Application of the AVMA Guidelines for the Depopulation of Animals to Biomedical Research



OLAW Online Seminar September 26, 2019

Samuel Cartner, DVM, PhD, DACLAM, University of Alabama at Birmingham Jennifer Pullium, MVB, DACLAM, New York University School of Medicine Axel Wolff, MS, DVM, NIH, Office of Laboratory Animal Welfare

PHS Policy and Euthanasia

PHS Policy on Humane Care and Use of Laboratory Animals (IV.C.1.g.)

"Methods of euthanasia used will be consistent with the recommendations of the American Veterinary Medical Association (AVMA) Panel on Euthanasia, unless a deviation is justified for scientific reasons in writing by the investigator." *

* AVMA Guidelines for Euthanasia of Animals: 2013 Edition or succeeding revised editions.



U.S. Department of Health and Human Services

National Institutes of Health Office of Laboratory Animal Welfare

Public Health Service Policy on Humane Care and Use of Laboratory Animals





AVMA Guidelines for the Depopulation of Animals: 2019 Edition

- Guides veterinarians in making humane decisions when large numbers of animals must be killed quickly in response to a disease outbreak or natural disaster.
- Depopulation refers to the rapid destruction of a population of animals in response to urgent circumstances with as much consideration given to the welfare of the animals as practicable.







AVMA Guidelines for the Depopulation of Animals: 2019 Edition and the Disaster Plan

 It is recommended that the disaster plan include contingencies for applying the AVMA Depopulation Guidelines in an emergency.

AVMA Guidelines for the **Depopulation of Animals:** 2019 Edition

Members of the Panel on Animal Depopulation

Steven Leary, DVM, DACLAM (Chair); Fidelis Pharmaceuticals, High Ridge, Missouri Raymond Anthony, PhD (Ethicist); University of Alaska Anchorage, Anchorage, Alaska Sharon Gwaltney-Brant, DVM, PhD, DABVT, DABT (Lead, Companion Animals Working Group); Veterinary Information Network, Mahomet, Illinois Samuel Cartner, DVM, PhD, DACLAM (Lead, Laboratory Animals Working Group); University of Alabama at Birmingham, Birmingham, Alabama Renee Dewell, DVM, MS (Lead, Bovine Working Group); Iowa State University, Ames, Iowa Patrick Webb, DVM (Lead, Swine Working Group); National Pork Board, Des Moines, Iowa Paul J. Plummer, DVM, DACVIM-LA (Lead, Small Ruminant Working Group); Iowa State University, Ames, Iowa Donald E. Hoenig, VMD (Lead, Poultry Working Group); American Humane Association, Belfast, Maine William Moyer, DVM, DACVSMR (Lead, Equine Working Group); Texas A&M University College of Veterinary Medicine, Billings, Montana Stephen A. Smith, DVM, PhD (Lead, Aquatics Working Group): Virginia-Maryland College of Veterinary Medicine, Blacksburg, Virginia Andrea Goodnight, DVM (Lead, Zoo and Wildlife Working Group); The Living Desert Zoo and Gardens, Palm Desert, California P. Gary Egrie, VMD (nonvoting observing member); USDA APHIS Veterinary Services, Riverdale, Maryland Axel Wolff, DVM, MS (nonvoting observing member); Office of Laboratory Animal Welfare (OLAW), Bethesda, Maryland AVMA Staff Consultants Cia L. Johnson, DVM, MS, MSc; Director, Animal Welfare Division Emily Patterson-Kane, PhD; Animal Welfare Scientist, Animal Welfare Division The following individuals contributed substantively through their participation in the Panel's Working Groups, and their assistance is sincerely appreciated. Companion Animals-Yvonne Bellay, DVM, MS; Allan Drusys, DVM, MVPHMgt; William Folger, DVM, MS, DABVP; Stephanie Janeczko, DVM, MS, DABVP, CAWA; Ellie Karlsson, DVM, DACLAM; Michael R. Moyer, VMD; Phillip Raclyn, DVM Laboratory Animals—Robert J. Adams, DVM, DACLAM; Michael Huerkamp, DVM, DACLAM; Kathleen Pritchett-Corning, DVM, DACLAM; Jennifer Pullium, MVB, DACLAM; Helen Valentine, DVM, MS, DACLAM Bovine-Joseph Clark, DVM; Steve Ensley, DVM, PhD; John Gilliam, DVM, MS, DACVIM, DABVP; Michael Gilsdorf, DVM, MS; Temple Grandin, PhD; Dee Griffin, DVM, MS; Michael Sanderson, DVM, MS, DACVPM (Epidemiology); Jan Shearer, DVM, MS, DACAW; David Sjeklocha, DVM, MS, DACVPM Swine—Peggy Anne Hawkins, DVM, MS; Robert Meyer, DVM, DACVAA; Alejandro Ramirez, DVM, MPH, PhD, DACVPM, Patricia V. Turner, DVM, DVSc, MS, DACLAM, DABT, DECAWBM; Sherrie Webb, MS Small Ruminant—Wyatt Frampton, DVM, MPH; Jan Shearer, DVM, MS, DACAW; Terry Taylor, DVM; Glen Zebarth, DVM Poultry-leff Erickson, DVM; Eric Benson, PhD; Michael Czarick III, MS; Brian Fairchild, PhD; Michelle Kromm, DVM, MPH, MAM, DACPV; Maureen Lee-Dutra, DVM, MPVM; Beth S. Thompson, JD, DVM; Bruce Webster, PhD; Eric Willinghan, DVM, MBA, MS; Kenneth Anderson, PhD Equine—Sam M. Crosby IV, DVM; Brandon Dominguez, DVM, MS, DACVPM; Noberto Espitia, PhD; Carl Heckendorf, DVM; Harold Kloeze, DVM, DVSc; Nathaniel Messer IV, DVM, DABVP-Equine; James Morehead, DVM; Harry Werner, VMD Agautics—Lori Gustafson, DVM, PhD; Kathleen Hartman, DVM, PhD Zoo and Wildlfe—David Miller, DVM, PhD, DACZW, Joe Gaudell, PhD; Anchony J. DeNicola, PhD; Dennis Ferraro, MS; Thomas Meehan, DVM; Lisa Pennisi, PhD; Julia Ponder, DVM, MPH; Stephen M. Vantassel, MS, MATS; Yvonne Nadler, DVM, MPH; Craig Harms, DVM, PhD, DACZM; Matthew Capitanio, DVM Expert At-Large-Gary Flory, BS AVWA GUIDELINES FOR THE DEPOPULATION OF ANIMALS: 2019 EDITION







