# GEORGIA Rabies Control Manual

May 2012 | Sixth Edition





Epidemiology Program | Department of Public Health

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#### **Foreword**

The purpose of this manual is to provide current information on the control of rabies in Georgia. It is designed to be used by county health departments, hospital emergency departments, private physicians and health care practitioners, veterinarians, and animal control programs. This manual should serve as an educational tool for use in all facets of community rabies control. Additionally, it is hoped that this manual will assist communities in standardizing rabies control practices within the state.

This document was prepared by Cherie L. Drenzek, DVM, MS, Julie Gabel, DVM, MPH, and Melissa Ivey, MPH. Credit is also given to authors of the following: 1) *Georgia Rabies Control Manual*, Third, Fourth, and Fifth Editions (1996, 2001, 2007); 2) National Association of State Public Health Veterinarians (NASPHV) *Compendium of Animal Rabies Prevention and Control 2011*, and 3) Use of a Reduced (4-Dose) Vaccine Schedule for Postexposure Prophylaxis to Prevent Human Rabies - Recommendations of the Advisory Committee on Immunization Practices (2010).

If you have any questions regarding this manual, please contact the Acute Disease Section, Epidemiology Program, Georgia Department of Public Health at (404) 657-2588.

#### **Important Phone Numbers**

#### **RABIES CONSULTATIONS**

404-616-9000
800-222-1222
See local phone directory
See local phone directory
404-657-2588
800-CDC-INFO (800-232-4636)

#### STATE PUBLIC HEALTH LABORATORIES

Georgia Public Health Laboratory (Decatur)	404-327-7900
Albany Regional Laboratory	229-430-4122
Waycross Regional Laboratory	912-285-6000

#### SOURCES FOR RABIES VACCINE

sanofi pasteur (Imovax<sup>®</sup> Rabies - HDCV)

Novartis Vaccines and Diagnostics (RabAvert<sup>®</sup> - PCEC) 800-VACCINE (800-822-2463) www.vaccineplace.com/products/

800-CHIRON8 (800-244-7668) www.rabavert.com

#### SOURCES FOR RABIES IMMUNE GLOBULIN

sanofi pasteur (Imogam® Rabies-HT) 800-VACCINE (800-822-2463) www.vaccineplace.com/products/

Talecris Biotherapeutics (HyperRab<sup>™</sup> S/D)

800-243-4153 www.talecris-pi.info

#### INDIGENT PATIENT RABIES VACCINE SUPPORT PROGRAMS

Both rabies vaccine manufacturers have patient assistance programs that provide vaccines and medications to uninsured and underinsured patients. These programs are administered through the *Rx Assist Patient Assistance Program Center* (www.rxassist.org/patients/default.cfm). The manufacturers may also be contacted directly for more information concerning eligibility requirements.

sanofi pasteur (Imovax <sup>®</sup> Rabies and Imogam <sup>®</sup> Rabies-HT)	866-801-5655
Novartis Vaccines and Diagnostics (RabAvert <sup>®</sup> )	800-589-0837

#### SEROLOGIC TESTING FOR HUMANS AND ANIMALS (see pages 37-38)

Atlanta Health Associates, Inc. 309 Pirkle Ferry Road, Suite D300 Cumming, GA 30040

Auburn University College of Veterinary Medicine Department of Pathobiology Virology Laboratory 261 Greene Hall Auburn University, AL 36849 ALS (see pages 37-36) Phone: 800-717-5612 Fax: 770-205-9021 http://atlantahealth.net

Phone: 331-844-2659 Fax: 334-844-2652 www.vetmed.auburn.edu/virology

Kansas State UniversityPhone: 785-532-4483College of Veterinary MedicineFax: 785-532-4474Veterinary Diagnostic Laboratory2005 Research Park CircleManhattan, Kansas 66502www.vet.ksu.edu/depts/dmp/service/rabies/index.htm

#### **RABIES TAGS**\*

Dogs, cats, and ferrets should be identified (e.g., metal or plastic tags or microchips) to allow for verification of rabies vaccination status.

\*Licenses/rabies tag requirements are County-based; please contact your County for specifics.

#### I. RABIES OVERVIEW

Rabies is a viral infection transmitted in the saliva of infected mammals. The virus enters the central nervous system of the host, causing an encephalomyelitis that is almost always fatal. Although all species of mammals are susceptible to rabies virus infection, only a few species are important as reservoirs for the disease in nature. In the United States, several distinct rabies virus variants have been identified in terrestrial mammals, including major terrestrial reservoirs in raccoons, skunks, foxes, and coyotes. In addition to the terrestrial reservoirs for rabies, several species of insectivorous bats also serve as reservoirs for the disease.

Wildlife is the most important potential source of infection for both humans and domestic animals in the United States. Reducing the risk of rabies in domestic animals and limiting contact with wild animals are central to the prevention of human rabies. Vaccination of all domestic dogs, cats, and ferrets, coupled with the systematic removal of stray animals that are at risk of exposure to rabid wildlife, are basic elements of a rabies control program. Georgia law (Rabies Control Law-O.C.G.A-31-19) requires that all owned dogs and cats be vaccinated against rabies by a licensed veterinarian using approved vaccines in accordance with the national *Compendium of Animal Rabies Prevention and Control* (see pages 51-64). Domestic ferrets also need to be vaccinated against rabies according to the national *Compendium of Animal Rabies Prevention and Control* (see pages 51-64) and Georgia law (O.C.G.A-27-5-5).

In the United States, indigenously acquired rabies among humans has declined markedly in recent years. The decline is, in part, due to vaccination and animal control programs begun in the 1940s that have practically eliminated the domestic dog as a reservoir of rabies and also to the development of effective human rabies vaccines and rabies immune globulin. During 2000-2008, a total of 31 cases of human rabies were reported in the United States (including one case in Georgia in 2000). Among the 28 cases for which rabies virus variants were obtained, 20 (71%) were associated with insectivorous bats, most commonly the Mexican free-tailed, silverhaired, and eastern pipistrelle bats. More than half of these human cases occurred during August-November, coincident with a seasonal increase in prevalence of rabid bats detected in the United States. Despite the substantial number of cases of human rabies attributable to bat exposure, the importance of these exposures is often overlooked or underestimated. In many of these cases, the bat bite was presumably not recognized nor the risk of rabies appreciated in order to seek appropriate medical attention.

Human rabies is a completely preventable disease if the risk of acquisition is appreciated and appropriate rabies post-exposure prophylaxis (consisting of wound care as well as both active and passive immunization) is obtained. Because rabies is a fatal disease, the goal of public health (in coordination with the medical community) is, first, to prevent human exposure to rabies by education and animal control measures and, second, to prevent the disease by administering rabies postexposure prophylaxis (PEP) if exposure occurs. Tens of thousands of people are successfully treated each year after being bitten by an animal that may have rabies. Although the decision to provide post-exposure prophylaxis rests with the patient and his or her physician, valuable consultations can be provided by the Georgia Poison Center, District and County health departments, or the Epidemiology Program, Department of Public Health (see page 2 for contact information).

#### **II. RABIES PREVENTION AND CONTROL**

#### A. Legal Authority

The primary responsibility for the control of rabies in Georgia rests with County Boards of Health. Chapter 31-19-1 of the Official Code of Georgia Annotated (O.C.G.A.) empowers and requires each County Board of Health to adopt and promulgate rules and regulations for the prevention and control of rabies (see pages 48-50).

#### **B. Principles of Rabies Control**

As a zoonotic disease, the foundations of rabies control rest upon preventing the disease in animals, preventing the disease in humans, and decreasing the likelihood of exposure between humans and animal rabies vectors. Public education regarding rabies exposure risk is paramount. The following principles apply:

- **Rabies Exposure.** Rabies is transmitted only when the virus is introduced into bite wounds, open cuts in skin, or onto mucous membranes.
- Human Rabies Prevention. Rabies in humans can be prevented either by eliminating exposures to rabid animals or by providing exposed persons with prompt local treatment of wounds combined with appropriate post-exposure prophylaxis (including both passive antibody administration and active immunization with cell culture vaccines). In addition, pre-exposure vaccination should be offered to persons in high-risk groups, such as veterinarians, animal handlers, and certain laboratory workers.
- Domestic Animals. Local governments should initiate and maintain effective programs to ensure vaccination of all dogs, cats, and ferrets and to remove strays and unwanted animals from the community. Recommended vaccination procedures and the licensed animal vaccines are specified in the *Compendium of Animal Rabies Prevention and Control* (see pages 51-62). In addition, adjunct procedures which enhance rabies control include: 1) identification systems (e.g., metal/plastic tags, microchips; please refer to individual County requirements) to verify animal rabies vaccination status; 2) local domestic animal licensure requirements; 3) requirement of interstate health certificates prior to domestic animal travel; 4) implementation of regulations governing imported domestic animals; and 5) establishment of a local animal control agency responsible for stray control, leash laws, and issuance of citations for failure to vaccinate animals.
- **Rabies in Wildlife.** The control of rabies among wildlife reservoirs is difficult. Vaccination of free-ranging wildlife or selective population reduction is not always feasible. Rabies control relies upon prevention of exposure to wildlife rabies reservoirs. This can be accomplished via public

education about wildlife rabies risk and recommendations regarding avoidance of contact with wild animals. Leash laws and other control of domestic animals will reduce exposure of pets to potentially rabid wildlife.

#### C. CONTROL METHODS IN ANIMALS

#### Animal Vaccination Protocols

Parenteral animal rabies vaccines should be administered only by a licensed veterinarian. This is the only way to ensure that a responsible person can be held accountable and to assure the public that the animal has been properly vaccinated. Within 28 days after primary vaccination, a peak rabies antibody titer is reached, and the animal can be considered immunized. An animal is currently vaccinated and is considered immunized if the primary vaccination was administered at least 28 days previously and vaccinations have been administered in accordance with the *Compendium of Animal Rabies Prevention and Control* (see pages 51-64). Regardless of the age of the animal at initial vaccination, a second vaccination should be administered 1 year later. Because a rapid anamnestic response is expected, an animal is considered currently vaccinated <u>immediately</u> after a booster vaccination.

- **Dogs, Cats, and Ferrets.** All dogs, cats, and ferrets should be vaccinated against rabies and revaccinated in accordance with the *Compendium of Animal Rabies Prevention and Control* (see pages 59-60). For many licensed vaccines, the age at primary vaccination is 3 months, but be aware that for some newer combination rabies vaccines, this age is 8 weeks. If a previously vaccinated animal is overdue for a booster, it should be revaccinated with a single dose of vaccine and placed on an annual or triennial schedule, depending on the type of vaccine used.
- Livestock. Vaccinating all livestock against rabies is neither economically feasible nor justified from a public health standpoint. However, livestock that are particularly valuable or that have frequent contact with humans, such as show animals or those in petting zoos, should be vaccinated against rabies (refer to *the Compendium of Animal Rabies Prevention and Control* for specific vaccines licensed for use in livestock, pages 59-60). Horses traveling interstate or with significant public contact (e.g., riding stables) should be currently vaccinated against rabies.
- Other Animals.

**Wild.** No parenteral rabies vaccine is licensed for use in wild animals. Because of the risk for rabies in wild animals (especially raccoons, skunks, coyotes, foxes, and bats), the Georgia Department of Natural Resources has rigid regulations which prohibit the keeping of wild and wild/domestic hybrids as pets. For further information, please see <u>www.dnr.state.ga.us</u>. Maintained in Exhibits and in Zoological Parks. Captive animals that are not completely excluded from all contact with rabies vectors can become infected with rabies. Moreover, wild animals might be incubating rabies when initially captured; therefore, wild-caught animals susceptible to rabies should be placed in strict isolation for a minimum of 6 months before being exhibited. Employees who work with animals at such facilities should receive pre-exposure rabies vaccination. The use of pre- or post-exposure rabies vaccinations for employees who work with animals at such facilities might reduce the need for euthanasia of captive animals. Carnivores and bats should be housed in a manner that precludes direct contact with the public.

#### Management of Animals Exposed to Rabies

Any animal potentially exposed to rabies virus by a wild, carnivorous mammal or a bat that is not available for testing should be regarded as having been exposed to rabies.

#### Dogs, Cats, and Ferrets

- **Unvaccinated** dogs, cats, and ferrets exposed to a rabid animal should be euthanized immediately. If the owner is unwilling to have this done, the animal should be placed in strict isolation for 6 months and vaccinated either upon entry to isolation OR 1 month before being released. Isolation in this context refers to confinement in an enclosure that precludes direct contact with humans and other animals. Animals overdue for a booster are evaluated on a case-by-case basis that should consider the severity of exposure, time elapsed since the animal's last vaccination, number of prior vaccinations, current health status of the animal, and local rabies epidemiology. Strict isolation should be conducted under the authority of the designated local rabies control agency in which the place, manner, and provisions of the confinement are specified. For example, strict isolation may take place in an animal control facility or an isolation pen at home, depending on local requirements. At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies.
- **Currently vaccinated** (see Definitions, pages 46-47) dogs, cats, and ferrets should be revaccinated immediately, kept under the owner's control, and observed at home for 45 days for clinical signs of rabies. During the observation period (see Definitions, pages 46-47) the animal should not be permitted to roam freely and should be restricted to leash walks, if applicable. At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies.

#### Livestock

- All species of livestock are susceptible to rabies; cattle and horses are the most frequently infected. Livestock exposed to a rabid animal and currently vaccinated with a vaccine approved by USDA for that species (see pages 59-60) should be revaccinated immediately and observed for 45 days.
- Unvaccinated livestock should be euthanized immediately. If the animal is not euthanized it should be kept under close observation for 6 months. Any illness in an animal under observation should be reported immediately to the local health department. If signs suggestive of rabies develop, the animal should be euthanized and the head shipped for testing as described in Part I.A.8.of the *Compendium of Animal Rabies Prevention and Control* (see page 53).
- Handling and consumption of tissues from exposed animals may carry a risk for rabies transmission. Risk factors depend in part on the site(s) of exposure, amount of virus present, severity of wounds, and whether sufficient contaminated tissue has been excised. If an exposed animal is to be slaughtered for consumption, it should be done immediately after exposure and all tissues should be cooked thoroughly.
- Barrier precautions should be used by persons handling the animal and tissues. Historically, federal guidelines for meat inspectors required that any animal known to have been exposed to rabies within 8 months be rejected for slaughter. USDA Food and Inspection Service (FSIS) meat inspectors should be notified if such exposures occur in food animals prior to slaughter.
- Rabies virus may be widely distributed in tissues of infected animals. Tissues and products from a rabid animal should not be used for human or animal consumption. However, pasteurization temperatures will inactivate rabies virus; therefore, drinking pasteurized milk or eating thoroughly cooked animal products does not constitute a rabies exposure.
- Multiple rabid animals in a herd or herbivore-to-herbivore transmission is uncommon; therefore, restricting the rest of the herd if a single animal has been exposed to or infected by rabies is usually not necessary.

#### **Other Animals**

• Other animals bitten by a rabid animal should be euthanized immediately. Animals maintained in USDA-licensed research facilities or accredited zoological parks should be evaluated on a case-by-case basis. Consultations can be provided by the Epidemiology Program, Department of Public Health.

#### Management of Animals that Bite Humans

#### Dogs, Cats, and Ferrets

- Rabies virus may be excreted in the saliva of infected dogs, cats, and ferrets during illness and/or for only a few days prior to illness or death. A healthy dog, cat, or ferret that bites a person should be confined (see Definitions, pages 46-47) and observed for 10 days, no matter if the animal is currently vaccinated or not. Administration of rabies vaccine is not recommended during the confinement period to avoid confusing signs of rabies with possible side effects of vaccine administration.
- Confinement (sometimes referred to as quarantine) conditions should prevent direct contact with other animals or persons. The confinement shall be conducted under the authority of the designated local rabies control agency in which the place, manner, and provisions of the confinement are specified. For example, confinement may take place in a kennel in a veterinary hospital, animal control facility, commercial boarding establishment, or a pen at home, depending on local requirements.
- At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies and the exposed person notified.
- Any stray or unwanted dog, cat, or ferret that bites a person may be euthanized immediately (or following the locally-specified impoundment period to give owners sufficient time to claim animals) and the head submitted for rabies examination.

Other biting animals (wild animals, animals maintained in zoological parks, canine or feline wild/domestic hybrids, etc.)

