

Regulatory Considerations for Using Pharmaceutical Products in Research Involving Laboratory Animals

Dorothy Bailey, DVM, and
Neal Bataller, DVM

Center for Veterinary Medicine
Food and Drug Administration

OLAW Online Webinar
June 4, 2015

The Terms “Pharmaceutical- and Non-pharmaceutical-grade Substances”

Animal Welfare Division, OPRR
November 22, 1993

The use of non-pharmaceutical grade chemical compounds

This is to provide an Office for Protection from Research Risks (OPRR) interpretation of the Public Health Service Policy on Humane Care and Use of Laboratory Animals regarding the use of non-pharmaceutical grade chemical compounds in physiological preparations involving laboratory animals.

The use of non-pharmaceutical grade chemical compounds in experimental animals under certain circumstances has been, and will continue to be, a necessary and acceptable component of biomedical research. Their use should be based on:

...acceptable veterinary...pharmaceutical grade product

1. scientific necessity,
2. non-availability of an acceptable veterinary or human pharmaceutical grade product, and
3. specific review and approval by the Institutional Animal Care and Use Committee (IACUC).

In preparing and reviewing proposals to use non-pharmaceutical grade products, investigators and IACUCs should consider a number of related animal welfare and scientific factors including safety, efficacy, and the inadvertent introduction of research complicating variables. While issues such as sterility, pyrogenicity, stability, pharmacokinetics and quality control can be assumed to have been addressed during the course of producing pharmaceutical grade drugs, the same can not always be said for substances produced in the research laboratory using non-pharmaceutical grade chemical compounds. ~~Cost savings alone is not~~ an adequate justification for using non-pharmaceutical grade compounds in animals.

Non-pharmaceutical grade products...

Although the potential animal welfare consequences of complications are less evident in non-survival studies, the scientific issues remain the same. The principles and need for professional judgement outlined above still apply.

Please call if I may provide further information.

Sincerely,



Nelson L. Garnett, D.V.M.
Director
Division of Animal Welfare, OPRR

Bridging Our Terminology

Send questions to
OLAWDPE@mail.nih.gov



Regulatory Considerations for Using Pharmaceutical Products in Research Involving Laboratory Animals



Today's Topics

- Investigational Use
- Legal Choices for Clinical Use
- Extralabel Drug Use
- Drug Compounding
- Euthanasia Issues

Definitions

- **Test article** – drug is focus of research.
- **Clinical use** – use of drug in the care of an animal or population of animals (drug is not the focus of research).
- **MUMS** – Minor Use & Minor Species.
- **Major species** – horses, dogs, cats, cattle, pigs, chickens, turkeys. All other species (except human) are considered minor by FDA.

Investigational Use (Test Articles)

1. Use in basic research
2. Use in pilot and pre-clinical studies for drug development
(21 CFR 312 for human drugs and 21 CFR 511.1(a) for animal drugs)
3. Use under an Investigational New Animal Drug (INAD) file to collect data to support an approval or conditional approval
(21 CFR 511.1(b))

Legal Choices for Clinical Care of Research Animals

- Use an approved or conditionally approved drug
- Use an indexed drug
- Use an approved drug extralabel
- Use a compounded drug made from an approved drug

Approved Drugs

- Well-controlled studies conducted to support safety and effectiveness.
- Manufacturing studies conducted to demonstrate strength and purity of drug.
- Data reviewed by FDA scientists.
- Approved drugs are manufactured in accordance with current Good Manufacturing Practices (cGMPs) to ensure quality and consistency.
- Extralabel use is allowed under certain circumstances (21 CFR 530).

How to Recognize Approved Drugs

- Drug label may have a New Animal Drug Application (NADA) number (XXXXXX).
- Generic drugs may have an Abbreviated New Animal Drug Application (ANADA) number (2XXXXX).
- If neither an NADA or ANADA number is listed in the labeling, a searchable database of approved animal drugs can be found at: <http://www.accessdata.fda.gov/scripts/animaldrugsatfda/>.
- Freedom of Information (FOI) summaries for approved drugs can be found at: <http://www.fda.gov/AnimalVeterinary/Products/ApprovedAnimalDrugProducts/FOIADrugSummaries/ucm056898.htm>.