Depopulation of Laboratory Animals

Samuel C. Cartner, DVM, PhD, DACLAM

Director, Animal Resources Program

University of Alabama at Birmingham

September 26, 2019



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Humane Endings Guidance: What is it? AVMA Guidelines on...

Euthanasia (2013)



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Humane Slaughter (2016)



Depopulation (2019)



"A method of killing that minimizes pain, distress, and anxiety experienced by the animal prior to loss of consciousness, and causes rapid loss of consciousness followed by cardiac or respiratory arrest and death".

Humane Endings Guidance: Depopulation What is Depopulation?

- Refers to the rapid destruction of a population of animals in response to urgent circumstances with as much consideration given to the welfare of the animals as practicable.
- May employ euthanasia techniques, but not all depopulation methods meet the [AVMA] criteria for euthanasia.
- Balances rapid response with prevention of further devastation and suffering with the most humane method possible.
- Occurs when doing nothing can result in greater animal suffering and endanger responders.



Humane Endings Guidance: Depopulation

Classifying Methods

Preferred

- These methods are given highest priority and should be utilized preferentially when developing emergency response plans and when circumstances allow reasonable implementation during emergencies.
- They may correspond to AVMA Guidelines for the Euthanasia of Animals or the AVMA Guidelines for the Humane Slaughter of Animals techniques but adjusted for situational considerations.



Humane Endings Guidance: Depopulation

Classifying Methods

- These methods are permitted only when the circumstances of the emergency are deemed to constrain the ability to reasonably implement a preferred method.
- Examples include zoonotic disease risk, response time, human safety, depopulation efficiency, deployable resources, equipment, animal access, disruption of infrastructure, and disease transmission risk.



Humane Endings Guidance: Depopulation

Classifying Methods

Not Recommended

- These methods should be considered ONLY when circumstances preclude the reasonable implementation of any of the "preferred or permitted in constrained circumstances" methods, <u>AND</u> when the risk of doing nothing is deemed likely to have a reasonable chance of resulting in significantly more animal suffering than that associated with the proposed depopulation technique.
- Examples of such situations include, but are not limited to, structural collapse or compromise of buildings housing animals, complete inability to safely access animals for prolonged period of time or any circumstance that poses a severe threat to human life.
- Not Recommended ≠ Unacceptable



Depopulation: Examples

Low-medium expansion foam for poultry

- Involves covering birds with blanket of foam, to a depth of 15-30 cm
- May be used in a variety of housing types (e.g., open sided, naturally ventilated; damaged structures that are unsafe to enter)
- Birds must be contained to the floor



