rapid-acting flea control product should be applied to hospitalized animals to prevent fleas from circulating in the clinic and transmitting *Y pestis* to patients and staff. Contact with exudates, respiratory droplets, oral secretions, tissues, or fleas carries a risk for infection.²² When working with a plague patient, or before beginning necropsy on a suspected case, veterinary staff should use personal protective equipment, including a gown, gloves, an N95 respirator, and protective eyewear, and adhere to hygiene and disinfection protocols.^{5,16,22,38} Cats are considered noninfectious after 72 hours of antibiotic therapy and when showing clinical improvement,^{16,38} at which time personal protective equipment can be relaxed to standard precautions.^{16,38}

Once alerted to a suspected case of plague by the clinician, health officials will conduct a home and environmental investigation, implement prevention measures at the source to prevent future cases, and work with pet owners and veterinary staff to

ensure proper medical evaluation and antibiotic prophylaxis.^{16,22,38} Because of the risk for droplet exposure, any human within 6 feet of an infected dog or cat should be medically evaluated.^{5,21}

Treatment

Because of the disease’s rapid progression, treatment with antibiotics (eg, gentamicin [IM or SC^{22,38}], doxycycline [PO], tetracycline [PO], chloramphenicol [PO]) should be initiated immediately, before a definitive diagnosis of plague has been obtained.^{16,38} Fluoroquinolones may also be effective against *Y pestis*, but supporting data are sparse to warrant their use over gentamicin.^{16,38} Oral antibiotics are only appropriate for less severe cases and as continued therapy after clinical improvement under parenteral gentamicin.²² Oral antibiotics can be administered once the clinical condition improves,¹⁹ usually within 72 hours.¹⁶ Patients should be treated with antibiotics for 10 to 21 days.³⁸

Prevention

To decrease the risk to pets and humans, flea control and prevention should be regularly applied and pets should be prevented from roaming or hunting outdoors.^{16,21,38} Brush, firewood, rock piles, clutter, garbage, and food sources (including pet food) around the home should be eliminated to reduce rodent infestation.^{18,22} Homes should also be made rodent-proof. Humans should avoid ill or dead animals, avoid feeding wildlife, and, when spending time outdoors, wear long pants and use insect repellent on skin and clothing.^{18,21}

Conclusion

Cats and, less commonly, dogs with plague pose a serious public health concern. Clinicians living in enzootic areas may be the first to recognize plague and have the opportunity to intervene and prevent human infections.³⁹ In addition to prompt recognition and treatment, robust application of infection prevention measures and early involvement of public health officials are vital in protecting human and animal life. ■