Finding Information About Drugs

How can one find detailed information about drugs and their formulations?

In addition to the resources listed in the previous slide, DailyMed (<http://dailymed.nlm.nih.gov/dailymed/index>) is a resource website from the National Library of Medicine. It provides information about marketed drugs in the US.

DailyMed is the official provider of FDA label information (package inserts). The drug labeling information on this website is the most recent submitted to the FDA.

Conditionally Approved Drugs

- Limited to minor uses (rare disease or condition in one of the 7 major species) and minor species.
- All approval requirements must be met to the current standard except effectiveness.
- FDA has determined there is reasonable expectation that the drug will be effective.
- The drug can be marketed for up to 5 years (through annual renewals) while the drug sponsor completes the effectiveness section to the full standard to receive a complete approval.
- Extralabel use is prohibited.

How to Recognize a Conditionally Approved Drug

- The labeling will contain a number for the Application for Conditional Approval of a New Animal Drug (CNADA).
- Conditionally approved indications cannot be combined in the same labeling with approved indications (i.e., labeling will only contain indications that are conditionally approved).

The Index of Legally Marketed Unapproved New Animal Drugs for Minor Species (Index)

- Indexing is an alternative pathway to legal marketing status.
- Only for non-food producing minor species and non-food early life stages of food-producing minor species.
- A **food-producing species** is any minor species of which some members are bred, cultured, farmed, ranched, caught, trapped or otherwise harvested for the purpose of having animals or edible products of the animals commercially distributed for consumption by humans or food-producing animals in the US.

Food-producing Species

Indexing is prohibited for a species that is considered to be a food-producing species, no matter where it is housed.

For example, use of an indexed product is prohibited in rabbits (even if the rabbits are being used in laboratory research) because they are considered a food-producing species in the US.

Adding a Drug to the Index

Adding a drug product to the Index is a 3-step process:

- Step 1 is a request for determination of eligibility which includes:
 - Drug information (intended use, conditions of use);
 - Summary of the manufacturing process; and
 - All safety information concerning the use of the drug except target animal safety.

Adding a Drug to the Index

- Step 2 is selection of the qualified expert panel:
 - At least three members;
 - Qualified by training and experience to review the target animal safety and effectiveness of the drug product; and
 - Members must not have a conflict of interest and may not be FDA employees.

Adding a Drug to the Index

- Step 3 is a request for addition to the Index which includes:
 - Findings of the expert panel, draft labeling, and a draft Freedom of Information (FOI) summary.
 - The expert panel must unanimously agree that the drug is safe and effective for the intended use.
 - If the FDA agrees with the expert panel, the drug is included in the Index.

Indexed Drugs

- The holder of an indexed product signs a commitment to manufacture the product using cGMPs to ensure quality and consistency.
- Indexed products have the same reporting standards as approved products (e.g., adverse drug events, annual reports).
- Extralabel use is prohibited.

How to Recognize an Indexed Drug

- The labeling will include a Minor Species Index File (MIF) number.
- Currently, the labeling of all indexed products includes the following language required by statute: “Not Approved by the FDA – legally marketed as an FDA Indexed Product under MIF 9XXXXX.”

Indexed Products List

The Index is maintained on the FDA website and includes the labeling and FOI summary for each drug product

<http://www.fda.gov/AnimalVeterinary/DevelopmentApprovalProcess/MinorUseMinorSpecies/ucm125452.htm>.

Two different extended release buprenorphine products for laboratory rats and mice:

- Currently not readily available.
- Holders are working on returning these to market.

What You Can Do

- This process is well-suited for drugs intended for use in laboratory animals, excluding rabbits.
- If there is a product you need, contact the drug manufacturer and suggest having the product indexed.
- Offer to sit on an expert panel for an indexed product.
- If you have questions, you can contact Dr. Dorothy Bailey at 240-402-0565 or dorothy.bailey@fda.hhs.gov.

What is the Difference?