- No parenteral rabies vaccines are licensed for use in animals other than dogs, cats, ferrets, and some livestock.
- Since the duration of clinical signs and the period of virus shedding are unknown for many species, confinement may not be a feasible management strategy. Most wild mammals that bite or otherwise expose persons should be **considered** for euthanasia and rabies examination. Prior vaccination of an animal might not preclude the necessity for euthanasia and testing if the period of virus shedding is unknown for that species.

• Management of animals other than dogs, cats, and ferrets depends on the species, the circumstances of the bite, the epidemiology of rabies in the area, and the biting animal's history, current health status, and potential for exposure to rabies.

The Epidemiology Program, Department of Public Health, should be consulted when circumstances warrant.

#### Wildlife

- Most wild mammals that bite or otherwise expose persons should be considered for euthanasia and rabies examination. Since the duration of clinical signs and the period of virus shedding are unknown for these species, an appropriate confinement or isolation period cannot be ascertained. Assessing rabies risk and the need for rabies diagnostic testing can be guided by the following:
  - Wild Carnivores. Raccoons, skunks and foxes are the terrestrial animals most often infected with rabies. All bites by such wildlife must be considered possible exposures to the rabies virus. Signs of rabies among wildlife cannot be interpreted reliably; therefore, any such animal that exposes a person should be euthanized at once (without unnecessary damage to the head) and the brain should be submitted for rabies testing.
  - Rodents and Lagomorphs. Squirrels, rats, mice, hamsters, guinea pigs, gerbils, chipmunks, and rabbits are almost never found to be infected with rabies and have not been known to transmit rabies to humans. Bites by these animals are usually not considered a rabies risk and do not warrant rabies testing unless the animal is sick or behaving in an unusual manner. Rodents that are considered to be a rabies risk include woodchucks or groundhogs (*Marmota monax*) because they are frequently large enough to survive the attack of a rabid carnivore. Approval must be obtained from the Georgia Public Health Laboratory or the Epidemiology Program of the Department of Public Health prior to submitting a rodent for rabies testing.
  - **Bats.** A bat that bites, scratches, or has any direct physical contact with a person should be safely captured (see page 41 for instructions), immediately euthanized, and the entire animal sent to the laboratory for rabies examination. People usually know when they have been bitten by a bat. However, because bats have small teeth that may leave marks that are not easily seen, there are situations in which rabies testing and medical advice should be sought even in the absence of an obvious bite wound. These include awakening to find a bat in the room, finding a bat in the room of an unattended child, having a bat physically brush against you, or finding a bat near a mentally impaired or intoxicated person. In

these situations a bite cannot be definitively ruled out. If physical contact occurs or the situations above occur and the bat is not available for testing (i.e., escapes from house), rabies post-exposure prophylaxis should be administered as soon as possible.

• Other wild animals. In most situations involving non-reservoir species (opossums, otters, polecats, beavers, weasels, etc.), the rabies risk is relatively low. The risk is higher and, consequently, rabies testing may be indicated if the animal is found in a rabies-endemic area, has opportunity for exposure to rabies reservoirs, is large enough to survive an attack by a rabid animal, or is ill or exhibiting abnormal behavior (for example, many rabid bobcats have been found in Georgia).

### PROTOCOL FOR LIVESTOCK POSSIBLY EXPOSED TO RABIES



- \* Consultations regarding animal exposures can be provided by the Epidemiology Program of the Department of Public Health at 404-657-2588.
- \*\* An animal is currently vaccinated if the primary rabies vaccine (USDA-approved for use in livestock species) was administered by a veterinarian at least 28 days previously and booster vaccines have been administered according to vaccine label.
- \*\*\*If bat or wild animal is NOT available for testing, must proceed as if result is positive.

## PROTOCOL FOR DOGS, CATS, AND FERRETS POSSIBLY EXPOSED TO RABIES



- \* Consultations regarding animal exposures can be provided by the Epidemiology Program of the Department of Public Health at 404-657-2588.
- \*\* An animal is currently vaccinated if the primary rabies vaccine was administered by a veterinarian at least 28 days previously and booster vaccines have been administered on an annual or triennial schedule. Animals overdue for a booster are evaluated on a case by case basis (e.g., severity of exposure, time elapsed since last vaccination, number of prior vaccinations, current health status, and local rabies epidemiology).
- \*\*\*If bat, attacking dog, or wild animal is NOT available for testing, must proceed as if result is positive.

#### PROTOCOL FOR COMPANION ANIMAL-TO-COMPANION ANIMAL EXPOSURES/ENCOUNTERS

Note: Because the United States has been declared free of canine rabies virus variant transmission, healthy dog-to-dog, dog-to-cat, or cat-to-cat encounters are not generally considered a rabies risk.





- \* Consultations regarding exposure can be provided by the Georgia Poison Center, 24 hours a day, 7 days a week, at 1-800-222-1222 or 404-616-4000.
- \*\* No parenteral rabies vaccines are licenses for use in wild animals or hybrids (the offspring of wild animals crossbred to domestic animals). Wild animals or hybrids should not be kept as pets. Prior vaccination of these animals does not preclude the necessity for euthanasia and testing.
- \*\*\* The following animals are NOT CONSIDERED LIKELY TO HAVE RABIES and will not be tested except by special arrangements with the Epidemiology Program of the Georgia Department of Public Health at 404-657-2588: chipmunk, gerbil, gopher, guinea pig, hamster, hare, mole, mouse, rabbit, rat, shrew, squirrel, and vole.

#### D. CONTROL METHODS IN HUMANS

Prevention of human rabies depends on eliminating exposure to rabid animals and providing exposed persons with prompt local treatment of their wounds, combined with appropriate rabies post-exposure prophylaxis (PEP) consisting of both passive antibody administration and immunization with cell culture vaccines. In addition, pre-exposure vaccination is recommended for persons in high-risk groups, such as veterinarians, animal handlers, and certain laboratory workers.

#### **Rabies Biologics**

In general, two types of rabies products are available in the United States, namely, rabies vaccines and rabies immune globulin. Rabies vaccines induce an active immune response that includes the production of virus neutralizing antibodies. This antibody response requires approximately 7-10 days to develop and usually persists for several years. Rabies immune globulin (RIG) provides a rapid, passive immunity that persists for only a short time (half-life of approximately 21 days) to bridge the gap until the production of active immunity in response to vaccine administration.

Two formulations of inactivated rabies vaccines are currently licensed for preexposure and post-exposure prophylaxis in the United States (see below). When used as indicated, both types of rabies vaccines are considered equally safe and efficacious. A full 1.0-mL intramuscular (IM) dose is used for both pre-exposure and post-exposure prophylaxis. There are no currently approved formulations for the intradermal dose and route for pre-exposure vaccination; all must be administered intramuscularly. Usually, an immunization series is initiated and completed with one vaccine product. No clinical studies were identified that document a change in efficacy or the frequency of adverse reactions when the series is completed with a second vaccine product.

Two rabies immune globulin (RIG) formulations are currently licensed and available in the United States (see below). In all post-exposure prophylaxis regimens, except for persons previously vaccinated, RIG should be administered concurrently with the first dose of vaccine.

#### A. Vaccines

- 1. Human Diploid Cell Vaccine (HDCV): HDCV is prepared from the Pitman-Moore strain of rabies virus grown on MRC-5 human diploid cell culture, concentrated by ultrafiltration, and inactivated with betapropiolactone. HDCV is formulated for IM administration in a single-dose vial containing lyophilized vaccine that is reconstituted in the vial with the accompanying sterile diluent to a final volume of 1.0 mL just before administration. Once dose of reconstituted vaccine contains <150  $\mu$ g neomycin sulfate, <100 mg albumin, and 20  $\mu$ g of phenol red indicator. It contains no preservative or stabilizer.
  - <u>Manufacturer</u>: sanofi pasteur
  - <u>Product name:</u> Imovax<sup>®</sup>Rabies

- 2. Purified Chick Embryo Cell Vaccine (PCECV): PCECV became available in the United States in 1997. The vaccine is prepared from the fixed rabies virus strain Flury LEP grown in primary cultures of chicken fibroblasts. The virus is inactivated with betapropiolactone and further processed by zonal centrifugation in a sucrose density gradient. It is formulated for IM administration in a single-dose vial containing lyophilized vaccine that is reconstituted in the vial with the accompanying sterile diluent to a final volume of 1.0 mL just before administration. One dose of reconstituted vaccine contains <12 mg polygeline, <0.3 mg human serum albumin, 1 mg potassium glutamate, and 0.3 mg sodium EDTA. No preservatives are added.
  - Manufacturer: Novartis Vaccines and Diagnostics
  - <u>Product name</u>: RabAvert<sup>®</sup>

#### B. Rabies Immune Globulin (RIG)

The two RIG products licensed in the United States, HyperRab<sup>™</sup> S/D and Imogam<sup>®</sup> Rabies-HT, are immunoglobulin (IgG) preparations concentrated by cold ethanol fractionation from plasma of hyper-immunized human donors. Both RIG products are standardized at an average potency value of 150 IU per mL, and supplied in 2-mL (300 IU) vials for pediatric use and 10-mL (1,500 IU) vials for adult use. The recommended dose is 20 IU/kg (0.133mL/kg) body weight. Both RIG preparations are considered equally efficacious when used as described.

These products are made from the plasma of hyperimmunized human donors that, in theory, might contain infectious agents. Nevertheless, the risk that such products will transmit an infectious agent has been reduced substantially by screening plasma donors for previous exposure to certain viruses, by testing for the presence of certain current virus infections, and by inactivating and/or removing certain viruses. No transmission of adventitious agents has been documented after administration of RIGs licensed in the United States.

• <u>Product names</u>: Imogam<sup>®</sup> Rabies-HT (sanofi pasteur) and HyperRab<sup>™</sup> S/D (Talecris Biotherapeutics)

Biologic	Product Name	Manufacturer	Dose	Route	Indications
<i>Human Rabies Vaccine</i> Human diploid cell vaccine (HDCV)	Imovax <sup>®</sup> Rabies*	sanofi pasteur 800-822-2463 http://www.vaccinep	<b>1mL</b> lace.com/produ	Intramuscular	Pre-exposure or post-exposure <sup>†</sup>
Purified chick embryo cell vaccine (PCECV)	RabAvert <sup>®</sup>	Novartis Vaccines 800-244-7668 www.rabavert.com	and Diagnos	tics	
Rabies Immune Globulin	Imogam <sup>®</sup> Rabies-HT	sanofi pasteur 800-822-2463 http://www.vaccinepl	20 IU/kg ace.com/produc	Local	Post-exposure only⁵
	HyperRab™ S/D	eutics roducts i.info			
*Imovax rabies I.D., admii	nistered intradermally	, is no longer availa	able in the U	nited States.	

<sup>§</sup>As much of the product as is anatomically feasible should be infiltrated into and around the wound. Any remaining product should be administered intramuscularly in the deltoid or quadriceps (at a location other than that used for vaccine inoculation to minimize potential interference).

Source: CDC. Use of a reduced (4-dose) vaccine schedule for post-exposure prophylaxis to prevent human rabies - recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR 2010; 59(02);1-9.

#### Sources for Rabies Prophylactic Biologics

Large hospitals routinely stock rabies biologics (i.e., rabies vaccine and immune globulin) and healthcare providers can order biologics from the manufacturer. To obtain prophylaxis, consult your healthcare provider.

#### **Pre-Exposure Vaccination**

Pre-exposure vaccination should be offered to persons in high-risk groups, such as veterinarians and their staff, animal handlers, rabies researchers, and certain laboratory workers. Pre-exposure vaccination should also be considered for other persons whose activities bring them into frequent contact with rabies virus or potentially rabid bats, raccoons, skunks, cats, dogs, or other species at risk for having rabies. In addition, international travelers might be candidates for pre-exposure vaccination if they are likely to come in contact with animals in areas where dog or other animal rabies is enzootic and immediate access to appropriate medical care, including rabies vaccine and immune globulin, might be limited.

Pre-exposure prophylaxis is administered for several reasons. First, although preexposure vaccination does not eliminate the need for additional medical evaluation after a rabies exposure, it simplifies management by eliminating the need for RIG and decreasing the number of doses of vaccine needed. This is particularly important for persons at high risk for being exposed to rabies in areas where modern immunizing products might not be available or where cruder, less safe biologics might be used, placing the exposed person at increased risk for adverse events. Second, pre-exposure prophylaxis might offer partial immunity to persons whose post-exposure prophylaxis is delayed. Finally, pre-exposure prophylaxis might provide some protection to persons at risk for unrecognized exposures to rabies.

- Pre-exposure vaccination regimens are as follows:
  - A. Intramuscular Primary Vaccination
    - Three 1.0-mL injections of HDCV or PCECV should be administered intramuscularly (deltoid area) -- one injection per day on days 0, 7, and 21 or 28. Vaccine preparations for intradermal (ID) administration are no longer available in the United States.

Type of Vaccination	Route	Regimen
Primary	Intramuscular	HDCV or PCECV; 1.0 mL (deltoid area), one each on days 0*, 7, and 21 or 28
Booster <sup>†</sup>	Intramuscular	HDCV or PCECV; 1.0 mL (deltoid area), day 0* only
HDCV= human diploid cell	vaccine; PCECV = purifie	d chick embryo cell vaccine
<sup>†</sup> Persons in the continuous antibody every 6 months, intramuscular booster dose value of at least complete test.	-risk category should hav and persons in the freque e of vaccine should be ac neutralization at a 1:5 s	e a serum sample tested for rabies virus neutralizing ent-risk category should be tested every 2 years. An Iministered if the serum titer fails to maintain a erum diluation by rapid fluorescent focus inhibition
Source: CDC. Use of a redu human rabies - recomment 2010; 59(02);1-9.	uced (4-dose) vaccine sch dations of the Advisory C	nedule for post-exposure prophylaxis to prevent ommittee on Immunization Practices (ACIP). MMWR
<u>Note</u> : Beca recommen serologic t	ause the antibody responded pre-exposure propresenting to confirm seroco	nse has been satisfactory after these phylaxis vaccine regimens, routine poversion is not necessary except for

#### B. Pre-Exposure Booster Doses of Vaccine

Following completion of the pre-exposure primary vaccination regimen, certain persons whose activities bring them into frequent contact with rabies virus or potentially rabid animals may need a **booster** dose of vaccine if their rabies neutralizing antibody level falls below an acceptable level (i.e., if the titer is less than complete neutralization at a 1:5 serum dilution by the RFFIT). The following table provides guidelines based upon level of risk.

Risk Category	Nature of Risk	Typical Populations	Pre-Exposure Recommendations
Continuous	Virus present continuously, often in high concentrations. Specific exposures likely to go unrecognized. Bite, nonbite, or aerosol exposure.	Rabies research laboratory workers; Rabies biologics production workers	Primary course. Serologic testing' every 6 months. Booster vaccination if antibody titer is below acceptable level. <sup>†</sup>
Frequent	Exposure usually episodic, with source recognized, but exposure also might be unrecognized. Bite, nonbite, or aerosol exposure.	Rabies diagnostic laboratory workers; Cavers, Animal control and wildlife workers in areas where rabies is enzootic; Veterinarians and staff; All persons who handle bats	Primary course. Serologic testing* every 2 years. Booster vaccination if antibody titer is below acceptable level. <sup>†</sup>
Infrequent	Exposure nearly always episodic with source recognized. Bite or nonbite exposure.	Veterinarians and animal control staff working with terrestrial animals in areas where rabies is uncommon to rare; Veterinary students; Travelers visiting areas where rabies is enzootic and immediate access to appropri- medical care, including biolog is limited	Primary course. No serologic testing or booster vaccinations. ate
Rare	Exposure always episodic with source recognized. Bite or nonbite exposure.	U.S. population at large, Including persons in areas where rabies is enzootic	No vaccination necessary.

\*Refer to pages 37-38 for information about serologic testing.

<sup>†</sup>Minimum acceptable antibody level is complete virus neutralization at a 1:5 serum dilution by the rapid fluorescent focus inhibition test (RFFIT). A booster dose should be administered if the titer falls below this level.

Source: CDC. Use of a reduced (4-dose) vaccine schedule for postexposure prophylaxis to prevent human rabies - recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR 2010; 59(02);1-9.