References

- World Health Organization. Plague: fact sheet. WHO website. <http://www.who.int/mediacentre/factsheets/fs267/en>. Published October 31, 2017. Accessed July 27, 2018.
- Dubyanskiy VM, Yeszhanov AB. Ecology of *Yersinia pestis* and the epidemiology of plague. *Adv Exp Med Biol*. 2016;918:101-170.
- Abbott RC, Rocke TE. Plague. https://pubs.usgs.gov/circ/1372/pdf/C1372_Plague.pdf. Published 2012. Accessed July 26, 2018.
- Dennis DT, Gage KL, Gratz N, Poland JD, Tikhomirov E. Plague manual: epidemiology, distribution, surveillance and control. http://www.who.int/csr/resources/publications/plague/WHO_CDS_CSR_EDC_99_2_EN/en. Published 1999. Accessed July 26, 2018.
- Gage KL, Dennis DT, Orloski KA, et al. Cases of cat-associated human plague in the Western US, 1977-1998. *Clin Infect Dis*. 2000;30(6):893-900.
- Centers for Disease Control and Prevention. Plague in the United States. CDC website. <https://www.cdc.gov/plague/maps/index.html>. Updated January 4, 2018. Accessed July 27, 2018.
- Kugeler KJ, Staples JE, Hinckley AF, Gage KL, Mead PS. Epidemiology of human plague in the United States, 1900-2012. *Emerg Infect Dis*. 2015;21(1):16-22.
- Plague: ecology and transmission. Centers for Disease Control and Prevention. <https://www.cdc.gov/plague/transmission/index.html>. Accessed July 27, 2018.
- Gage KL, Kosoy MY. Natural history of plague: perspectives from more than a century of research. *Annu Rev Entomol*. 2005;50:505-528.
- Hubbert WT, Goldenberg MI. Natural resistance to plague: genetic basis in the vole (*Microtus californicus*). *Am J Trop Med Hyg*. 1970;19(6):1015-1019.
- Quan SF, Kartman L. The resistance of *Microtus* and *Peromyscus* to infection by *Pasteurella pestis*. *Trans R Soc Trop Med Hyg*. 1956;50(1):104-105.
- Holdenried R, Quan SF. Susceptibility of New Mexico rodents to experimental plague. *Public Health Rep*. 1956;71(10):979-984.
- Orloski KA, Eidson M. *Yersinia pestis* infection in three dogs. *J Am Vet Med Assoc*. 1995;207(3):316-318.
- Mead PS. *Yersinia* species (including plague). In: Bennett J, Dolin R, Blazer M, eds. *Mandell, Douglas, and Bennett's Principles and Practice of Infectious Diseases*. 8th ed. Philadelphia, PA: Elsevier Saunders; 2015:2607-2618.
- Eidson M, Tierney LA, Rollag OJ, Becker T, Brown T, Hull HF. Feline plague in New Mexico: risk factors and transmission to humans. *Am J Public Health*. 1988;78(10):1333-1335.
- Orloski KA, Lathrop SL. Plague: a veterinary perspective. *J Am Vet Med Assoc*. 2003;222(4):444-448.
- Bin Saeed AA, Al-Hamdan NA, Fontaine RE. Plague from eating raw camel liver. *Emerg Infect Dis*. 2005;11(9):1456-1457.
- Kwit N, Nelson C, Kugeler K, et al. Human plague - United States, 2015. *MMWR Morb Mortal Wkly Rep*. 2015;64(33):918-919.
- Wong D, Wild MA, Walburger MA, et al. Primary pneumonic plague contracted from a mountain lion carcass. *Clin Infect Dis*. 2009;49(3):e33-e38.
- von Reyn CF, Barnes AM, Weber NS, Hodgins UG. Bubonic plague from exposure to a rabbit: a documented case, and a review of rabbit-associated plague cases in the United States. *Am J Epidemiol*. 1976;104(1):81-87.
- Centers for Disease Control and Prevention. Human plague—four states, 2006. *MMWR Morb Mortal Wkly Rep*. 2006;55(34):940-943.
- Centers for Disease Control and Prevention. Plague: information for veterinarians. CDC website. <https://www.cdc.gov/plague/healthcare/veterinarians.html>. Updated September 27, 2016. Accessed July 26, 2018.
- Wang H, Cui Y, Wang Z, et al. A dog-associated primary pneumonic plague in Qinghai Province, China. *Clin Infect Dis*. 2011;52(2):185-190.
- Werner SB, Weidmer CE, Nelson BC, Nygaard GS, Goethals RM, Poland JD. Primary plague pneumonia contracted from a domestic cat at South Lake Tahoe, Calif. *J Am Med Assoc*. 1984;251(7):929-931.
- Runfola JK, House J, Miller L, et al. Outbreak of human pneumonic plague with dog-to-human and possible human-to-human transmission — Colorado, June–July 2014. *MMWR Morb Mortal Wkly Rep*. 2015;64(16):429-434.
- Doll JM, Zeitz PS, Etestad P, Bucholtz AL, Davis T, Gage K. Cat-transmitted fatal pneumonic plague in a person who traveled from Colorado to Arizona. *Am J Trop Med Hyg*. 1994;51(1):109-114.
- Thornton DJ, Tustin RC, Pienaar BJ, Pienaar WN, Bubb HD. Cat bite transmission of *Yersinia pestis* infection to man. *J S Afr Vet Assoc*. 1975;46(2):165-169.
- Weniger BG, Warren AJ, Forseth V, et al. Human bubonic plague transmitted by a domestic cat scratch. *J Am Med Assoc*. 1984;251(7):927-928.
- Kassem AM, Tengelsen L, Atkins B, et al. Notes from the field: plague in domestic cats - Idaho, 2016. *MMWR Morb Mortal Wkly Rep*. 2016;65(48):1378-1379.
- Centers for Disease Control and Prevention. Two cases of human plague—Oregon, 2010. *MMWR Morb Mortal Wkly Rep*. 2011;60(7):214.
- Gould LH, Pape J, Etestad P, Griffith KS, Mead PS. Dog-associated risk factors for human plague. *Zoonoses Public Health*. 2008;55(8-10):448-454.
- Centers for Disease Control and Prevention. Plague: symptoms. CDC website. <https://www.cdc.gov/plague/symptoms/index.html>. Updated September 14, 2015. Accessed January 31, 2018.
- Eidson M, Thilsted JP, Rollag OJ. Clinical, clinicopathologic, and pathologic features of plague in cats: 119 cases (1977-1988). *J Am Vet Med Assoc*. 1991;199(9):1191-1197.
- Watson RP, Blanchard TW, Mense MG, Gasper PW. Histopathology of experimental plague in cats. *Vet Pathol*. 2001;38(2):165-172.
- Gasper PW, Barnes AM, Quan TJ, et al. Plague (*Yersinia pestis*) in cats: description of experimentally induced disease. *J Med Entomol*. 1993;30(1):20-26.
- Centers for Disease Control and Prevention. Plague: resources for clinicians. CDC website. <https://www.cdc.gov/plague/healthcare/clinicians.html>. Updated October 5, 2015. Accessed January 31, 2018.
- Etestad P. Plague in dogs. Merck Veterinary Manual website. <https://www.merckvetmanual.com/dog-owners/disorders-affecting-multiple-body-systems-of-dogs/plague-in-dogs#v3208803>. Accessed July 31, 2018.
- Etestad P. Overview of plague. Merck Veterinary Manual website. <https://www.merckvetmanual.com/generalized-conditions/plague/overview-of-plague#v9162260>. Accessed July 31, 2018.
- Nichols MC, Etestad PJ, Vinhaton ES, et al. *Yersinia pestis* infection in dogs: 62 cases (2003-2011). *J Am Vet Med Assoc*. 2014;244(10):1176-1180.