Photo: Thornber PM, Rubira RJ, Styles DK. Humane killing of animals for disease control purposes. Rev. sci. tech. Off. int. Epiz., 2014;33(1):303-310. Available at: https://pdfs.semanticscholar.org/a897/4bbf1be6178312f0a7d2863409767980d20e.pdf



Planning for Emergency Killing in Research Settings

Michael J. Huerkamp, DVM, DACLAM Director and Attending Veterinarian Division of Animal Resources Emory University Atlanta, GA

2018 Humane Endings Symposium November 4, 2018 Chicago, IL



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Acknowledgments

Panel on Depopulation

• Laboratory Animal Working Group

- Dr. Sam Cartner, Chair (UAB, Birmingham, AL)
- Dr. Bob Adams (Johns Hopkins, Baltimore, MD)
- Dr. Emily Patterson-Kane (AVMA, Schaumburg, IL)
- Dr. Kate Pritchett-Corning (Harvard, Cambridge, MA)
- Dr. Jennifer Pullium (NYU Langone Med Ctr, NY, NY)
- Dr. Helen Valentine (U of Illinois, Urbana-Champaign, IL)
- Dr. Michael Huerkamp (Emory, Atlanta, GA)
- AVMA: Dr. Cia Johnson, Lindsey McKinney

Images: Dr. Ryan Curtis, Dr. Bob Livingston



Depopulation

Small laboratory and wild-caught rodents

Preferred: Euthanasia techniques

- Carbon Dioxide (CO₂)
- Isoflurane overdose
- Intraperitoneal (IP) euthanasia solution
- Physical methods

- Compatible: confluent monolayer + CO₂
- Incompatible: CO₂ in home cage
- Pre-charged chamber + high-flow







Depopulation

Laboratory dogs, cats, ferrets, rabbits, sheep, goats, and swine

Preferred: Injectable anesthetic overdose or injectable euthanasia solutions

- Two-step method:
 - 1. anesthesia followed by
 - 2. physical or adjunctive method or injection of Potassium chloride (KCl) or Magnesium sulfate (MgSO₄)

- Use of effective compounded or non-pharmaceutical-grade, expired anesthetic or euthanasia agent
- Reuse of needles (5x)



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Depopulation Nonhuman Primates

Preferred: Euthanasia or slaughter techniques

- Two-step method:
 - 1. anesthesia followed by
 - 2. physical or adjunctive method or injection of KCl or MgSO₄

- Use of effective compounded or non-pharmaceutical-grade (NPG), expired anesthetic, or euthanasia agent
- Gunshot by trained operator in appropriate outdoor environment



Depopulation Aquatic Vertebrates

Preferred: Euthanasia or slaughter techniques

- Rapid chilling of small tropical fish
- Anesthesia followed by adjunctive method
- Pithing, blunt force followed by adjunctive method

Permitted in Constrained Circumstances

• Use of effective compounded or non-pharmaceutical-grade, expired anesthetic, euthanasia, or toxic agent



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Depopulation Avian and Poultry

Preferred: Euthanasia or slaughter techniques

- Inhaled or injectable agents
- Anesthesia followed by adjunctive method

Permitted in Constrained Circumstances

• Use of effective compounded or non-pharmaceutical-grade, expired anesthetic, or euthanasia agent



Depopulation Special Considerations

Dangerous animals

• Keep nets, snake hooks, darting apparatus on hand

Animals exposed to hazardous agents (e.g., BSL3 or 4)

• Should not require responders to enter primary containment

Fetal or neonatal animals

- Altricial: physical methods (e.g., cervical dislocation, decapitation)
- Precocial: euthanized as adults of the same species

Embryonated poultry eggs

Cooling or freezing



Agricultural vs. Laboratory Settings



Facility & Program Attributes Research Facilities

- Physical plant
- Protective distances
- High value, sometimes irreplaceable animals
- High population density
- Epizootic approach
- Extensive experience
- Mandatory emergency preparedness plans
- Attending Veterinarian



Facility & Program Attributes Physical Plant

- Indoors containment, resilient construction, system redundancy
- Utilities: hardened supply with or without self-generation, emergency power









Facility & Program Attributes Distance Matters

- Biosecurity: isolated from different institutional animals
- Minimal fomites such as service vehicles between institutions



Photo: Unsplash @roryhennessey



Facility & Program Attributes High Value Census

- Irreplaceable attributes and high value of many research models
 - Novel founder mice: \$7,500 each
 - Cryopreservation of genetics