- **Law**
 - Act of Congress
 - Federal Food, Drug & Cosmetic Act
- **Regulation**
 - Proposed by the FDA
 - Has the effect of law
- **FDA Policy / Guidance**
 - Presents FDA current thinking
 - Not legally binding
- **Case Law**
 - Reported or written decisions of courts
 - Interpretation of the law (precedent)

Legal Animal Drugs

To be legally marketed, an animal drug must be the subject of:

- **approved** new animal drug application (NADA)
- **approved generic** application
[abbreviated new animal drug application]
(ANADA)
- **conditionally approved** new animal drug
- **Indexed** animal drug

Legal Human Drugs

To be legally marketed, a human drug must be the subject of:

- **approved** new drug application (NDA)
- **approved generic** application [abbreviated new drug application] (ANDA)
- **OTC (over-the-counter) Drug Monograph**

Unapproved Animal Drugs

- Animal drugs illegally marketed in the US that have not been approved by the FDA.
- Unapproved animal drugs have not been reviewed under FDA's legal review processes.

Unapproved Animal Drugs

- No pre-approval review.
- No post-approval monitoring.
- May unfairly compete against approved products.
- Reduces the availability of legal, reviewed, tested and monitored drug products.

Compounded Animal Drugs

- If compounded from **approved** drugs (as starting ingredients) **MAY** be legal under ELU (extralabel use) regulations
- If compounded from bulk ingredients **IS** considered an unapproved animal drug
- Zero tolerance for compounding from bulk ingredients for food-producing animals

Compounded Animal Drugs

- Unknown quality control / manufacturing standards.
- Unknown assurance of purity, potency, or stability.
- Potential animal safety and efficacy issues.
- Potential for unknown or unsafe residues.
- Difficult to monitor – no reporting requirements (even less than for other unapproved animal drugs).

Extralabel Drug Use

- Part 21, Code of Federal Regulations, Section 530 (21 CFR 530).
- Extralabel is the use of an **approved** drug in a manner that is not in accordance with the approved label directions.
 - e.g., indication, species, dosage level, frequency, route of administration
- Allows veterinarians to prescribe extralabel uses of certain **approved animal drugs** and **approved human drugs**.

Extralabel Drug Use

“Such use is limited to treatment modalities when the health of an animal is threatened or suffering or death may result from failure to treat.”

Extralabel Drug Use (Final Reg)

“The agency considers the clause ‘when the health of animals is threatened,’ in § 530.2, to include the concept of minimizing animal pain and suffering.”

Extralabel Drug Use

- Must be by, or on, the order of a veterinarian within the context of a “valid veterinarian-client-patient relationship”.
- Must not result in violative residues.
- Must be in conformance with all other parts of extralabel drug use regulation (21 CFR Part 530).
- Must not use drugs prohibited from extralabel use.

Extralabel Drug Use

- Approved animal drugs
- Approved human drugs
- Conditionally approved animal drugs
- Indexed animal drugs
- Medicated feeds
- VFD drugs (veterinary feed directive)
- Human OTC monographed drugs
- Unapproved animal drugs
- Drugs compounded from approved drugs
- Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- Approved human drugs
- Conditionally approved animal drugs
- Indexed animal drugs
- Medicated feeds
- VFD drugs (veterinary feed directive)
- Human OTC monographed drugs
- Unapproved animal drugs
- Drugs compounded from approved drugs
- Drugs compounded from bulk

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- Conditionally approved animal drugs
- Indexed animal drugs
- Medicated feeds
- VFD drugs (veterinary feed directive)
- Human OTC monographed drugs
- Unapproved animal drugs
- Drugs compounded from approved drugs
- Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
 - Indexed animal drugs
 - Medicated feeds
 - VFD drugs (veterinary feed directive)
 - Human OTC monographed drugs
 - Unapproved animal drugs
 - Drugs compounded from approved drugs
 - Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
 - Medicated feeds
 - VFD drugs (veterinary feed directive)
 - Human OTC monographed drugs
 - Unapproved animal drugs
 - Drugs compounded from approved drugs
 - Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
 - VFD drugs (veterinary feed directive)
 - Human OTC monographed drugs
 - Unapproved animal drugs
 - Drugs compounded from approved drugs
 - Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
- ✗ VFD drugs (veterinary feed directive)
 - Human OTC monographed drugs
 - Unapproved animal drugs
 - Drugs compounded from approved drugs
 - Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
- ✗ VFD drugs (veterinary feed directive)
- ✗ Human OTC monographed drugs
- Unapproved animal drugs
- Drugs compounded from approved drugs
- Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
- ✗ VFD drugs (veterinary feed directive)
- ✗ Human OTC monographed drugs
- ✗ Unapproved animal drugs
 - Drugs compounded from approved drugs
 - Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
- ✗ VFD drugs (veterinary feed directive)
- ✗ Human OTC monographed drugs
- ✗ Unapproved animal drugs
- ✓ Drugs compounded from approved drugs
- Drugs compounded from bulk ingredients