#### C. Post-Exposure Prophylaxis for Previously Vaccinated Persons

If a person is exposed to rabies, local wound care remains an important part of post-exposure prophylaxis, even for previously vaccinated persons. If exposed to rabies, persons who have been previously vaccinated with either the recommended pre-exposure OR post-exposure regimen should receive **TWO** IM doses of vaccine (1.0 mL each in the deltoid), one immediately and one 3 days later. Administration of RIG is unnecessary and should <u>not</u> be administered to these persons because the administration of passive antibody might inhibit the relative strength or rapidity of an expected anamnestic (or "memory") immune response.

For previously vaccinated persons who are exposed to rabies, determining the rabies virus neutralizing antibody titer for decision-making about prophylaxis is inappropriate for at least three reasons. First, several days will be required to collect the serum and determine the test result. Second, no "protective" titer is known. Finally, although rabies virus neutralizing antibodies are important components, other immune effectors also are operative in disease prevention.

#### Post-Exposure Vaccination

In general, post-exposure prophylaxis (PEP) is indicated for persons exposed to a rabid animal in order to prevent infection with rabies virus. In the United States, the PEP regimen consists of local wound treatment, administration of one dose of immune globulin (with the exception of persons who have previously received complete vaccination regimens, either pre-exposure or post-exposure), and 4 doses of rabies vaccine over a 14-day period. Rabies immune globulin (RIG) and the first dose of rabies vaccine should be given as soon as possible after exposure. Additional doses of rabies vaccine should be given on days 3, 7, and 14 after the first vaccination. A 5dose regimen (days 0, 3, 7, 14, and 28) of rabies vaccine should be administered for persons with altered immunocompetence, as they may experience a substantially reduced immune response to rabies vaccines. See chart on the next page for specific schedule and administration instructions.

If RIG was not administered when vaccination was begun (i.e., day 0), it can be administered up to and including day 7 of the post-exposure prophylaxis series. Beyond the seventh day, RIG is not indicated because an antibody response to cell culture vaccine is presumed to have occurred.

Vaccination Status	Treatment	Regimen*
Not previously vaccinated	Local wound cleansing	PEP should <b>always</b> begin with immediate cleansing of all wounds with soap and water. If available, a virucidal agent such as a povidone-iodine solution should be used to irrigate the wounds.
	RIG	Administer 20 IU/kg body weight. If anatomically feasible, the <b>full</b> dose should be infiltrated around the wound(s and any remaining volume should be administered IM at an anatomical site distant from vaccine administration. RIG should <b>not</b> be administered in the same syringe as vaccine. Because RIG might partially suppress active production of antibody, no more than the recommended dose should be given.
	Vaccine	HDCV or PCECV 1.0 mL, IM (deltoid area) <sup>\$</sup> , one each on days 0 <sup>#</sup> , 3, 7,and 14 <sup>‡</sup> (and 28 if person is immunocompromised).
Previously vaccinated <sup>†</sup>	Local wound cleansing	PEP should <b>always</b> begin with immediate cleansing of all wounds with soap and water. If available, a virucidal agent such as a povidone-iodine solution should be used to irrigate the wounds.
	RIG	RIG should not be administered.
	Vaccine	HDCV or PCECV 1.0 mL, IM (deltoid area) <sup>§</sup> , one each on days $0^{#}$ and $3^{\ddagger}$ .
*These regimens are applicable	for all age groups, including childr	en.
<sup>†</sup> Any person with a history of a rabies vaccine adsorbed, or preantibody response to the prior	complete pre-exposure or post-exp evious vaccination with any other ty vaccination.	osure vaccination regimen with HDCV, PCECV, or pe of rabies vaccine and a documented history of
<sup>s</sup> The deltoid area is the only ac outer aspect of the thigh may l	ceptable site of vaccination for ado be used. Vaccine should never be a	ults and older children. For younger children, the dministered in the gluteal area.
<sup>#</sup> Day 0 is the day the first dose	of vaccine is administered.	
<sup>‡</sup> Vaccination should correspond vaccine on the appropriate day person cannot receive the seco possible). The schedule of subs consistent with recommendatio days 8 and 15) to ensure adequ Georgia Department of Public H	with this regimen. However, in rar r, the time between doses should be nd dose of vaccine on day 3, it sho equent doses of vaccine should be ons (e.g., if vaccine was administer ate and proper immune response. I dealth at 404-657-2588 for consulta	re cases when it is not possible to administer the e lengthened rather than shortened (e.g., if a uld be administered as soon after day 3 as adjusted so that the time between doses is ed on day 4, subsequent doses should be given on Please contact the Epidemiology Program of the tions.
Source: CDC. Use of a reduced recommendations of the Adviso	(4-dose) vaccine schedule for post- ory Committee on Immunization Pra	exposure prophylaxis to prevent human rabies - actices (ACIP). MMWR 2010; 59(02);1-9.

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#### Assessing the Need for PEP

Administration of rabies PEP is a medical urgency, not a medical emergency. Persons who have been bitten by animals suspected or proven to be rabid should begin PEP as soon as possible. However, very long incubation periods (up to 1 year) have been reported in humans. Thus, when a documented or likely exposure has occurred, PEP is indicated regardless of the length of the delay, provided the clinical signs of rabies are not present. Under most circumstances, PEP should not be initiated while the biting, healthy dog, cat, or ferret is still in 10-day confinement. However, during the 10-day confinement period, begin PEP at the first sign of rabies in a dog, cat, or ferret that has bitten someone.

Healthcare providers should evaluate each possible exposure to rabies and when necessary consult with the Georgia Poison Center or public health officials regarding the need for rabies PEP.

In the United States, the following factors should be considered in the rabies risk assessment before PEP is initiated:

- type of exposure (bite or nonbite)
- the geographic location of the incident
- the type of animal that was involved
- circumstances of the exposure (provoked or unprovoked)
- the vaccination status of the animal
- whether the animal can be safely captured and tested for rabies

In general, the highest risk of rabies transmission is associated with bite exposure from terrestrial wild carnivores or bats (see **Decision Trees A-1** and **A-2**). Raccoons, skunks, and foxes are the terrestrial animals most often infected with rabies. Suggestive clinical signs of rabies among wildlife cannot be interpreted reliably. All bites by such wildlife must be considered possible exposures to the rabies virus. PEP should be initiated as soon as possible following exposure to wildlife, unless the animal is available for testing and shows no evidence of rabies (e.g., a negative test).

In addition, bats are increasingly implicated as important wildlife reservoirs for variants of rabies virus transmitted to humans. In all instances of potential human exposures involving bats, the bat in question should be safely collected, if possible, and submitted for rabies diagnosis. Rabies PEP is recommended for all persons with bite, scratch, or mucous membrane exposure to a bat, unless the bat is available for testing and shows no evidence of rabies (e.g., a negative test). PEP might also be appropriate even if a bite, scratch, or mucous membrane exposure is not apparent when there is reasonable probability that such exposure might have occurred (see pages 40-42 for more specific information about bats and rabies).

The likelihood of rabies in a domestic animal varies by region; hence, the need for PEP also varies. In the continental United States, rabies among dogs has been reported sporadically along the United States-Mexico border and in areas of the United States with enzootic wildlife rabies. During 2000-2006, more cats than dogs were reported rabid in the United States. The majority of these cases were associated with the epizootic of rabies among raccoons in the eastern United States. The large number of rabid cats compared with other domestic animals might be attributed to a lower vaccination rate among cats because of less stringent cat vaccination laws, fewer confinement or leash laws, and the nocturnal activity patterns of cats placing them at greater risk for exposure to infected raccoons, skunks, foxes, and bats. In certain developing countries, dogs remain the major reservoir and vector of rabies and represent an increased risk for rabies exposure in such countries.

In the United States, a currently vaccinated dog, cat, or ferret is unlikely to become infected with rabies (see **Decision Tree B**). Although all species of livestock are susceptible to rabies, they are infrequently found to be infected (see **Decision Tree C**). Cattle and horses are among the most frequently reported rabid livestock; in many cases these animals have a previously reported history of exposure to a wildlife rabies reservoir, such as raccoon, skunk, or bobcat.

Small rodents (e.g., squirrels, hamsters, guinea pigs, gerbils, chipmunks, rats, and mice) and lagomorphs (including rabbits and hares) are rarely infected with rabies and have not been known to transmit rabies to humans (see **Decision Tree D**). In all cases involving rodents, Georgia Poison Center or public health officials should be consulted before a decision is made to initiate PEP.

An unprovoked attack by an animal might be more likely than a provoked attack to indicate that the animal is rabid. Bites inflicted on a person attempting to feed or handle an apparently healthy animal should generally be regarded as provoked.

Refer to the chart on the next page and to the Decision Trees on pages 27-31 for specific guidelines.

### Rabies Post-Exposure Prophylaxis and Animal Guide

Animal Type	Evaluation and Disposition of Animal	Post-Exposure Prophylaxis Recommendations			
Dogs, cats, and ferrets	Healthy and available for 10-day confinement	Persons should not begin PEP unless animal develops clinical signs of rabies.*			
	Rabid or suspected rabid	Immediately begin PEP.			
	Unknown (e.g., escaped)	Consult Georgia Poison Center or public health officials.			
Skunks, raccoons, bobcats, foxes and most other carnivores; bats	Regarded as rabid unless animal proven negative by laboratory tests <sup>†</sup>	Consider immediate PEP.			
Livestock, small rodents, lagomorphs (rabbits and hares), large rodents (woodchucks and beavers), and other mammals	Consider individually.	Consult Georgia Poison Center or public health officials. Bites from squirrels, hamsters, guinea pigs, gerbils, chipmunks, rats, other small rodents, rabbits, and hares almost never require PEP. Larger rodents may be a risk.			

<sup>†</sup>The animal should be euthanized and tested as soon as possible. Holding for observation is not recommended.

tested.

## **Decision Tree A-1 HIGH RISK ANIMALS** Wild Carnivore (Raccoon, Fox, Skunk, etc.) Exposure Did an exposure occur? YES NO Rabies post-exposure prophylaxis (PEP) not Is animal available necessary for testing? NO YES Begin rabies PEP ASAP Are results POSITIVE? NO YES Rabies PEP not Begin rabies PEP ASAP necessary



\*Any direct contact between a person and a bat should be evaluated for an exposure. If the person can be reasonably certain a bite, scratch, or mucous membrane exposure did not occur, or if the bat is available for testing and is negative for presence of rabies virus, post-exposure prophylaxis is not necessary. Other situations that might qualify as exposures include finding a bat in the same room as a person who might be unaware that a bite or direct contact had occurred (e.g., a deeply sleeping person awakens to find a bat in the room or an adult witnesses a bat in the room with a previously unattended child, mentally disabled person, or intoxicated person). These situations should not be considered exposures if rabies is ruled out by diagnostic testing of the bat, or circumstances suggest it is unlikely that an exposure took place. Other household members who did not have direct contact with the bat or were awake and aware when in the same room as the bat should not be considered as having been exposed to rabies.







#### **III. LABORATORY DIAGNOSIS OF RABIES**

#### A. General Principles of Rabies Diagnosis in Animals

The rapid and accurate laboratory diagnosis of rabies infections in animals is essential for timely administration of rabies post-exposure prophylaxis and may also aid in defining current epidemiologic patterns of rabies and in recognizing the need for the development of rabies control programs. In Georgia, animal rabies diagnosis is provided by the three laboratories of the Georgia Public Health Laboratory (GPHL) in accordance with the established national standardized protocol for rabies testing (http://www.cdc.gov/rabies/pdf/RabiesDFASPv2.pdf).

The direct fluorescent antibody test (dFA) is most frequently used to diagnose rabies in animals. All rabies laboratories in the United States perform this test on the brain tissue of animals suspected of having rabies. This test has been thoroughly evaluated for more than 40 years and is recognized as the most rapid and reliable of the tests for routine use. The dFA test is based on the principle that an animal infected by rabies virus will have rabies virus protein (antigen) present in its tissue. Because rabies is present in nervous tissue (and not blood like many other viruses) the ideal tissue to test for the presence of rabies antigen is brain. The most important part of a dFA test is flourescein-labeled anti-rabies antibody. When labeled antibody is added to rabies-suspect brain tissue, it will bind to rabies antigen if it is present. Unbound antibody can be washed away and the areas where the antigen has bound antibody will appear as a bright fluorescent apple green color when viewed with a fluorescence microscope. If rabies virus is absent, there will be no staining.

#### **B.** Specimen Collection, Labeling, and Submission

A key factor in obtaining reliable laboratory results is the condition of the specimen when received by the laboratory. Shipping of specimens should be coordinated with the county health department or animal control officer. Containers for shipment are available from county health departments or from GPHL Laboratory Supply (404-327-7904).

#### • Submission Guidelines

- 1. Only specimens received in good condition with at least two identifiable brain parts are approved for reporting test results.
- 2. For a specimen to be accepted for testing, there must have been exposure of a human or domestic animal to the suspected rabid animal.
- 3. The laboratories are not equipped to handle whole carcasses: only the HEAD is accepted as a specimen, except for bats and animals of similar size, which should be submitted whole. Whole carcasses of any larger animal will be returned to the sender for resubmission of the HEAD ONLY.

- 4. The following guidelines are recommended for the removal of animal heads (whenever possible, this procedure should be performed by a person who has received pre-exposure rabies vaccine).
  - Rubber gloves and protective clothing as well as face and eye protection should be worn while the head is being removed and packaged.
  - Sever the head between the foramen magnum and the atlas. Local veterinarians or trained animal control personnel can assist in this removal.
  - Allow fluids and blood to drain from the head. Keep as clean as possible and place the head in a double plastic bag for transport to the laboratory.
  - If fleas or ticks are present, spray insecticide into the plastic bag containing the head before closing. Do not send maggots.
  - Cutting surfaces and instruments should be thoroughly cleaned with detergent and water and disinfected. Gloves should also be cleaned and disinfected or discarded following use.
- 5. Only brain material (not the entire head) of very large animals (e.g., cows, horses) will be accepted due to limitations for handling in the laboratory. Removal of the brain should only be attempted by a veterinarian. Whole heads of large animals received by the laboratory will be returned to the sender for resubmission of the BRAIN ONLY.
- Rodents (e.g., rats, mice, gerbils, hamsters, guinea pigs, chipmunks, voles, squirrels, moles) and rabbits are not usually involved in the rabies cycle and will not be accepted for testing without prior arrangements with the Epidemiology Program (404-657-2588) or the Georgia Public Health Laboratory to which the specimen is being sent (Atlanta (Decatur): 404-327-7900; Albany: 229-430-4122; Waycross: 912-285-6000.)
- 7. If specimens cannot be delivered to the laboratory immediately, refrigerate but DO NOT FREEZE. Frozen specimens cannot be tested until they thaw, which may cause a delay in reporting.
- 8. Do NOT send tissue in a preservative such as formalin, as rabies testing cannot be performed on such specimens.

- Laboratory Submission Form
  - A Rabies Submission Form #3062 should accompany each specimen submitted for rabies examination. This form should be filled out completely and legibly, making sure to include accurate addresses and phone numbers for use in reporting results. If you do not have a GPHL submitter code, please call GPHL at 404-327-7900 to have one assigned to you prior to submission. Veterinary clinics and hospitals should not submit specimens directly to GPHL without a submitter code. Veterinarians should contact the local health department or animal control agency for assistance in submitting specimens for rabies testing.
  - Blank forms may be found on page 36 of this manual and also on the Department of Public Health website at: <u>http://dph.georgia.gov/lab</u>

#### • Specimen Shipment Guidelines

Containers for shipment are available from county health departments or from GPHL Laboratory Supply (404-327-7904). Rabies testing is available Monday through Friday.