Facility & Program Attributes Dense, Mobile Populations

- High concentration of large numbers
 - Mice: ~30-40/Net Square Foot (NSF), 100K+/institution
 - Broilers: 1-2/NSF (National Chicken Council)
- Mobility of some housing systems: racks and pens on wheels





Facility & Program Attributes Epizootics

- Different impact
- Usually managed in-place
- Isolate, test, and cull by cage, room or greater scale





Emergency Planning Regulations and Requirements

- "Facilities must ... have a disaster plan" (p. 35)
- "Animals that cannot be relocated or protected...<u>must</u> be humanely euthanized." (p. 35)
- May involve ranking of species
- Defined decision tree
- Communication most likely disrupted





Emergency Planning Attending Veterinarian

- Required by the Guide for the Care and Use of Laboratory Animals (Guide)
 - Responsible for the health and well-being of all laboratory animals (p. 14)
 - Facilities must have a disaster plan (p. 35)
- Allowed to deviate from conventional practices, AVMA guidelines
 - Professional judgement
- Depopulation guidelines
 - IP injection of 70% ethanol to mice, immature rats
 - Use of compounded or non-pharmaceutical-grade injectable anesthetics, euthanasia agents
 - Expired drug deployment
 - Needle reuse





Disaster Response Plan





Natural Hazards

HAZARD AND VULNERABILITY ASSESSMENT TOOL NATURALLY OCCURRING EVENTS								
	PROBABILITY	SEVERITY = (MAGNITUDE - MITIGATION)						
EVENT		HUMAN IMPACT Possibility of death or injury	PROPERTY IMPACT Physical losses and damages	BUSINESS IMPACT	PREPARED- NESS Preplanning	INTERNAL RESPONSE Time, effectivness, resouces	EXTERNAL RESPONSE Community/ Mutual Aid staff and supplies	RISK.
Tornado	3	2	3	2	2	2	2	72%
Pandemic Flu	1 - T	3	0	2	2	2	2	20%
evere Drought with	1	1	1	2	3	3	3	24%
Ice Storm	2	1	2	2	2	2	1	37%
Severe Thunderstorm	3	1	1	1	1	1	1	33%
Earthquake	1 1	1	1	1	3	3	3	22%
Hurricane	1	1	2	2	2	2	2	20%
Blizzard	1	1	1	2	2	2	1	17%
Flood, External	1	O	1	1	3	2	2	17%
Snow Fall	1	1	i .	1	1	1	1	11%
Temperature Extremes	1	1	1	1	1	1	1	11%
Tidal Wave	0	1			1.		1	0%
Wild Fire	0							0%
Landslide	0							0%
Dam Inundation	0				1		1	0%

2008 Analysis Adapted From Kaiser Foundation HVA Emory University Office of Critical Event Preparedness and Response



Research Institution Census

(approximate rank order by head-count)

- 1. Mice
- 2. Zebra fish
- 3. Rats
- 4. Guinea pig
- 5. Rabbit
- 6. Hamster
- 7. NHP

- 8. Livestock
- 9. Dog

10. Cat

- 11. Amphibian
- 12. Ferret
- 13. Gerbil
- Other species of birds, amphibians, fish and rodents



"Big Academe" Scenario Mass Euthanasia / Total Depopulation

Pandemic flu

- Flu: state quarantined, animal food milled out-of-state and depleted
- 20,000 mouse cage depopulation
- Euthanex SmartBox[™] chambers x 7
 - 16 cages + 400 L CO₂ per cycle
 - 12,310 L per 50-lb CO₂ tank
 - Mass depopulation requires 21 *FULL* 50-lb tanks,
 - 45 hrs x 14 people
- 35 gallon waste containers x 20 stations x 40 people = 20 hrs and 20 50-lb CO₂ tanks







Sizable Populations of Large Animals To anesthetize 3,000 NHPs requires ~240 bottles of Telazol or Ketamine HCl







Conclusions

- Complete pre-emptive depopulation event is unlikely
 - Hardened facilities/programs + unpredictable, fast-moving event
- Adapt and apply the Panel on Euthanasia & Panel on Humane Slaughter guidelines
 - AV can assess effectiveness of non-pharmaceutical-grade and expired drugs
- Include in disaster planning & practice table top exercises
- Inventory anesthesia and euthanasia agents
- Advocate for cryo storage





Disaster Response and Recovery

Jennifer K. Pullium, MVB, DACLAM Senior Director, Division of Comparative Medicine NYU School of Medicine

OSR OFFICE OF SCIENCE & RESEARCH

Topics Covered

Disaster response

- Incident command
- Vivarium
 - Animals
 - Staffing
- Disaster recovery
 - Short-term measures
 - Other considerations and innovative planning