Extralabel Drug Use

- ✓ Approved animal drugs (Rx or OTC)
- ✓ Approved human drugs
- ✗ Conditionally approved animal drugs
- ✗ Indexed animal drugs
- ✗ Medicated feeds
- ✗ VFD drugs (veterinary feed directive)
- ✗ Human OTC monographed drugs
- ✗ Unapproved animal drugs
- ✓ Drugs compounded from approved drugs
- ✗ Drugs compounded from bulk ingredients

CPG Sec. 650.100

Animal Drugs for Euthanasia

- Euthanasia, or painless killing, is widely practiced by veterinarians to destroy unwanted pets and other animals.
- It is accomplished by a variety of methods, one of which involves drug injection.
- Pest control is not considered euthanasia, and the substances used in this practice are usually regulated by the Environmental Protection Agency.

CPG Sec. 650.100

Animal Drugs for Euthanasia

- FDA CVM considers products intended for animal euthanasia to conform to the definition of a drug under section 201(g) of the Act since they are clearly intended to **affect the function of the body** by inducing death.
- Moreover, with one exception, namely sodium pentobarbital, FDA CVM considers all such products to be new animal drugs, within the meaning of section 201(w) which may be legally **marketed** only after approval of an NADA.
- The data must establish that the product results in a humane and painless death.

Animal Drugs Approved for Euthanasia

NADA/ANADA	Sponsor	Proprietary Name	Ingredient
<u>100-809</u>	Hoechst Roussel Vet	T-61 Euthanasia Solution	Embutramid; Tetracaine; Mebezonium
<u>119-807</u>	Intervet, Inc.	Beuthanasia-D Special	Pentobarbital; Phenytoin
<u>128-967</u>	Zoetis Inc.	Repose Euthanasia Solution	Secobarbital; Dibucaine
<u>141-245</u>	Bayer HealthCare LLC, Animal Health Division	Tributame™ Euthanasia Solution	Embutramid; Chloroquine; Lidocaine
<u>200-071</u>	Virbac AH, Inc.	Euthasol®	Pentobarbital; Phenytoin
<u>200-280</u>	Med-Pharmex, Inc.	Euthanasia-III Solution	Pentobarbital; Phenytoin

Animal Drugs Approved for Euthanasia

NADA/ANADA	Sponsor	Proprietary Name	Ingredient
<u>100-809</u>	Hoechst Roussel Vet	T-61 Euthanasia Solution	Embutramid ; Tetracaine; Mebezonium
<u>119-807</u>	Intervet, Inc.	Beuthanasia-D Special	Pentobarbital; Phenytoin
<u>128-967</u>	Zoetis Inc.	Repose Euthanasia Solution	Secobarbital; Dibucaine
<u>141-245</u>	Bayer HealthCare LLC, Animal Health Division	Tributame™ Euthanasia Solution	Embutramid; Chloroquine; Lidocaine
<u>200-071</u>	Virbac AH, Inc.	Euthasol®	Pentobarbital; Phenytoin
<u>200-280</u>	Med-Pharmex, Inc.	Euthanasia-III Solution	Pentobarbital; Phenytoin

Animal Drugs Unapproved for Euthanasia

NADA/ANADA	Sponsor	Proprietary Name	Ingredient
		Fatal-Plus	Pentobarbital
		Sleepaway	Pentobarbital
		Socumb	Pentobarbital
		Pentasol	Pentobarbital
		Pentobarbital	Pentobarbital

Medical Gases

Medical gases are certified as drugs only for the following indications:

- **Oxygen:** for treatment or prevention of hypoxemia or hypoxia.
- **Nitrogen:** for use in hypoxic challenge testing.
- **Nitrous oxide:** for analgesia.
- **Carbon dioxide:** for use in extracorporeal membrane oxygenation therapy or respiratory stimulation.
- **Helium:** for treatment of upper airway obstruction or increased airway resistance.
- **Medical air:** to reduce the risk of hyperoxia.
- **Carbon monoxide:** for use in lung diffusion.