- Properly package the specimen by placing the severed animal head in a double plastic bag and secure the bag by twisting and knotting. For bats or similar size animals, do not remove the head, but submit whole. For large animals (e.g., cows, horses) submit the BRAIN ONLY (consult the attending veterinarian).
- Place the large plastic bag into the Styrofoam container. Add cold packs. DO NOT USE DRY ICE.
- Place the sealed bag containing the specimen on top of the cold packs in the container. Seal the Styrofoam shipper. Place the completed submission form in the brown envelope, and tape to the lid of the sealed shipper. Place the shipper in the cardboard box and tape the address for shipment. Do not seal the box until shipment so that the agent can inspect the container.
- The package should be shipped PREPAID to the nearest Public Health Laboratory using the method of shipment that will assure prompt delivery. CONTAINERS WITH SPECIMENS CANNOT BE SENT THROUGH THE MAIL. Addresses and telephone numbers of laboratories are as follows:

Albany Regional Laboratory 1109 N. Jackson Street Albany, Georgia 31701-2022 Georgia Public Health Laboratory 1749 Clairmont Road Decatur, Georgia 30033-4050 Waycross Regional Laboratory 1751 Gus Karle Parkway Waycross, Georgia 31503

Telephone: 912-388-7050

- Any bite case with a strong probability of human rabies exposure should be handled with utmost speed. Where possible, hand deliver such specimens after telephoning ahead to advise the laboratory of the expected time of arrival.
- Avoid shipping specimens on weekends or holidays unless prior approval has been obtained from the laboratory manager. Special instructions regarding labeling will be needed to ensure that weekend courier or security personnel are notified to receive the specimen from the carrier. A better alternative is to place the specimen in double plastic bags as described above and refrigerate until shipment can be made when the laboratory is in operation Monday through Friday, unless the test result is urgent.

#### C. Reporting and Interpreting Results

Rabies testing is available Monday through Friday. Due to the time required for tissue fixation, reports will ordinarily be issued the next business day following receipt of the specimen, provided that the specimen is received by 10:00 a.m. Reporting will be delayed on specimens that are frozen.

- Specimens received on Friday or those involved in emergency situations (i.e., severe human head or neck exposures or human exposures for which emergency testing has been approved by the Epidemiology Program at 404-657-2588) will be tested and reported the same day received, provided they arrive in the laboratory by 10:00 a.m. Otherwise, results will be reported the following business day.
- If the brain is decomposed or damaged to the point that the laboratory is uncertain as to whether the specimen is the appropriate brain tissue, testing will not be done unless there is human exposure. Report will read "UNSATISFACTORY" with the comment: "Test requires at least two identifiable brain parts." With human exposure, routine testing is performed. If POSITIVE, the report will so state. If NEGATIVE, a report of "UNSATISFACTORY" will be made with the comment: "Test requires at least two identifiable brain parts." In this situation, an unsatisfactory test result should be managed as if POSITIVE.
- All positive, negative, and unsatisfactory rabies results are immediately telephoned or electronically reported to the submitter listed on the Rabies Submission Form, with a hard copy of the report sent by mail. Electronic reporting is available for all submitters. Please contact GPHL (404-327-7900) to initiate electronic reporting.



#### GEORGIA PUBLIC HEALTH RABIES SUBMISSION FORM

Use Only When SendSS is Offline

Complete a separate form for each test requested

Laboratory	use only
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Accession #

Results: 🗖 Positive 🗖 Negative

Unsatisfactory Reason:

ChooseLab to Perform Test

INVESTIGATOR INFORMATION SAMPLE SOURCE										
SubmitterCode			Send SS Offline Ref. ID	CountyofIncident		Incident Date				
								1	1	
Submitter/ClinicName				Victim/Owner Last Name	FirstName	First Name		MI County		
								_		
Street Address				Victim/Owner Phone:	Victim/Owner Phone: Work Phone:		Cell Phone:			
City		State	Zip	Incident Address	Cit		State	Zip		
				COLLECTION AND SH	COLLECTION AND SHIPPING INFORMATION					
Clinic Phone Number Fax Number		Sample Type	Sample Type		Date of Collection					
				Brain / /			1			
Submitter POC Name (require	d to ensure i	notice of	results)	Time		Time o	ne of Collection			
		Other AM		I □ PM						
Submitter POC Phone Number (required information)			Ship	Shipped Condition:						
			Avoid freezing speci room temperatur delivered to PH Lab on	Avoid freezing specimens, and any room temperature item must be delivered to PH Lab on <u>collection date</u>			mmended)			
SELF PAY		(Submit	ter will be bi	lled if a valid code is not provided	0					
(SUBMITTER WILL BE INVOICED) APPROVAL CODE				)DE: _						

SPECIMEN INFORMATION

BITE NUMBER (EPI) BI/A#	Animal Species	Reason for Testing (mandatory, check all that apply)		
County of Animal Origin	Cat Dog (Breed:) Fox Skunk Raccoon Bat	Human Severity Exposure Bite-deep Bite-superficial Domestic Non-Bite Exposure(fluids)		
Date of Death	Other:	Exposure Unknown		
1 1		Epidemiological Reasons		
Classification	Vaccinated Animal?	U Other:		
Pet 🔲 Wild 🔲 Stray	□Yes □No □Unknown			

ADDITIONAL CONTACTS RELATED TO INCIDENT

First Name	Last Name	MI	County	Home Phone	Other Phone	DOB(Vicams)
Attach additional pages for any other contacts related to this specimen						

A correlating list of test and prices is located at http://dph.georgia.gov/lab Page 1 of \_\_\_\_ - Form 3583B (Revised March 2014)

#### D. Serologic Testing

All persons tested during several CDC studies 2-4 weeks after completion of preexposure and post-exposure rabies prophylaxis in accordance with ACIP guidelines have demonstrated an adequate antibody response to rabies. Therefore, serum samples from patients completing pre-exposure or post-exposure prophylaxis do not need to be tested to document seroconversion unless the person is immunosuppressed. If titers are obtained, specimens collected 2-4 weeks after completing the pre-exposure or post-exposure prophylaxis regimen should completely neutralize challenge virus at a 1:5 serum dilution by the Rapid Fluorescent Focus Inhibition Test (RFFIT). Although antibody levels do not define a person's immune status, they are markers of continuing immune response.

In animals, neutralizing antibody titers have been shown to be imperfect markers of protection. Antibody titers will vary with time since the last vaccination. Titers do not directly correlate with protection because other immunologic factors also play a role in preventing rabies and our abilities to measure and interpret those other factors are not well developed. Therefore, evidence of circulating rabies virus antibodies should not be used as a substitute for current vaccination in managing rabies exposures or determining the need for booster vaccinations in animals.

Although virus neutralizing antibody levels may not definitively determine a person's susceptibility or protection from a rabies virus exposure, titers in persons at risk for exposure are used to monitor the relative rabies immune status over time. Considering these issues, serologic testing to quantitate antibody levels after rabies vaccination in humans and animals is applicable in the following cases:

- A person at "continuous risk" of exposure to rabies should have a serum sample tested for rabies antibody every six months (see page 21). This includes rabies research laboratory workers and rabies biologics production workers.
- A person at "frequent risk" of exposure to rabies should have a serum sample tested for rabies antibody every two years (see page 21). This includes: rabies diagnostic laboratory workers; cavers; veterinarians and staff; animal control and wildlife workers in areas where rabies is enzootic; and persons who frequently handle bats.
- Some "rabies-free" jurisdictions may require evidence of vaccination and rabies antibodies in domestic animals (dogs and cats) for importation purposes. CONTACT INDIVIDUAL COUNTRIES FOR IMPORT REQUIREMENTS. Keep in mind there is not an established "protective" titer in animals. Titers do not directly correlate with protection because other immunologic factors also play a role in preventing rabies and our abilities to measure and interpret those other factors are not well developed. Therefore, evidence of circulating rabies virus antibodies should not be used as a substitute for

current vaccination in managing rabies exposures or determining the need for booster vaccinations in animals.

There are two types of RFFIT tests depending on the request: a **screen** test simply tells the patient/client if a booster of rabies vaccine is indicated and serum is tested at two dilutions. An **end-point** titer is used to determine the exact titer and is tested at serial five-fold dilutions until an end-point is reached. This test is indicated for those who want to know their exact titer and for animals being exported to some rabies-free countries. Testing requires two milliliters (mls) of serum.

#### • Laboratories conducting rabies serologic testing

<u>Note</u>: Phoning the laboratory in advance for correct forms, testing costs, and proper instructions is recommended.

☆	Kansas State University Rabies Laboratory 2005 Research Park Circle Manhattan KS 66502 Phone: 785-532-4483 Fax: 785-532-4474 www.vet.ksu.edu/depts/dmp/service/rabies/index.htm
*	Atlanta Health Associates, Inc. 309 Pirkle Ferry Road, Suite D300 Cumming, GA 30040 Phone: 800-717-5612 Fax: 770-205-9021 http://atlantahealth.net
☆	Auburn University Virology Laboratory (animals only) College of Veterinary Medicine Department of Pathobiology Virology Laboratory 261 Greene Hall Auburn University, AL 36849 Phone: 334-844-2659 Fax: 334-844-2652 www.vetmed.auburn.edu/virology

#### IV. RABIES CONTROL DURING DISASTER RESPONSE

Animals may be displaced during and after manmade or natural disasters and require emergency sheltering. Animal rabies vaccination and exposure histories are often not available for displaced animals and disaster response creates situations where animal caretakers may lack appropriate training and previous vaccination. For these situations it is critical to implement and coordinate rabies prevention and control measures to reduce the risk of rabies transmission and the need for human PEP. Public health officials and other response partners should consider the following control measures, when feasible:

- Examine each animal at a triage site for signs of rabies.
- Isolate animals exhibiting signs of rabies pending evaluation by a veterinarian.
- Ensure that all animals have a unique identifier.
- Administer a rabies vaccination to all dogs, cats and ferrets unless reliable proof of vaccination exists.
- Adopt minimum standards for animal caretakers that include personal protective equipment, previous rabies vaccination, and appropriate training in animal handling.
- Maintain documentation of animal disposition and location (e.g., returned to owner, died or euthanized, adopted, relocated to another shelter, address of new location).
- Provide facilities to confine and observe animals involved in exposures.
- Report human exposures to appropriate public health authorities.

#### V. BATS AND RABIES

The most common rabies virus variants responsible for human rabies in the United States are bat-related; therefore, any potential exposure to a bat requires a thorough evaluation. During 1990-2007, a total of 34 naturally acquired bat-associated human cases of rabies were reported in the United States. In 6 cases, a bite was reported; in 2 cases, contact with a bat and a probable bite were reported. In 15 cases, physical contact was reported (e.g., the removal of a bat from the home or workplace or the presence of a bat in the room where the person had been sleeping) but no bite was documented. In 11 cases, no bat encounter was reported; in these cases, an unreported or undetected bat bite remains the most plausible hypothesis because the genetic sequences of the human rabies viruses closely matched those of specific species of bats. Clustering of human cases associated with bat exposures has never been reported in the United States (e.g., within the same household or among a group of campers where bats were observed during their activities). The risk for rabies resulting from an encounter with a bat may be difficult to determine because of the limited injury inflicted by a bat bite (compared with more obvious wounds caused by the bite of terrestrial carnivores), an inaccurate recall of a bat encounter that may have occurred several weeks or months earlier, and evidence that some bat-related rabies viruses may be more likely to result in infection after inoculation into superficial epidermal layers. For these reasons, any direct contact between a human and a bat should be evaluated for an exposure.

Awareness of the facts about bats and rabies can help people protect themselves, their families, and their pets.

- Bat Rabies Prevention Tips
  - It is not possible to tell if a bat has rabies by looking at it. Rabies can be confirmed only in a laboratory. However, any bat that is active by day, is found in a place where bats are not usually seen (for example, in a room in the house or on the lawn), or is unable to fly is far more likely than others to be rabid. Such bats are often the most easily approached. Therefore, it is best never to handle any bat.
  - Bat bites are not always visible. Therefore, in situations in which a bat is physically present and there is a possibility of exposure, the person should seek medical advice and the bat should be safely captured (see next page) and submitted to a rabies laboratory for testing. If rabies cannot be ruled out by laboratory testing, or if the bat is not available for testing, people with a reasonable probability of an exposure may be recommended for rabies post-exposure prophylaxis. Scenarios that may indicate a reasonable probability of exposure to rabies include:

- a child picks up a live bat
- an adult touches a bat without seeing the part of the body they touched
- a bat flies into a person and touches bare skin
- a person steps on a bat with bare feet
- a deeply sleeping person awakens to find a bat in the room
- a bat is found near an infant, toddler, or mentally impaired or intoxicated person.

Assistance with bat capture may be provided by a local animal control agency or health department. If professional help is immediately unavailable, the bat may be safely captured by following these steps:

#### Safe Bat Capture

- Equipment needed: leather work gloves; small box or coffee can; piece of cardboard; tape.
- When the bat lands, approach it slowly while wearing the gloves and place the box or coffee can over it. Slide the cardboard under the container to trap the bat inside.
- Tape the cardboard to the container securely and punch very small holes (1/8 inch or less in diameter) in the cardboard, allowing the bat to breathe.
- If any possible contact between the bat and a person or domestic animal has occurred, do not release the bat. Contact the health department or animal control agency to make arrangements for rabies testing.
- If no human or pet exposure has occurred, take the container outdoors immediately and release the bat away from people and pets.
- Some bats live in buildings, and there may be no reason to evict them if there is little chance for contact with people. However, bats should always be prevented from entering living quarters or occupied spaces in homes, churches, schools, and other similar areas where they might contact people and pets. Assistance with "bat-proofing" homes can be provided by an animal control or wildlife conservation agency. Another excellent resource is Bat Conservation International at <u>www.batcon.org</u>.

• If there is suspicion that a pet or domestic animal has been bitten by a bat, contact a veterinarian or health department for assistance immediately and have the bat tested for rabies. Remember to keep vaccinations current for cats, dogs, ferrets, and other animals.

Citation is given to the Centers for Disease Control for information contained in the brochure, "Bats and Rabies: A Public Health Guide"

#### VI. FREQUENTLY-ASKED QUESTIONS (FAQ) ABOUT RABIES

#### What is the incubation period of rabies in animals and humans?

The incubation period is the time between exposure and onset of clinical signs of disease. The incubation period may vary from a few days to several years, but typically lasts 1 to 3 months. This period is quite long because the rabies virus spreads slowly through the nerves to the spinal cord and brain. There are no signs of illness during the incubation period; rabies virus is not transmissible during this time. When the virus reaches the brain, it multiplies rapidly and passes to the salivary glands. At this point, clinical signs of rabies are evident and rabies virus can be transmitted via saliva.

#### How can I protect my pet from rabies?

First, visit your veterinarian with your pet on a regular basis and keep rabies vaccinations up-to-date for all dogs, cats, and ferrets. Second, maintain control of your pets by keeping cats and ferrets indoors and keeping dogs under direct supervision. Third, spay or neuter your pets to help reduce the number of unwanted pets that may not be properly cared for or vaccinated regularly. Lastly, call animal control to remove all stray animals from your neighborhood since these animals may be unvaccinated or ill.

#### Why does my pet need the rabies vaccine?

Although the majority of rabies cases occur in wildlife, most humans are given rabies vaccine as a result of exposure to domestic animals. This explains the tremendous cost of rabies prevention in domestic animals in the United States. While wildlife are more likely to be rabid than are domestic animals in the United States, the amount of human contact with domestic animals greatly exceeds the amount of contact with wildlife. Your pets and other domestic animals can be infected when they are bitten by rabid wild animals. When "spillover" rabies occurs in domestic animals, the risk to humans is increased. Pets are therefore vaccinated by your veterinarian to prevent them from acquiring the disease from wildlife and thereby transmitting it to humans.

### My dog just fought with a raccoon and I picked him up to see whether he had any wounds. Am I at risk for rabies?

This would be considered of minimal risk but the first line of defense is to always wash hands with soap and water. Nonbite exposures (other than organ or tissue transplants) have rarely been proven to cause rabies and post-exposure prophylaxis is not indicated unless saliva or other potentially infectious material was directly introduced into fresh, open cuts in the skin or onto mucous membranes. Rabies virus is inactivated by desiccation, ultraviolet irradiation, and other factors and does not persist in the environment (e.g., on a dog's fur).

#### Can a vaccinated animal ever get rabies?

Rabies is rare in vaccinated animals. If such an event is suspected, it should be reported immediately to District public health officials and the Epidemiology Program. The laboratory diagnosis should be confirmed and the virus characterized by a rabies reference laboratory. A thorough epidemiologic investigation should be conducted.

### Can I use rabies titers as a substitute for current vaccination or in the management of domestic animals exposed to rabies?

No, rabies titers alone are only one marker of immunity and may not indicate absolute protection. Titers do not directly correlate with protection because other immunologic factors also play a role in preventing rabies and our abilities to measure and interpret those other factors are not well developed.