Incident Command

- Integrated institutional disaster response
- Participants:
 - Animal Resources Director/AV
 - Facilities and environmental health and safety
 - Information technology
 - Institutional senior leadership
 - Public relations
 - Purchasing
 - Human resources and labor relations





Responsibilities

- Everyone in the institution has their area of focus
 - Don't assume everyone will (or is able) to drop everything and assist you
- Need to ensure multiple people are capable of leading the vivarium disaster response
 - Don't assume the Director or AV will be onsite



Responsibilities

- Issues that previously would have required multiple meetings and much hand wringing – now just require decisions
- Remember indecision is a decision
- Differences between clinical and research staff become more apparent





- Personnel safety comes first
 - May be working in an unfamiliar site
 - Need to know the emergency egress(es)
- Began by rounding all animal facilities and satellites for: animal health, food, water, +/- bedding, temp, relative humidity
 - Had already stockpiled food, bedding, etc.
 - Bottled water (water cooler) for everything



- Consider a stash of cash (few thousand)
- Relocated satellite animals for easier care
- Store emergency supplies in multiple locations





- To what point do you allow environmental parameters outside the *Guide* or Animal Welfare Act before euthanasia?
 - Days? Week(s)?
 - Assuming the animals are clinically normal...
 - Are you going to euthanize everything because there are no air exchanges but temp and relative humidity are OK?



- Just because the storm/earthquake/event has passed...doesn't mean the disaster is over
 - Couple weeks after the storm, another building was not going to resume utilities
 - ~4,000 cages; 8 different species; 50 PIs
 - Winter is coming...
- Recommend local institutions determine what housing can be offered in a disaster



Smilow rescue











Staffing

- Don't assume people will come to work just because they are "essential personnel"
- Effective disciplinary program
- Have people staying onsite in advance
- Make it as comfortable as possible
 - Hotel room(s), food (before, during, after)
- Satellite phones



Press





Press

- Myths from my residency training:
 - Lab animal vets don't speak to the press
 - Your Public Relations (PR) people will take care of everything
 - Just give them a briefing about animal research
- Realities I learned the hard way:
 - No one knows your operation as well as you
 - A briefing to PR folks is not enough
 - If you want something done right, do it yourself



Press

- News cycle isn't aware that you've been working almost 48 hrs with no sleep
- "Not available for comment" = "Not sitting by the phone with nothing to do but answer endless questions"
- Encourage staff not to read internet comments
- And if you still think briefing PR folks is enough...



Personnel

- Now members of an "exclusive club"
 - Find a way to communicate this fact to outsiders
- Mandatory Post Traumatic Stress Disorder (PTSD) counseling
- Look for hidden opportunities with previously difficult staff members
- Look after each other
 - Make sure people are eating, sleeping, home OK



Recovery: Short-term Measures

- Replacement animals
 - Cryo makes it much easier
 - Takes time to bring back and need housing
- Replacement facilities
 - Temporary and permanent
 - Better hope you've been nice to people



What can we do to really plan?

- Tabletop exercises are <u>not</u> enough
 - Reading comprehension test of disaster plan
 - Doesn't tell you how anyone would perform in an actual emergency
- Tactical Decision Games (TDGs)
 - Developed by the military
 - Civilian use by industrial psychologists
 - Crichton, M. Horiz. Psychol. 10, 7–22 (2001)
 - Can't simulate a disaster, but failure is not an option





What can we do to really plan?

- What really matters are the human factors (non-technical skills)
- Participants are given an emergency scenario and asked to make decisions under stress
 - Distractions, time limits, role play
 - Recommended for ALL levels of employees
 - Pullium et al., Nature 2014, Oct 23
- Leadership for Disaster Response (LeDR)
 - Conducted TDGs throughout the US, as well as UK, Ireland, Australia (academia and industry)



Depopulation guidelines

- We were fortunate in not having to depopulate for Hurricane Sandy
- Useful guidelines if we had to depopulate:
 - Combining cages of mice for euthanasia (perhaps in large containers)
 - Using all available pentobarbital (possibly expired)
 - Re-using needles until dull



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Submit to the chat box in the GoToMeeting control panel







How can depopulation be accomplished to minimize compassion fatigue?







Who should be included in the decision to depopulate?





Question 3

Please talk more about networking with other nearby Institutions during an emergency. How do you initiate and maintain arrangements so that they are in place when a disaster occurs?





Question 4

Can you share some strategies for developing talking points on depopulation for media communications?







Submit to the chat box in the GoToMeeting control panel





21st Century Cures Act: Next Steps



OLAW Online Seminar December 5, 2019