Enforcement Discretion

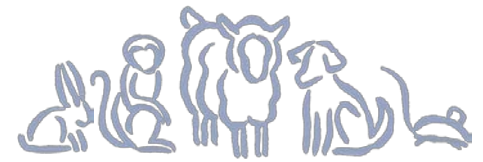
OLAW



Updated Definition, June 4, 2015:

A pharmaceutical-grade substance is any active or inactive drug, biologic, reagent, etc., manufactured under Good Manufacturing Practices (GMP) which is approved, conditionally approved, or indexed by the FDA or for which a chemical purity standard has been written or established by a recognized compendia [e.g., US Pharmacopeia and National Formulary (USP-NF), British Pharmacopeia, (BP)].

OLAW



- OLAW and USDA agree that pharmaceutical-grade substances, when available, must be used to avoid toxicity or side effects that may threaten the health and welfare of vertebrate animals and / or interfere with the interpretation of research results.
- It is frequently necessary to use investigational substances, veterinarian- or pharmacy-compounded drugs, and / or Schedule I controlled substances to meet scientific and research goals. The IACUC is responsible for evaluating the potential adverse consequences of such substances when used for research.

OLAW



- Cost savings alone are not an adequate justification for the use of non-pharmaceutical-grade substances in animals.
- Unavailability or shortages of pharmaceutical-grade substances may lead to cost increases. The IACUC may determine that this justifies the use of a non-pharmaceutical-grade substitution.

OLAW FAQ F4



Guide for the Care and Use of Laboratory Animals



- Pharmaceutical-grade substances should be used when available.
- The use of non-pharmaceutical-grade substances should be described and justified in the animal use protocol and be approved by the IACUC.
- Examples:
 - when necessary to meet the scientific goals of a project.
 - when a pharmaceutical-grade product is unavailable.



Policy 3 - Veterinary Care



Pharmaceutical-grade substances (PGS) are expected to be used in USDA regulated species when available, even in acute procedures.

- **This includes but is not limited to:**
 - compounds, medications, drugs, vehicles, and diluents.
- **APHIS recognizes that some substances are only available as non-pharmaceutical-grade (NPG):**
 - test articles, novel compounds, and those from compounding.
- **IACUC approval is required for the use of a NPG substance.**
 - Cost savings alone is not sufficient justification.
 - Approval may be given if the shortage or unavailability of the PGS has resulted in cost increases.

- NPG use - IACUC approved
- Clinical use:
 - Drugs for clinical treatment of animals (including analgesics and anesthetics)
 - Whenever possible, pharmaceutical-grade drugs must be used
- Research use:
 - Compounds used to accomplish scientific aims of the study
 - Pharmaceutical-grade is preferred, but when NPG's used:
 - Scientific justification
 - Consideration of: purity, grade stability, pH, osmolality, etc.

Case Study

May a PI use a commercial euthanasia solution at 50mg/kg, a dose less than recommended for euthanasia, for a terminal procedure in rabbits or must he use a pharmaceutical-grade anesthetic formulation of sodium pentobarbital?

- As part of IACUC approved research, rabbits undergo a terminal Langendorff procedure during which the rabbit is under deep anesthesia and a cardiectomy (heart removal) is performed.
- Once the animal is under deep anesthesia, the procedure is completed in under a minute. No other procedures are performed until after the heart is removed and the animal is dead.
- No other anesthetic /euthanasia agent is acceptable as it will compromise cardiac physiology needed for the studies that will be conducted on the myocardium after the heart is removed.