#### Will the rabies vaccine make me sick?

Adverse reactions to rabies vaccine and immune globulin are not common. Newer vaccines in use today cause fewer adverse reactions than previously available vaccines. Mild, local reactions to the rabies vaccine, such as pain, redness, swelling, or itching at the injection site, have been reported. Rarely, symptoms such as headache, nausea, abdominal pain, muscle aches, and dizziness have been reported. Local pain and low-grade fever may follow injection of rabies immune globulin.

#### What if I cannot get rabies vaccine on the day I am supposed to get my next dose?

Consult with your doctor or state or local public health officials for recommended times if there is going to be a change in the recommended schedule of shots. Rabies prevention is a serious matter and changes should not be made in the schedule of doses.

#### Should I be concerned about rabies when I travel outside the United States?

Yes. Rabies and rabies-like viruses occur in animals anywhere in the world. When traveling, it is always prudent to avoid approaching any wild or domestic animal.

The developing countries in Africa, Asia, and Latin America have additional problems in that dog rabies is common there and human PEP may be difficult to obtain. The importance of rabid dogs in these countries, where tens of thousands of people die of the disease each year, cannot be overstated. Unlike programs in developed countries, dog rabies vaccination programs in developing countries have not always been successful. Before traveling abroad, consult a health care provider, travel clinic, or health department about your risk of exposure to rabies and how to handle an exposure should it arise. Medical assistance should be obtained as soon as possible after an exposure.

#### Can rabies be transmitted from one person to another?

The only documented cases of rabies caused by human-to-human transmission, although extremely rare, occurred among recipients of transplanted corneas and other solid organs. Organ and tissue transplantation resulting in rabies transmission has occurred among 16 transplant recipients from corneas (n=8), solid organs (n=7), and vascular tissue (n=1). The 16 cases occurred in six countries: the United States (5 cases: one cornea, three solid organs, and one vascular tissue), Germany (4 cases), Thailand (2 cases), India (2 cases), Iran (2 cases), and France (1 case). Investigations revealed that the donors had died of an illness compatible with or proven to be rabies. Stringent guidelines for acceptance of donor corneas have reduced this risk. No documented laboratory-diagnosed cases of human-to-human transmission have been documented from a bite or nonbite exposure other than the transplant cases. Casual contact, such as touching a person with rabies or contact with non-infectious fluid or tissue (i.e., urine, blood, and feces) does not constitute an exposure and does not require PEP. In addition, contact with someone who is receiving rabies PEP does not constitute rabies exposure and does not require post-exposure prophylaxis.

Citation is given to the Centers for Disease Control for information contained in their rabies website: http://www.cdc.gov/rabies.

#### VII. REFERENCES

#### A. Definitions

- **Currently Vaccinated Against Rabies**. An animal is "currently vaccinated" and is considered immunized against rabies if a vaccination certificate documents that the animal received a USDA-approved primary rabies vaccine from a licensed veterinarian at least 28 days previously and that booster vaccinations have been administered on an annual or triennial schedule, in accordance with the *Compendium of Animal Rabies Prevention and Control* (see pages 51-64) or as described on the individual vaccine label.
- **Exposure.** Rabies exposure occurs when the virus is introduced into bite wounds or open cuts in skin or onto mucous membranes. Two categories of exposure, bite and nonbite, should be considered.
  - **Bite**. Any penetration of the skin by teeth constitutes a bite exposure. All bites, regardless of location, represent a potential risk of rabies transmission. Keep in mind that bites by some animals, such as bats, can inflict minor injury and thus be undetected.
  - Nonbite. The contamination of open wounds, abrasions, mucous membranes, or theoretically, scratches, with saliva or other potentially infectious material (such as neural tissue) from a rabid animal constitutes a nonbite exposure. Nonbite exposures from terrestrial animals rarely cause rabies. However, occasional reports of transmission by nonbite exposure suggest that such exposures constitute sufficient reason to consider post-exposure prophylaxis.
- Non-Exposure. Other contact by itself, such as being in the vicinity of, petting or handling an animal, or coming in contact with blood, urine, or feces does NOT constitute an exposure and does NOT require PEP. Because desiccation and ultraviolet irradiation inactivate the rabies virus, in general, if the material containing the virus is dry, the virus can be considered noninfectious.
- **Confinement.** A general term referring to the restriction of an animal to a building, pen, or other escape-proof enclosure to monitor for clinical signs of rabies. There are **two specific types of confinement**, depending upon the circumstances of the encounter.
  - Quarantine (for animal-human encounters). This is a **10-day** period of confinement for a domestic animal (dog, cat, or ferret **only**) which has bitten a person, **no matter if the animal is currently vaccinated or not**. Quarantine conditions should prevent direct contact with other animals or persons. The quarantine shall be conducted under the authority of the designated local rabies control agency in which the place, manner, and provisions of the quarantine are specified. For example, quarantine may

take place in a kennel in a veterinary hospital, animal control facility, commercial boarding establishment, or a pen at home, depending on local requirements. At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies and the exposed person notified.

• Strict Isolation (for animal-animal encounters). This is the confinement of an animal exposed or potentially exposed to rabies in a manner that prevents direct contact with other animals or persons. In most cases, this term applies to an **unvaccinated** domestic animal exposed to a rabid wild animal; the duration of strict isolation should be **six months**. Strict isolation should be conducted under the authority of the designated local rabies control agency in which the place, manner, and provisions of the confinement are specified. For example, strict isolation may take place in an animal control facility, or an isolation pen at home, depending on local requirements. At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies and the exposed person notified.

<u>Note</u>: The animal should be vaccinated against rabies upon entry into isolation OR one month prior to isolation exit.

- Observation period. In animal-animal encounters involving currently vaccinated domestic animals (dogs, cats, ferrets, and in some cases, livestock) exposed to a rabid wild animal, the observation period is the 45-day period in which the animal is kept under the owner's control to monitor for clinical signs of rabies to develop. During the observation period, the animal should not be permitted to roam and should be restricted to leash walks, if applicable. At the first sign of illness or behavioral change in the animal, the local rabies control agency should be notified and the animal should be evaluated by a veterinarian. If clinical signs are suggestive of rabies, the animal should be immediately euthanized and tested for rabies and the exposed person notified.
- **Provoked Attack**. An attack is considered to be "provoked" if a domestic animal is placed in a situation such that an expected reaction would be to bite or attack. Examples include invasion of an animal's territory, attempting to pet or handle an unfamiliar animal, startling an animal, breaking up an animal fight, running or bicycling past an animal, assisting an injured or sick animal, trying to capture an animal, or removing food, water, or other objects in the animal's possession.
- **Unprovoked Attack**. An attack or bite is considered to be "unprovoked" when none of the above conditions for a "provoked" attack are met; essentially, the animal strikes for no apparent reason.

#### B. Georgia Rabies Control Law

- I. OPINIONS OF THE ATTORNEY GENERAL
  - Control of rabies generally is delegated to county boards of health, and control of dangerous drugs is vested with the State Board of pharmacy and state drug inspector (now director of Georgia Drugs and Narcotics Agency). 1975 Op. Atty. Gen. No. 75-23.
  - Expense of confining animals included in county board's budget—Local County Boards of Health should prescribe rules for prevention and control of rabies by providing for vaccination, tagging, and certification of dogs, and for confinement of any animal which exhibits any signs of rabies; cost of such confinement would be an expense of County Board of Health to be included in its budget which is submitted to local taxing authorities under provision of section 31-3-14, 1965-66 Op. Atty. Gen. No. 65-21.
  - Responsibility of County Boards of Health regarding strays and unwanted dogs—Local County Boards of Health should adopt rules and regulations relative to catching and impounding of strays and unwanted dogs. 1965-66 Op. Atty. Gen. No. 65-21.

#### II. OFFICIAL CODE 31-19, CONTROL OF RABIES

#### 31-19-1. Responsibility for Control

Each county board of health shall have primary responsibility for the control of rabies within its jurisdiction. Such boards, in addition to their other powers, are empowered and required to adopt and promulgate rules and regulations for the prevention and control of such disease.

#### 31-19-2. Powers of department in infected area.

The department (DPH) may declare any County or any area therein or any group of counties or areas therein where rabies exists to be an infected area and may provide for immunization and such other measures as shall be indicated for the prevention and control of the disease.

#### 31-19-3. Licensing and regulation of animals by local authorities.

The governing authorities of each county and municipality are authorized and required, in the control of rabies, to require regulation or licensing of animals.

#### 31-19-4. Duty of notification.

It shall be the duty of any person bitten by any animal reasonably suspected of being rabid immediately to notify the appropriate county board of health. It shall be the duty of the owner, custodian, or person having possession and knowledge of any animal which has bitten any person or animal or of any animal which exhibits any signs of rabies to notify the appropriate county board of health and to confine such animal in accordance with rules and regulations of the county board of health.

#### 31-19-5. Inoculation of canines and felines against rabies.

The county boards of health are empowered and required to adopt and promulgate rules and regulations requiring canines and felines to be inoculated against rabies and to prescribe the intervals and means of inoculation, the fees to be paid in county sponsored clinics, that procedures be in compliance with the recommendations of the National Association of State Public Health Veterinarians for identifying inoculated canines and felines, and all other procedures applicable thereto. As used in this chapter, the term "inoculation against rabies" means the administering by a licensed veterinarian of antirabies vaccine approved by the department.

#### 31-19-6. Certificates of inoculation: tags.

Reserved. Repealed by Ga. L. 1992, p. 2089, sec. 2, effective July 1, 1992.

#### 31-19-7. County rabies control officer.

(a) The County board of health shall appoint a person who is knowledgeable of animals to be the County rabies control officer. It shall be the duty of the County rabies control officer to enforce this chapter and other laws which regulate the activities of dogs.

(b) The County governing authority of each County is authorized to levy a fee not to exceed 50 cents for each dog, such fee to be collected by the veterinarian administering the antirabies vaccine required by this chapter. This fee shall be in addition to that provided for in Code Section 31-19-5. If any County has no resident veterinarian, the out-of-county veterinarian administering the antirabies vaccine and collecting the fee provided for by this Code section shall forward to the treasurer of the County of the dog owner's residence the fee prescribed by that County's governing authority.

(c) The fees collected under this Code section shall be used to help in paying the salary of the County rabies control officer.

#### 31-19-8. Joint administration of chapter by adjoining counties.

The governing authority of each County may devise and implement plans whereby this chapter, as amended, is administered jointly with one or more adjoining counties.

#### 31-19-9. Applicability to municipalities with rabies control laws.

This chapter shall not apply to municipalities which already have a rabies control law unless and until such law is repealed.

#### 31-19-10. Penalty.

Any person who violates any provision of this chapter or any rule or regulation adopted pursuant thereto shall be guilty of a misdemeanor.

#### C. Compendium of Animal Rabies Prevention and Control, 2011\*; National Association of State Public Health Veterinarians, Inc. (NASPHV)

Rabies is a fatal viral zoonosis and a serious public health problem (1). All mammals are believed to be susceptible to the disease, and for purposes of this document, use of the term "animal" refers to mammals. The disease is an acute, progressive encephalitis caused by a lyssavirus. Rabies virus is the most important lyssavirus globally. In the United States, multiple rabies virus variants are maintained in wild mammalian reservoir populations such as raccoons, skunks, foxes, and bats. Although the U.S. has been declared free of canine rabies virus variant transmission, there is always a risk of reintroduction of these variants (2-6).

The virus is usually transmitted from animal to animal through bites. The incubation period is highly variable. In domestic animals it is generally 3-12 weeks, but can range from several days to months, rarely exceeding 6 months (7). Rabies is communicable during the period of salivary shedding of rabies virus. Experimental and historic evidence document that dogs, cats, and ferrets shed virus a few days prior to clinical onset and during illness. Clinical signs of rabies are variable and include inappetance, dysphagia, cranial nerve deficits, abnormal behavior, ataxia, paralysis, altered vocalization, and seizures. Progression to death is rapid. There are currently no known effective rabies antiviral drugs.

The recommendations in this compendium serve as a basis for animal rabies prevention and control programs throughout the United States and facilitate standardization of procedures among jurisdictions, thereby contributing to an effective national rabies control program. This document is reviewed and revised as necessary. The most current version replaces all previous versions. These recommendations do not supersede state and local laws or requirements. Principles of rabies prevention and control are detailed in Part I; recommendations for parenteral vaccination procedures are presented in Part II; and all animal rabies vaccines licensed by the United States Department of Agriculture (USDA) and marketed in the United States are listed and described in Part III.

#### The NASPHV Committee

Catherine M. Brown, DVM, MSc, MPH, Chair Lisa Conti, DVM, MPH Paul Ettestad, DVM, MS Mira J. Leslie, DVM, MPH Faye E. Sorhage, VMD, MPH Ben Sun, DVM, MPVM

#### **Consultants to the Committee**

Donald Hoenig, VMD; AVMA Donna M. Gatewood, DVM, MS; USDA Center for Veterinary Biologics Lorraine Moule; NACA Barbara Nay; Animal Health Institute Raoult Ratard, MD, MS, MPH; CSTE Charles E. Rupprecht, VMD, MS, PhD; CDC Dennis Slate, MS, PhD; USDA Wildlife Services James Powell, MS; APHL Burton Wilcke, Jr., PhD; APHA

#### **Endorsed by:**

American Public Health Association (APHA) American Veterinary Medical Association (AVMA) Association of Public Health Laboratories (APHL) Council of State and Territorial Epidemiologists (CSTE) National Animal Control Association (NACA)

#### \*Address all correspondence to:

Catherine M. Brown, DVM, MSc, MPH State Public Health Veterinarian Massachusetts Department of Public Health Hinton State Laboratory Institute 305 South St. Jamaica Plain, MA 02130

#### Part I. Rabies Prevention and Control

#### A. PRINCIPLES OF RABIES PREVENTION AND CONTROL

**1. CASE DEFINITION:** An animal is determined to be rabid after diagnosis by a qualified laboratory as specified in Part I.A.9. The national case definition for animal rabies requires laboratory confirmation by either:

• A positive direct fluorescent antibody test (preferably performed on central nervous system tissue); or

• Isolation of rabies virus (in cell culture or in a laboratory animal (8).

**2. RABIES EXPOSURE:** Rabies is transmitted when the virus is introduced into bite wounds, open cuts in skin, or onto mucous membranes from saliva or other potentially infectious material such as neural tissue (9). Questions regarding possible exposures should be directed promptly to state or local public health authorities.

**3. PUBLIC HEALTH EDUCATION:** Essential components of rabies prevention and control include ongoing public education, responsible pet ownership, routine veterinary care and vaccination, and professional continuing education. The majority of animal and human exposures to rabies can be prevented by raising awareness concerning: rabies transmission routes, avoiding contact with wildlife, and following appropriate veterinary care. Prompt recognition and reporting of possible exposures to medical professionals and local public health authorities is critical.

**4. HUMAN RABIES PREVENTION:** Rabies in humans can be prevented either by eliminating exposures to rabid animals or by providing exposed persons with prompt local treatment of wounds combined with the appropriate administration of human rabies immune globulin and vaccine. Exposure assessment should occur before postexposure rabies prophylaxis (PEP) is initiated and should include discussion between medical providers and public health officials. The rationale for recommending preexposure prophylaxis and details of both pre- and post-exposure prophylaxis administration can be found in the current recommendations of the Advisory Committee on Immunization Practices (ACIP) (9,10). These recommendations, along with information concerning the current local and regional epidemiology of animal rabies and the availability of human rabies biologics, are available from state health departments.

**5. DOMESTIC ANIMAL VACCINATION:** Multiple vaccines are licensed for use in domestic animal species. Vaccines available include: inactivated or modified live virus vectored products; products for intramuscular and subcutaneous administration; products with durations of immunity from one to 4 years; and products with varying minimum age of vaccination. The recommended vaccination procedures and the licensed animal vaccines are specified in Parts II and III of this compendium, respectively. Local

governments should initiate and maintain effective programs to ensure vaccination of all dogs, cats, and ferrets and to remove strays and unwanted animals. Such procedures in the United States have reduced laboratory confirmed cases of rabies in dogs from 6,949 in 1947 to 93 in 2009 (2). Because more rabies cases are reported annually involving cats (274 in 2009) than dogs, vaccination of cats should be required (2). Animal shelters and animal control authorities should establish policies to ensure that adopted animals are vaccinated against rabies.

**6. RABIES IN VACCINATED ANIMALS:** Rabies is rare in vaccinated animals (*11-13*). If such an event is suspected, it should be reported to public health officials; the vaccine manufacturer; and USDA, Animal and Plant Health Inspection Service, Center for Veterinary Biologics (Internet: http://www.aphis.usda.gov/animal\_health/vet\_biologics/vb\_adverse\_event.shtml; telephone: 800-752-6255). The laboratory diagnosis should be confirmed and the virus variant characterized by the Centers for Disease Control and Prevention (CDC) rabies reference laboratory. A thorough epidemiologic investigation 5 including documentation of the animal's vaccination history and a description of potential rabies exposures should be conducted.

**7. RABIES IN WILDLIFE:** The control of rabies among wildlife reservoirs is difficult (*14*). Vaccination of free-ranging wildlife or selective population reduction is useful in some situations (*15*), but the success of such procedures depends on the circumstances surrounding each rabies outbreak (see Part I. C.). Because of the risk of rabies in wild animals (especially raccoons, skunks, coyotes, foxes, and bats), the American Veterinary Medical Association, American Public Health Association, Council of State and Territorial Epidemiologists, National Animal Control Association and the National Association of State Public Health Veterinarians strongly recommend the enactment and enforcement of state laws prohibiting their importation, distribution, translocation, and private ownership.

**8. RABIES SURVEILLANCE**: Enhanced laboratory-based rabies surveillance and variant typing are essential components of rabies prevention and control programs. Accurate and timely information and reporting is necessary to: guide human PEP decisions; determine the management of potentially exposed animals; aid in emerging pathogen discovery; describe the epidemiology of the disease; and assess the need for and effectiveness of vaccination programs for domestic animals and wildlife. Every animal submitted for rabies testing should be reported to CDC to evaluate surveillance trends. Electronic laboratory reporting and notification of animal rabies surveillance data should be implemented (*16*). Optimal information on animals submitted for rabies testing should include species, point location, vaccination history, rabies virus variant (if rabid), and human or domestic animal exposures. Rabid animals with a history of importation within 60 days into the United States are immediately notifiable by state health departments to CDC; all indigenous cases should follow standard notification protocols (*17*). Integration with standard public health reporting and notification systems should facilitate the transmission of the above data elements.

#### 9. RABIES DIAGNOSIS:

a) The direct fluorescent antibody (DFA) test is the gold standard for rabies diagnosis. The DFA test should be performed in accordance with the established national standardized protocol (http://www.cdc.gov/rabies/docs/standard\_dfa\_protocol\_rabies.pdf) by a qualified laboratory that has been designated by the local or state health department (18, 19). Animals submitted for rabies testing should be euthanized (20, 21) in such a way as to maintain the integrity of the brain so that the laboratory can recognize the anatomical parts. Except in the case of very small animals, such as bats, only the head or brain (including brain stem) should be submitted to the laboratory. To facilitate prompt laboratory testing, submitted specimens should be stored and shipped under refrigeration without delay. The need to thaw frozen specimens will delay testing. Chemical fixation of tissues should be avoided to prevent significant testing delays and because it might preclude reliable testing. Questions about testing of fixed tissues should be directed to the local rabies laboratory or public health department.

b) Rabies testing should be available on an emergency basis to expedite exposure management decisions (18). When confirmatory testing is needed by state health departments (e.g., inconclusive results, unusual species, mass exposures), the CDC rabies laboratory can provide results within 24 hours of submission (22).

c) A direct rapid immunohistochemical test (DRIT) is being used by trained field personnel in

surveillance programs for specimens not involved in human or domestic animal exposures (23-26). All positive DRIT results need to be confirmed by DFA testing at a qualified laboratory.

d) Currently, there are no USDA licensed rapid test kits commercially available for rabies diagnosis. Unlicensed tests should not be used due to several concerns: the sensitivity/specificity are not known; the tests have not been validated against current standard methods; the excretion of virus in the saliva is intermittent and the amount varies over time; any test result would need to be confirmed by more 6 reliable methods such as DFA testing on brain tissue; and the interpretation of results may place exposed animals and persons at risk.

**10. RABIES SEROLOGY:** Some jurisdictions require evidence of vaccination and rabies virus antibodies for animal importation purposes. Rabies virus antibody titers are indicative of a response to vaccine or infection. Titers do not directly correlate with protection because other immunologic factors also play a role in preventing rabies, and our abilities to measure and interpret those other factors are not well-developed. Therefore, evidence of circulating rabies virus antibodies in animals should not be used as a substitute for current vaccination in managing rabies exposures or determining the need for booster vaccinations (27-30).

**11. RABIES RESEARCH:** Information derived from well-designed studies is essential for the development of science-based recommendations. Data are needed in several areas including: viral shedding periods for domestic livestock and lagomorphs; potential shedding of virus in milk; earliest age at which rabies vaccination is effective and protective effect of maternal antibody; duration of immunity; postexposure prophylaxis protocols for domestic animals; models for treatment of clinical rabies; extra label vaccine use in domestic animals and wildlife rabies reservoirs; host-pathogen adaptations and dynamics; and the ecology of wildlife rabies reservoir species, especially in relation to the use of oral rabies vaccines.

#### **B. PREVENTION AND CONTROL METHODS IN DOMESTIC AND CONFINED ANIMALS**

**1. PREEXPOSURE VACCINATION AND MANAGEMENT:** Parenteral animal rabies vaccines should be administered only by or under the direct supervision of a licensed veterinarian on premises. Rabies vaccinations may also be administered under the supervision of a licensed veterinarian to animals held in animal control shelters before release. The veterinarian signing a rabies vaccination certificate must ensure that the person administering vaccine is identified on the certificate and is appropriately trained in vaccine storage, handling, administration, and in the management of adverse events. This practice assures that a qualified and responsible person can be held accountable for properly vaccinating the animal. Within 28 days after initial vaccination, a peak rabies virus antibody titer is reached, and the animal can be considered immunized (*29,31-33*). An animal is currently vaccinated and is considered immunized if the initial vaccination was administered at least 28 days previously or booster vaccinations have been administered in accordance with this compendium.

Regardless of the age of the animal at initial vaccination, a booster vaccination should be administered 1 year later (see Parts II and III for vaccines and procedures). No laboratory or epidemiologic data exist to support the annual or biennial administration of 3- or 4-year vaccines after the initial series. Because a rapid anamnestic response is expected, an animal is considered currently vaccinated immediately after a booster vaccination (*34*).

#### a) DOGS, CATS AND FERRETS

All dogs, cats, and ferrets should be vaccinated against rabies and revaccinated in accordance with Part III of this compendium. If a previously vaccinated animal is overdue for a booster, it should be revaccinated. Immediately after the booster, the animal is considered currently vaccinated and should be placed on a booster schedule, depending on the labeled duration of the vaccine used.

#### b) LIVESTOCK

All horses should be vaccinated against rabies (35). Livestock, including species for which licensed vaccines are not available, that have frequent contact with humans (e.g., in petting zoos, fairs, and other public exhibitions) should be vaccinated against rabies (36,37). Consideration should also be given to vaccinating livestock that are particularly valuable.

c) CAPTIVE WILD ANIMALS AND HYBRIDS (the offspring of wild animals crossbred to

domestic animals).

(1) Wild animals or hybrids should not be kept as pets (38-40). No parenteral rabies vaccines are licensed for use in wild animals or hybrids (41).

(2) Animals that are maintained in exhibits and in zoological parks and are not completely excluded from all contact with rabies vectors can become infected. Moreover, wild animals might be incubating rabies when initially captured; therefore, wild-caught animals susceptible to rabies should be quarantined for a minimum of 6 months. Employees who work with animals at such facilities should receive preexposure rabies vaccination. The use of pre- or postexposure rabies vaccinations for handlers who work with animals at such facilities might reduce the need for euthanasia of captive animals that expose handlers. Carnivores and bats should be housed in a manner that precludes direct contact with the public (*36,37*).

**2. STRAY ANIMALS:** Stray dogs, cats, and ferrets should be removed from the community. Local health departments and animal control officials can enforce the removal of strays more effectively if owned animals are required to have identification and are confined or kept on leash. Strays should be impounded for at least 3 business days to determine if human exposure has occurred and to give owners sufficient time to reclaim animals.

#### 3. IMPORTATION AND INTERSTATE MOVEMENT OF ANIMALS:

a) INTERNATIONAL. CDC regulates the importation of dogs and cats into the United States (5). Importers of dogs must comply with rabies vaccination requirements (42 CFR, Part 71.51[c] [http://www.cdc.gov/animalimportation/dogs.html]) and complete CDC form 75.37 (http://www.cdc.gov/animalimportation/pdf/dog-import.pdf). These regulations require dogs imported from rabies endemic countries to be vaccinated for rabies and confined for varying timeframes depending on age, prior vaccination status, and country of origin. The appropriate health official of the state of destination should be notified within 72 hours of the arrival of any imported dog required to be placed in confinement under these regulations. Failure of the owner to comply with these confinement requirements should be promptly reported to the Division of Global Migration and Quarantine, CDC (telephone: 404-639-4528 or 404-639-4537).

Federal regulations alone are insufficient to prevent the introduction of rabid animals into the United States (3,4,42,43). All imported dogs and cats are subject to state and local laws governing rabies and should be currently vaccinated against rabies in accordance with this compendium. Failure of the owner to comply with state or local requirements should be referred to the appropriate state or local official.

b) AREAS WITH DOG-TO-DOG RABIES TRANSMISSION. Canine rabies virus variants have been eliminated in the United States (2,6). Rabid dogs have been introduced into the continental United States from areas with dog-to-dog rabies transmission (3,4,42,43). The movement of dogs for the purposes of adoption or sale from areas with dog-dog rabies transmission increases the risk of introducing canine-transmitted rabies to areas where it does not currently exist and should be prohibited.

c) INTERSTATE. Before interstate (including commonwealths and territories) movement, dogs, cats, ferrets, and horses should be currently vaccinated against rabies in accordance with this compendium's recommendations (see Part I. B.1.). Animals in transit should be accompanied by a currently valid NASPHV Form 51, Rabies Vaccination Certificate (http://www.nasphv.org/Documents/RabiesVacCert.pdf). When an interstate health certificate or certificate of veterinary inspection is required, it should contain the same rabies vaccination information as Form 51.

**4. ADJUNCT PROCEDURES:** Methods or procedures that enhance rabies control include the following (http://www.rabiesblueprint.com/spip.php?article119):

a) IDENTIFICATION. Dogs, cats, and ferrets should be identified (e.g., metal or plastic tags or

microchips) to allow for verification of rabies vaccination status.

b) LICENSURE. Registration or licensure of all dogs, cats, and ferrets is an integral component of an effective rabies control program. A fee is frequently charged for such licensure, and revenues collected are used to maintain rabies or animal control activities. Evidence of current vaccination should be an essential prerequisite to licensure.

c) CANVASSING. House-to-house canvassing by animal control officials facilitates enforcement of vaccination and licensure requirements.

d) CITATIONS. Citations are legal summonses issued to owners for violations, including the failure to vaccinate or license their animals. The authority for officers to issue citations should be an integral part of each animal control program.

e) ANIMAL CONTROL. All local jurisdictions should incorporate stray animal control, leash laws, animal bite prevention, and training of personnel in their programs.

f) PUBLIC EDUCATION. All local jurisdictions should incorporate education covering responsible pet ownership, bite prevention, and appropriate veterinary care in their programs.

**5. POSTEXPOSURE MANAGEMENT:** This section refers to any animal exposed (see Part I.A.2.) to a confirmed or suspected rabid animal. Wild mammalian carnivores or bats that are not available or suitable for testing should be regarded as rabid animals.

a) DOGS, CATS AND FERRETS. Any illness in an exposed animal should be reported immediately to the local health department. If signs suggestive of rabies develop (e.g., paralysis, seizures, etc.), the animal should be euthanized and the head shipped for testing as described in Part I.A.9.

(1) Dogs, cats, and ferrets that have never been vaccinated and are exposed to a rabid animal should be euthanized immediately. If the owner is unwilling to have this done, the animal should be placed in strict isolation for 6 months. Isolation in this context refers to confinement in an enclosure that precludes direct contact with people and other animals. Rabies vaccine should be administered upon entry into isolation or up to 28 days before release to comply with preexposure vaccination recommendations (see Part I.B.1.a.). There are currently no USDA licensed biologics for postexposure prophylaxis of previously unvaccinated domestic animals, and there is evidence that the use of vaccine alone will not reliably prevent the disease in these animals (44).

(2) Animals overdue for a booster vaccination should be evaluated on a case-by-case basis based upon severity of exposure, time elapsed since last vaccination, number of previous vaccinations, current health status, and local rabies epidemiology to determine need for euthanasia or immediate revaccination and observation/isolation.

(3) Dogs, cats, and ferrets that are currently vaccinated should be revaccinated immediately, kept under the owner's control, and observed for 45 days. The rationale for an observation period is based in part on the potential for: overwhelming viral challenge, incomplete vaccine efficacy, improper vaccine administration, variable host immunocompetence, and immune-mediated fatality (i.e., early death phenomenon) (12,45-47).

b) LIVESTOCK. All species of livestock are susceptible to rabies; cattle and horses are the most frequently reported infected species (2). Any illness in an exposed animal should be reported immediately to the local health and agriculture officials. If signs suggestive of rabies develop, the animal should be euthanized and the head shipped for testing as described in Part I.A.9.

(1) Unvaccinated livestock should be euthanized immediately. If the animal is not euthanized, it should be observed and confined on a case-by-case basis for 6 months.

(2) Livestock exposed to a rabid animal and currently vaccinated with a vaccine approved by USDA for that species should be revaccinated immediately and observed for 45 days.

(3) Multiple rabid animals in a herd or herbivore-to-herbivore transmission are uncommon (48); therefore, restricting the rest of the herd if a single animal has been exposed to or infected by rabies is usually not necessary.

(4) Handling and consumption of tissues from exposed animals might carry a risk for rabies transmission. Risk factors depend in part on the site(s) of exposure, amount of virus present, severity of wounds, and whether sufficient contaminated tissue has been excised. If an exposed animal is to be custom or home-slaughtered for consumption, it should be done immediately after exposure, and all tissues should be cooked thoroughly. Persons handling exposed animals, carcasses, and tissues should use barrier precautions (49,50). Historically, federal guidelines for meat inspectors required that any animal known to have been exposed to rabies within 8 months be rejected for slaughter (51). USDA Food and Inspection Service (FSIS) and state meat inspectors should be notified if such exposures occur in food animals before slaughter.

Rabies virus is widely distributed in tissues of rabid animals (52-54). Tissues and products from a rabid animal should not be used for human or animal consumption (55,56) or transplantation (57). Pasteurization and cooking will inactivate rabies virus (58); therefore, inadvertently drinking pasteurized milk or eating thoroughly cooked animal products does not constitute a rabies exposure.

c) OTHER ANIMALS. Other mammals exposed to a rabid animal should be euthanized immediately. Animals maintained in USDA-licensed research facilities or accredited zoological parks should be evaluated on a case-by-case basis in consultation with public health authorities. Management options may include isolation, observation, or administration of rabies biologics.

#### 6. MANAGEMENT OF ANIMALS THAT BITE HUMANS:

a) Dogs, Cats, and Ferrets. Rabies virus is excreted in the saliva of infected dogs, cats, and ferrets during illness and/or for only a few days before illness or death (59-61). Regardless of rabies vaccination status, a healthy dog, cat, or ferret that exposes a person should be confined and observed daily for 10 days from the time of the exposure (62); administration of rabies vaccine to the animal is not recommended during the observation period to avoid confusing signs of rabies with rare adverse reactions (13). Any illness in the animal should be reported immediately to the local health department. Such animals should be evaluated by a veterinarian at the first sign of illness during confinement. If signs suggestive of rabies develop, the animal should be euthanized and the head submitted for testing as described in Part I.A.9. Any stray or unwanted dog, cat, or ferret that exposes a person may be euthanized immediately and the head submitted for rabies examination.

b) Other Animals. Other animals that might have exposed a person to rabies should be reported immediately to the local health department. Management of animals other than dogs, cats, and ferrets depends on the species, the circumstances of the exposure, the epidemiology of rabies in the area, the exposing animal's history, current health status, and the animal's potential for exposure to rabies. The shedding period for rabies virus is undetermined for most species. Previous vaccination of these animals might not preclude the necessity for euthanasia and testing.