Does this fall into the category of:

- **a terminal surgery (euthanasia solution is not permitted); or**
- **a two-step euthanasia (euthanasia solution may be used, but at a lower dose since the adjunctive cardiectomy confirms death)?**

OLAW and USDA Position

Euthanasia solution is to be used only during humane termination of an animal's life. Any other use is contrary to the label instructions and considered extralabel use.

OLAW and USDA have stated that euthanasia solution is **NOT** to be used as an anesthetic for survival or non-survival procedures.

OLAW and USDA have stated that a euthanasia procedure may be performed in which an animal is given euthanasia solution after which an irreversible procedure such as a terminal perfusion, exsanguination, or tissue harvest is immediately conducted to ensure death.

Under these circumstances the euthanasia solution is being used as intended with the final outcome being death.

Any delay in the terminal steps to conduct other potentially painful procedures would change the euthanasia to non-survival surgery and be an unacceptable use of the euthanasia solution.

Question 1

The use of saturated KCl is an acceptable method of euthanasia according to the AVMA Panel on Euthanasia.

A pharmaceutical-grade saturated KCl preparation is not commercially available. What is the position of each of the organizations speaking today on the use of KCl from a chemical supply company for euthanasia while the animal is under a general plane of anesthesia?

Question 2

Dr. Bataller listed 6 FDA-approved products with pentobarbital as the active ingredient, only 3 of which are available.

How do these products differ from Fatal-Plus which is a commercial product but not FDA approved?

Summary

- FDA regulations and guidance focus on safety and efficacy of commercial animal and human drug products.
- OLAW, USDA, and AAALAC focus on research animal welfare.
- Guidance by FDA, OLAW, USDA, and AAALAC overlap during both clinical and research use in lab animals.
- Legal choices provide the least risk but may not be appropriate or available in all cases.
- Use of non-pharmaceutical-grade substances should be described and justified in the animal use protocol and be approved by the IACUC. Cost savings alone is not sufficient justification.

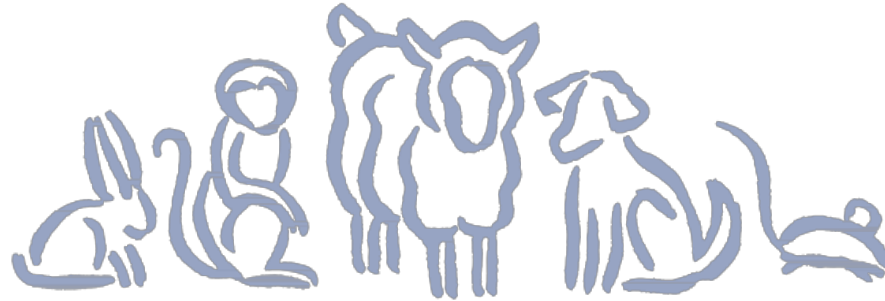
References

- Animaldrugs@FDA.gov
<http://www.accessdata.fda.gov/scripts/animaldrugsatfda/>
- FOI Summaries for Approved Drugs
<http://www.fda.gov/AnimalVeterinary/Products/ApprovedAnimalDrugProducts/FOIADrugSummaries/ucm056898.htm>
- MUMS
<http://www.fda.gov/AnimalVeterinary/DevelopmentApprovalProcess/MinorUseMinorSpecies/ucm070206.htm>
- Indexing
<http://www.fda.gov/AnimalVeterinary/DevelopmentApprovalProcess/MinorUseMinorSpecies/ucm125452.htm>
- Unapproved Drugs
<http://www.fda.gov/AnimalVeterinary/GuidanceComplianceEnforcement/ComplianceEnforcement/UnapprovedAnimalDrugs/default.htm>

References



- OLAW FAQ
http://grants.nih.gov/grants/olaw/faqs.htm#useandmgmt_4
- USDA Policy 3
http://www.aphis.usda.gov/animal_welfare/downloads/Animal%20Care%20Policy%20Manual.pdf
- AAALAC FAQ C.9.
http://aaalac.org/accreditation/faq_landing.cfm#B9
- ILAR Guide, 8th edition, 2011, page 31
http://www.nap.edu/openbook.php?record_id=12910



National Institutes of Health
Office of Laboratory Animal Welfare

OLAW Online Seminar

Topic: TBD (suggestions welcome)

September 24, 2015