**7. OUTBREAK PREVENTION AND CONTROL.** The emergence of new rabies virus variants or the introduction of non-indigenous viruses poses a significant risk to humans, domestic animals, and wildlife (63-70). A rapid and comprehensive response includes the following measures (71):

- a) Characterize the virus at the national reference laboratory.
- b) Identify and control the source of the introduction.
- c) Enhance laboratory-based surveillance in wild and domestic animals.
- d) Increase animal rabies vaccination rates.

- e) Restrict the movement of animals.
- f) Evaluate the need for vector population reduction.
- g) Coordinate a multiagency response.
- h) Provide public and professional outreach and education.

**8. DISASTER RESPONSE:** Animals might be displaced during and after man-made or natural disasters and require emergency sheltering (http://www.bt.cdc.gov/disasters/petshelters.asp and http://www.avma.org/disaster/default.asp) (72). Animal rabies vaccination and exposure histories often are not available for displaced animals. Disaster response creates situations where animal caretakers might lack

appropriate training and preexposure vaccination. In such situations, it is critical to implement and coordinate rabies prevention and control measures to reduce the risk of rabies transmission and the need for human PEP. Such measures include actions to:

a) Coordinate relief efforts of individuals and organizations with the local emergency operations center before deployment.

b) Examine each animal at a triage site for possible bite injuries or signs of rabies.

c) Isolate animals exhibiting signs of rabies, pending evaluation by a veterinarian.

d) Ensure that all animals have a unique identifier.

e) Administer a rabies vaccination to all dogs, cats and ferrets unless reliable proof of vaccination exists.

f) Adopt minimum standards for animal caretakers as feasible, including personal protective equipment, preexposure rabies vaccination, and appropriate training in animal handling (73).g) Maintain documentation of animal disposition and location (e.g., returned to owner, died or euthanized, adopted, relocated to another shelter, and address of new location).

h) Provide facilities to confine and observe animals involved in exposures (see Part I.B.6.).

i) Report human exposures to appropriate public health authorities (see Part I.A.3.).

#### C. PREVENTION AND CONTROL METHODS RELATED TO WILDLIFE

The public should be warned not to handle or feed wild mammals. Wild mammals and hybrids that expose persons, pets, or livestock should be considered for euthanasia and rabies diagnosis. A person exposed by any wild mammal should immediately report the incident to a healthcare provider who, in consultation with public health authorities, can evaluate the need for PEP (9, 10).

Translocation of infected wildlife has contributed to the spread of rabies (63-68,74); therefore, the translocation of known terrestrial rabies reservoir species should be prohibited. Whereas state regulated wildlife rehabilitators and nuisance wildlife control operators may play a role in a comprehensive rabies control program, minimum standards for persons who handle wild mammals should include rabies vaccination, appropriate training, and continuing education.

**1. CARNIVORES:** The use of oral rabies vaccines (ORV) for the mass vaccination of free-ranging wildlife should be considered in selected situations, with the approval of the appropriate state agencies (*14*,*75*). There have been documented successes using ORV to control rabies in wildlife in North America (*75-78*). The currently licensed vaccinia-vectored ORV is labeled for use in raccoons and coyotes. The distribution of ORV should be based on scientific assessments of the target species and followed by timely and appropriate analysis of surveillance data; such results should be provided to all stakeholders. In addition, parenteral vaccination (trap-vaccinate-release) of wildlife rabies reservoirs may be integrated into coordinated ORV programs to enhance their effectiveness. Continuous and persistent programs for trapping or poisoning wildlife are not effective in reducing wildlife rabies reservoirs on a statewide basis. However, limited population control in high-contact areas (e.g., picnic grounds, camps, and suburban areas) might be indicated for the removal of selected high-risk species of wildlife. State agriculture, public health, and wildlife agencies should be consulted for planning, coordination, and evaluation of vaccination or population reduction programs (*14*).

**2. BATS:** From the 1950's to date, indigenous rabid bats have been reported from every state except Hawaii and have caused rabies in at least 43 humans in the United States (79-92). Bats should be excluded appropriately from houses, public buildings, and adjacent structures to prevent direct association with humans (93,94). Such structures should then be made bat-proof by sealing entrances used by bats.

Controlling rabies in bats through programs designed to reduce bat populations is neither feasible nor desirable.

#### Part II: Recommendations for Parenteral Rabies Vaccination Procedures

**A. VACCINE ADMINISTRATION:** All animal rabies vaccines should be restricted to use by or under the direct supervision of a veterinarian (95), except as recommended in Part I.B.1.

**B. VACCINE SELECTION:** Part III lists all vaccines licensed by USDA and marketed in the United States at the time of publication. New vaccine approvals or changes in label specifications made subsequent to publication should be considered as part of this list. Any of the listed vaccines can be used for revaccination, even if the product is not the same as previously administered. Vaccines used in state and local rabies control programs should have at least a 3-year duration of immunity. This constitutes the most effective method of increasing the proportion of immunized dogs and cats in any population (96). No laboratory or epidemiologic data exist to support the annual or biennial administration of 3- or 4-year vaccines following the initial series.

**C. ADVERSE EVENTS:** Currently, no epidemiologic association exists between a particular licensed vaccine product and adverse events (*13,97-98*). Although rare, adverse events including vomiting, injection site swelling, lethargy, hypersensitivity, and rabies in a previously vaccinated animal have been reported. Adverse events should be reported to the vaccine manufacturer and to USDA, Animal and Plant Health Inspection Service, Center for Veterinary Biologics (Internet:

http://www.aphis.usda.gov/animal\_health/vet\_biologics/vb\_adverse\_event.shtml; telephone: 800-752-6255). No contraindication to rabies vaccination exists. Animals with a previous history of anaphylaxis can be medically managed and observed after vaccination (46).

**D. WILDLIFE AND HYBRID ANIMAL VACCINATION:** The safety and efficacy of parenteral rabies vaccination of wildlife and hybrids have not been established, and no rabies vaccines are licensed for these animals. Zoos or research institutions may establish vaccination programs to attempt to protect valuable animals, but these should not replace appropriate public health activities that protect humans (see Part I.B.1.c.2).

**E. ACCIDENTAL HUMAN EXPOSURE TO VACCINE:** Human exposure to parenteral animal rabies vaccines listed in Part III does not constitute a risk for rabies virus infection. Human exposure to vacciniavectored oral rabies vaccines should be reported to state health officials (100,101).

**F. RABIES CERTIFICATE:** All agencies and veterinarians should use NASPHV Form 51 (revised 2007), Rabies Vaccination Certificate, or an equivalent. This form can be obtained from vaccine manufacturers, NASPHV (http://www.nasphv.org/Documents/RabiesVacCert.pdf), or CDC

(http://www.cdc.gov/rabies/pdf/nasphv\_form51.pdf). The form must be completed in full and signed by the administering or supervising veterinarian. Computer generated forms containing the same information are also acceptable.

#### Age at Primary Route of Vaccinationa Inoculation Product Name For Use Ir Booster Recommended Produced By Marketed By Dosage A) MONOVALENT (Inactivated) IM<sup>c</sup> or SC RABVAC 1 Boehringer Ingelheim Boehringer Ingelheim Dogs 1 ml 3 months Annually Vetmedica, Inc. Vetmedica. Inc. Cats 1 ml3 months Annually IM or SC Licence No. 112 RABVAC 3 Boehringer Ingelheim 1 year later & triennially IM or SC Boehringer Ingelheim Dogs 1 ml 3 months 1 year later & triennially Vetmedica, Inc. Vetmedica, Inc Cats 1 ml 3 months IM or SC Licence No. 112 Horse 2 ml3 months Annually IM RABVAC 3 TF Boehringer Ingelheim 1 year later & triennially IM or SC Boehringer Ingelheim Dogs 1 ml 3 months 1 year later & triennially IM or SC Vetmedica. Inc. Vetmedica. Inc. Cats 1 ml3 months Licence No. 112 Horses 3 months Annually IM 2 mlCONTINUUM RABIES Intervet, Incorporated Intervet, Incorporated Dogs 1 ml3 months 1 year later & triennially SC 1 year later & quadrennially License No. 165A Cats 1 ml3 months SC EQUI-RAB Intervet, Incorporated Intervet, Incorporated Horses 1 ml 4 months Annually IM License No. 165A PRORAB-1 Intervet, Incorporated 3 months Annually IM or SC Intervet, Incorporated Dogs 1 ml License No. 165A Annually IM or SC Cats 1 ml 3 months Annually Sheep 2 ml3 months IM DEFENSOR 1 Pfizer, Incorporated Pfizer, Incorporated Dogs 1 ml3 months Annually IM or SC License No. 189 Cats 1 ml 3 months Annually SC Pfizer, Incorporated **DEFENSOR 3** Pfizer, Incorporated 1 ml3 months 1 year later & triennially IM or SC Dogs License No. 189 Cats 3 months 1 year later & triennially SC 1 ml Sheep 2 ml 3 months Annually IM Cattle 2 ml3 months Annually IM RABDOMUN Pfizer, Incorporated Schering-Plough Animal 1 ml 3 months 1 year later & triennially IM or SC Dogs License No. 189 1 year later & triennially SC Health Cats 3 months 1 mlSheep 2 ml3 months Annually IM Annually Cattle 2 ml3 months IM **RABDOMUN** 1 Pfizer, Incorporated Schering-Plough Animal Dogs 1 ml3 months Annually IM or SC License No. 189 Health Cats 1 ml 3 months Annually SC IMRAB 1 Merial, Incorporated Merial, Incorporated Dogs 1 ml 3 months Annually SC License No. 298 Cats 1 ml 3 months Annually SC IMRAB 1 TF Merial, Incorporated Dogs 3 months SC Merial, Incorporated 1 ml Annually License No. 298 Cats 1 ml3 months Annually SC IMRAB 3 1 year later & triennially IM or SC Merial, Incorporated Merial, Incorporated Dogs 1 ml3 months License No. 298 Cats 1 ml 3 months 1 year later & triennially IM or SC Sheep 2 ml3 months 1 year later & triennially IM or SC Cattle 2 ml3 months Annually IM or SC Horses 2 ml 3 months Annually IM or SC Ferrets 1 ml3 months Annually SC IMRAB 3 TF Merial, Incorporated Merial, Incorporated 1 year later & triennially IM or SC Dogs 1 ml 3 months License No. 298 Cats 1 ml3 months 1 year later & triennially IM or SC 3 months Annually SC Ferrets 1 ml Merial, Incorporated License No. 298 IMRAB Merial, Incorporated Cattle 2 ml3 months Annually IM or SC Large Animal Horses 2 ml3 months Annually IM or SC 2 ml1 year later & triennially IM or SC Sheep 3 months B) MONOVALENT (Rabies glycoprotein, live canary pox vector) PUREVAX Feline Merial, Incorporated Merial, Incorporated Cats Annually SC 1 ml 3 months Rabies License No. 298 C) COMBINATION (Inactivated rabies) CONTINUUM DAP-R Intervet, Incorporated Intervet, Incorporated Dogs 1 ml3 months 1 year later & triennially SC License No. 165A **CONTINUUM** Feline Intervet, Incorporated Intervet, Incorporated Cats 1 ml 3 months 1 year later & triennially SC HCP-R License No. 165A Equine POTOMAVAC + IM Merial, Incorporated Merial, Incorporated Horses 1 ml 3 months Annually IMRAB License No. 298 D) COMBINATION (Rabies glycoprotein, live canary pox vector) PUREVAX Feline 3/ Merial, Incorporated Merial, Incorporated Cats 1 ml 8 weeks Every 3 weeks until 3 months SC Rabies License No. 298 & annually 3 weeks later and annually 3 months PUREVAX Feline 4/ Merial, Incorporated Merial, Incorporated Cats 1 ml 8 weeks Every 3 weeks until 3 months SC License No. 298 & annually Rabies 3 months 3 weeks later and annually E) ORAL (Rabies glycoprotein, live vaccinia vector) - RESTRICTED TO USE IN STATE AND FEDERAL RABIES CONTROL PROGRAMS RABORAL V-RG Merial, Incorporated Merial, Incorporated Coyotes N/A N/A As determined by local Oral License No. 298 Raccoons authorities a. Minimum age (or older) and revaccinated one year later d. Subcutaneously b. One month = 28 days e. Fort Dodge Animal Health was recently acquired by Boehringer Ingelheim Vetmedica, Inc. c. Intramuscularly

Part III: Rabies Vaccines Licensed and Marketed in the U.S., 2011

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Manufacturer	Phone Number	Internet Address	
Boehringer Ingelheim Vetmedica, Inc	800-638-2226	Not available	
Intervet, Incorporated	800-441-8272	http://www.intervetusa.com	
Merial, Incorporated	888-637-4251	http://us.merial.com/	
Pfizer, Incorporated	800-366-5288	http://www.pfizerah.com	

#### **Rabies Vaccine Manufacturer Contact Information**

**ADVERSE EVENTS:** Adverse events should be reported to the vaccine manufacturer and to USDA, Animal and Plant Health Inspection Service, Center for Veterinary Biologics (Internet: http://www.aphis.usda.gov/animal\_health/vet\_biologics/vb\_adverse\_event.shtml; telephone: 800-752-6255;).

#### **REFERENCES:**

- 1. Rabies. In: Heymann D, ed. Control of communicable diseases manual. 19th ed. Washington, DC: American Public Health Association; 2008:498-508.
- 2. Blanton JD, Palmer D, Christian KA, Rupprecht CE. Rabies surveillance in the United States during 2009. J Am Vet Med Assn 2010;237(6):646-657. Available at: ttp://www.cdc.gov/rabies/resources/publications/index.html.
- 3. Castrodale L, Walker V, Baldwin J, Hofmann J, Hanlon C. Rabies in a puppy imported from India to the USA, March 2007. Zoonoses Public Health 2008;55(8-10):427-430.
- 4. CDC. Rabies in a Dog Imported from Iraq -- New Jersey, June 2008. MMWR 2008; 57:1076-1078. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5739a3.htm.
- 5. McQuiston JH, Wilson T, Harris S, et al. Importation of dogs into the United States: risks from rabies and other zoonotic diseases. Zoonoses Public Health 2008;55(8-10):421-426.
- Velasco-Villa A, Reeder SA, Orciari LA, et al. Enzootic rabies elimination from dogs and reemergence in wild terrestrial carnivores, United States. Emerg Infect Dis 2008;14(12):1849-1854. Available at: http://www.cdc.gov/EID/content/14/12/1849.htm.
- 7. Beran GW. Rabies and infections by rabies-related viruses. In: Beran GW (ed.) Handbook of zoonoses section B: Viral, second ed. Boca Raton, FL: CRC Press; 1994:307-57.
- Council of State and Territorial Epidemiologists. Public Health Reporting and National Notification for Animal Rabies. Infectious Disease Positions Statements, June 2009. CSTE, Atlanta, GA. Available at: http://www.cste.org/ps2009/09-ID-12.pdf.
- CDC. Human rabies prevention—United States, 2008. Recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR 2008;57(No. RR-3):1-28. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/rr57e507a1.htm.
- CDC. Use of reduced (4-dose) vaccine schedule for postexposure prophylaxis to prevent human rabies. Recommendations of the Advisory Committee on Immunization Practices (ACIP). MMWR 2010;59(No. RR-2):1-12. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/rr5902a1.htm.
- 11. McQuiston J, Yager PA, Smith JS, Rupprecht CE. Epidemiologic characteristics of rabies virus variants in dogs and cats in the United States, 1999. J Am Vet Med Assoc 2001;218:1939–42.
- 12. Murray KO, Holmes KC, Hanlon CA. Rabies in vaccinated dogs and cats in the United States, 1997-2001. J Am Vet Med Assoc 2009;235:691-695.
- 13. Frana TS, Clough NE, Gatewood DM, Rupprecht CE. Postmarketing surveillance of rabies vaccines for dogs to evaluate safety and efficacy. J Am Vet Med Assoc 2008;232:1000-1002.
- 14. Hanlon CA, Childs JE, Nettles VF, et al. Recommendations of the Working Group on Rabies. Article III: rabies in wildlife. J Am Vet Med Assoc 1999;215:1612–8.
- 15. Slate D, Algeo TD, Nelson KM, et al. Oral rabies vaccination in North America: opportunities, complexities, and challenges. PLoS Negl Trop Dis 2009;3(12):1-9
- 16. Council of State and Territorial Epidemiologists. Electronic laboratory reporting in the US: underfunded and under potential, or, recommendations for the implementation of ELR in the US. Policy Positions Statements, June 2009. CSTE, Atlanta, GA. Available at: http://www.cste.org/ps2009/09-SI-03.pdf.
- Council of State and Territorial Epidemiologists. Process statement for immediately nationally notifiable conditions. Policy Positions Statements, June 2009. CSTE, Atlanta, GA. Available at: http://www.cste.org/ps2009/09-SI-04.pdf.

- 18. Hanlon CA, Smith JS, Anderson GR, et al. Recommendations of the Working Group on Rabies. Article II: laboratory diagnosis of rabies. J Am Vet Med Assoc 1999;215:1444–6.
- 19. Rudd RJ, Smith JS, Yager PA, et al. A need for standardized rabies-virus diagnostic procedures: effect of coverglass mountant on the reliability of antigen detection by the fluorescent antibody test. Virus Res 2005;111:83–8.
- 20. American Veterinary Medical Association. AVMA guidelines on euthanasia, June2007. Schaumburg, IL: American Veterinary Medical Association; 2007. Available at: http://www.avma.org/issues/animal\_welfare/euthanasia.pdf.
- 21. Michigan Rabies Working Group. Humane euthanasia of bats for public health rabies testing. 2008. Available at: http://www.michigan.gov/documents/emergingdiseases/Humane\_Euthanasia\_of\_Bats-Final\_244979\_7.pdf.
- 22. CDC. Public health response to a potentially rabid bear cub -- Iowa, 1999. MMWR 1999;48:971-3. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm4842a5.htm.
- 23. Niezgoda M, Rupprecht CE. Atlanta: US Department of Health and Human Services, Centers for Disease Control and Prevention 1-16; 2006. Standard operating procedure for the direct rapid immunohistochemistry test for the detection of rabies virus antigen. National Laboratory Training Network Course. Available at: http://www.rabiesblueprint.com/IMG/pdf/DRIT\_SOP.pdf.
- 24. Lembo T, Niezgoda M, Velasco-Villa A, Cleaveland S, Ernest E, Rupprecht CE. Evaluation of a direct, rapid immunohistochemical test for rabies diagnosis. Emerg Infect Dis. 2006. Feb;12(2):310-3.
- 25. Dürr S, Naïssengar S, Mindekem R, et al. Rabies diagnosis for developing countries. PLoS Negl Trop Dis. 2008. Mar 26;2(3):e206.
- 26. Saturday GA, King R, Fuhrmann L. Validation and operational application of a rapid method for rabies antigen detection. US Army Med Dep J. 2009. Jan-Mar:42-5.
- 27. Tizard I, Ni Y. Use of serologic testing to assess immune status of companion animals. J Am Vet Med Assoc 1998;213:54–60.
- 28. Greene CE, ed. Rabies and other lyssavirus infections. In: Infectious diseases of the dog and cat. 3rd ed. London, England: Saunders Elsevier; 2006;167–83.
- 29. Rupprecht CE, Gilbert J, Pitts R, Marshall K, Koprowski H. Evaluation of an inactivated rabies virus vaccine in domestic ferrets. J Am Vet Med Assoc 1990;196:1614–6.
- 30. Moore SM, Hanlon CA. Rabies-specific antibodies: measuring surrogates of protection against a fatal disease. PLoS Negl Trop Dis. 2010. Mar 9;4(3):e595.
- 31. Aubert MF. Practical significance of rabies antibodies in cats and dogs. Rev Sci Tech 1992;11:735-60.
- 32. Muirhead TL, McClure JT, Wichtel JJ, et al. The effect of age on serum antibody titers after rabies and influenza vaccination in healthy horses. J Vet Intern Med 2008;22:654-661.
- 33. Shimazaki Y, Inoue S, Takahashi C, et al. Immune response to Japanese rabies vaccine in domestic dogs. J Vet Med B 2003;50:95-8.
- 34. Cliquet F, Verdier Y, Sagné L, et al. Neutralising antibody titration in 25,000 sera of dogs and cats vaccinated against rabies in France, in the framework of the new regulations that offer an alternative to quarantine. Rev Sci Tech 2003;22:857–66.
- 35. Rabies. In: Guidelines for the vaccination of horses. American Association of Equine Practitioners; 2009. Available at: http://www.aaep.org/rabies.htm.
- 36. National Association of State Public Health Veterinarians. Compendium of measures to prevent disease and injury associated with animals in public settings, 2007. MMWR 2007;56(RR05);1-13. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/rr5805a1.htm.
- Bender J, Schulman S. Reports of zoonotic disease outbreaks associated with animal exhibits and availability of recommendations for preventing zoonotic disease transmission from animals to people in such settings. J Am Vet Med Assoc 2004;224:1105–9.
- American Veterinary Medical Association. Private ownership of wild animals. Schaumburg, IL: American Veterinary Medical Association; 2006. Available at: http://www.avma.org/issues/policy/wild\_animal\_ownership.asp.
- American Veterinary Medical Association. Position on canine hybrids. Schaumburg, IL: American Veterinary Medical Association; 2008. Available at: http://www.avma.org/issues/policy/canine\_hybrids.asp.
- 40. Siino BS. Crossing the line: the case against hybrids. American Society for the Prevention of Cruelty to Animals, Animal Watch; 2000:22–9. Available at: http://www.petfinder.com/before-pet-adoption/case-against-hybrids.html?page-index=1&query=hybrids.
- 41. Jay MT, Reilly KF, DeBess EE, Haynes EH, Bader DR, Barrett LR. Rabies in a vaccinated wolf-dog hybrid. J Am Vet Med Assoc 1994;205:1729–32.
- 42. CDC. An imported case of rabies in an immunized dog. MMWR 1987;36:94–6. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/00000874.htm.
- 43. CDC. Imported dog and cat rabies—New Hampshire, California. MMWR 1988;37:559–60. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/00001275.htm.

- 44. Hanlon CA, Niezgoda MN, Rupprecht CE. Postexposure prophylaxis for prevention of rabies in dogs. Am J Vet Res 2002;63:1096–100.
- 45. US Government Printing Office. 9CFR113.209. Available at: http://edocket.access.gpo.gov/cfr\_2003/9cfr113.209.htm.
- 46. Greene CE, ed. Immunoprophylaxis. In: Infectious diseases of the dog and cat. 3rd ed. London, England: Saunders Elsevier; 2006;1069-1119.
- 47. Willoughby, RE. "early death" and the contraindication of vaccine during rabies treatment. Vaccine 2009;27:7173-7177.
- 48. Mansfield K, McElhinney L, Hübschle O, et al. A molecular epidemiological study of rabies epizootics in kudu (Tragelaphus strepsiceros) in Namibia. BMC Vet Res 2006;2:2.
- Viral agents. In: U.S. Department of Health and Human Services. Biosafety in Microbiological and Biomedical Laboratories. 5<sup>th</sup> edition. Washington, D.C.: U.S. Government Printing Office; 2007:234-235. Available at: http://www.cdc.gov/biosafety/publications/bmbl5/BMBL5\_sect\_VIII\_e.pdf.
- 50. Wertheim HFL, Nguyen TQ, Nguyen KAT, et al. Furious rabies after an atypical exposure. PLoS Med 2009;6(3):0264-8.
- 51. Ante-mortem inspection. In: U.S. Meat and Poultry Inspection Program. Meat and poultry inspection manual. Washington, D.C.: U.S. Government Printing Office; 1973:314 p.
- 52. Debbie JG, Trimarchi CV. Pantropism of rabies virus in free-ranging rabid red fox (*Vulpes fulva*). J Wildl Dis 1970;6(4):500-6.
- 53. Fekadu M, Shaddock JH. Peripheral distribution of virus in dogs inoculated with two strains of rabies virus. Am J Vet Res 1984;45(4):724-729.
- 54. Charlton, KM. The pathogenesis of rabies and other lyssaviral infections: recent studies. Curr Top Microbiol Immunol 1994;187:95–119.
- 55. Afshar, A. A review of non-bite transmission of rabies virus infection. Br Vet J 1979;135:142-8.
- 56. CDC. Mass treatment of humans who drank unpasteurized milk from rabid cows—Massachusetts, 1996–1998. MMWR 1999;48:228–9. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/00056759.htm.
- CDC. Public health service guideline on infectious disease issues in xenotransplantation. MMWR 2001;50(No. RR-15):1-56.
- 58. Turner GS, Kaplan C. Some properties of fixed rabies virus. J Gen Virol 1967;1:537-551.
- 59. Vaughn JB, Gerhardt P, Paterson J. Excretion of street rabies virus in saliva of cats. J Am Med Assoc 1963;184:705.
- 60. Vaughn JB, Gerhardt P, Newell KW. Excretion of street rabies virus in saliva of dogs. J Am Med Assoc 1965;193:363–8.
- 61. Niezgoda M, Briggs DJ, Shaddock J, Rupprecht CE. Viral excretion in domestic ferrets (Mustela putorius furo) inoculated with a raccoon rabies isolate. Am J Vet Res 1998;59:1629–32.
- 62. Tepsumethanon V, Lumlertdacha B, Mitmoonpitak C, Sitprija V, Meslin FX, Wilde H. Survival of naturally infected rabid dogs and cats. Clin Infect Dis 2004;39:278–80.
- Jenkins SR, Perry BD, Winkler WG. Ecology and epidemiology of raccoon rabies. Rev Infect Dis 1988;10(Suppl 4):S620–5.
- 64. CDC. Translocation of coyote rabies—Florida, 1994. MMWR 1995;44:580–7. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/00038451.htm.
- 65. Rupprecht CE, Smith JS, Fekadu M, Childs JE. The ascension of wildlife rabies: a cause for public health concern or intervention? Emerg Infect Dis 1995;1:107–14. Available at: http://www.cdc.gov/ncidod/eid/vol1no4/rupprech.htm.
- 66. Constantine DG. Geographic translocation of bats: known and potential problems. Emerg Infect Dis 2003;9:17–21. Available at: http://www.cdc.gov/ncidod/EID/vol9no1/02-0104.htm.
- 67. Krebs JW, Strine TW, Smith JS, Rupprecht CE, Childs JE. Rabies surveillance in the United States during 1993. J Am Vet Med Assoc 1994;2051695–709.
- 68. VF Nettles, JH Shaddock, RK Sikes, CR Reyes. Rabies in translocated raccoons. Am J Public Health 1979;69:601–2.
- 69. RM Engeman, KL Christensen, MJ Pipas, DL Bergman. Population monitoring in support of a rabies vaccination program for skunks in Arizona. J Wildl Dis 2003;39:746–50.
- 70. Leslie MJ, Messenger S, Rohde RE, et al. Bat-associated rabies virus in skunks. Emerg Infect Dis 2006;12:1274–7. Available at: http://www.cdc.gov/ncidod/EID/vol12no08/05-1526.htm.
- 71. Rupprecht CE, Hanlon CA, Slate D. Control and prevention of rabies in animals: paradigm shifts. Dev Biol (Basel). 2006;125:103-11.
- 72. Pets Evacuation and Transportations Standards Act of 2006. Available at: http://frwebgate.access.gpo.gov/cgibin/getdoc.cgi?dbname=109\_cong\_public\_laws&docid=f:publ308.109.pdf.
- 73. National Animal Control Association guidelines. Available at: http://www.nacanet.org/guidelines.html.

- 74. Chipman R, Slate D, Rupprecht C, Mendoza M. Downside Risk of Translocation. Dodet B, Fooks AR, Muller T, Tordo N, and the Scientific & Technical Department of the OIE (eds): Towards the Elimination of Rabies in Eurasia. Dev Biol. Basel, Karger 2008;131:223-232.
- 75. Slate D, Rupprecht CE, Rooney JA, Donovan D, Lein DH, Chipman RB. Status of oral rabies vaccination in wild carnivores in the United States. Virus Res 2005;111:68–76.
- 76. Sidwa TJ, Wilson PJ, Moore GM, et al. Evaluation of oral rabies vaccination programs for control of rabies epizootics in coyotes and gray foxes: 1995-2003. J Am Vet Med Assoc 2005;227:785-792.
- 77. MacInnes CD, Smith SM, Tinline RR, et al. Elimination of rabies from red foxes in eastern Ontario. J Wildl Dis 2001;37:119-132.
- 78. Rosatte RC, Power MJ, Donovan D, et al. Elimination of arctic variant of rabies in red foxes, metropolitan Toronto. Emerg Infect Dis 2007;13(1)25-27. Available at: http://www.cdc.gov/ncidod/EID/13/1/25.htm.
- 79. Messenger SL, Smith JS, Rupprecht CE. Emerging epidemiology of bat-associated cryptic cases of rabies in humans in the United States. Clin Infect Dis 2002;35:738–47.
- 80. CDC. Human rabies—California, 2002. MMWR 2002;51:686–8. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5131a4.htm.
- 81. CDC. Human rabies—Tennessee, 2002. MMWR 2002;51:828–9. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5137a2.htm.
- 82. CDC. Human rabies—Iowa, 2002. MMWR 2003;52:47–8. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5203a3.htm.
- 83. CDC. Human death associated with bat rabies—California, 2003. MMWR 2004;53:33–5. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5302a4.htm.
- 84. CDC. Recovery of a patient from clinical rabies, Wisconsin, 2004. MMWR 2004;53:1171–3. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5350a1.htm.
- 85. CDC. Human rabies—Mississippi, 2005. MMWR 2006;55:207–8. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5508a4.htm.
- 86. CDC. Human rabies—Indiana and California, 2006. MMWR 2007;56:361–5. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5615a1.htm.
- 87. CDC. Human rabies—Minnesota, 2007. MMWR 2008;57:460-462. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5717a3.htm.
- 88. CDC. Human rabies—Missouri, 2008. MMWR 2009;58:1207-9. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5843a3.htm.
- 89. CDC. Human rabies—Kentucky/Indiana, 2009. MMWR 2010;59:393-6. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5913a3.htm.
- 90. CDC. Human rabies—Virginia, 2009. MMWR 2010;591236-8. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5938a3.htm.
- 91. CDC. Presumptive abortive human rabies—Texas, 2009. MMWR 2010;59:185-90. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5907a1.htm.
- 92. CDC. Human rabies-Michigan 2009. MMWR 2011;60:437-40. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm6014a1.htm?s\_cid=mm6014a1\_w
- 93. Greenhall AM. House bat management. US Fish and Wildlife Service, Resource Publication 143;1982. Jamestown, ND: Northern Prairie Wildlife Research Center Online. Available at: http://www.npwrc.usgs.gov/resource/mammals/housebat/index.htm.
- 94. Greenhall, AM. Frantz, SC. Bats. In: Hygnstrom SE, Timm RM, Larson GE, eds. Prevention and Control of Wildlife Damage 1994. Available at: http://icwdm.org/handbook/mammals/bats.asp.
- 95. American Veterinary Medical Association. Model rabies control ordinance. Schaumburg, IL: American Veterinary Medical Association 2008. Available at: http://www.avma.org/issues/policy/AVMA-Model-Rabies-Ordinance.pdf.
- 96. Bunn TO. Canine and feline vaccines, past and present. In Baer GM, ed. The natural history of rabies. 2nd ed. Boca Raton, FL: CRC Press; 1991:415–25.
- 97. Macy DW, Hendrick MJ. The potential role of inflammation in the development of postvaccinal sarcomas in cats. Vet Clin North Am Small Anim Pract 1996;26:103–9.
- 98. Gobar GM, Kass PH. World wide web-based survey of vaccination practices, postvaccinal reactions, and vaccine site-associated sarcomas in cats. J Am Vet Med Assoc 2002;220:1477–82.
- 99. Kass PH, Spangler WL, Hendrick MJ, et al. Multicenter case-control study of risk factors associated with development of vaccine-associated sarcomas in cats. J Am Vet Med Assoc 2003;223:1283–92.
- 100. Rupprecht CE, Blass L, Smith K, et al. Human infection due to recombinant vaccinia-rabies glycoprotein virus. N Engl J Med 2001;345:582–6.
- 101. CDC. Human vaccinia infection after contact with a raccoon rabies vaccine bait— Pennsylvania, 2009. MMWR 2009; 58:1204-7. Available at: http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5843a2.htm.