

UNITED STATES DEPARTMENT OF AGRICULTURE  
ANIMAL AND PLANT HEALTH INSPECTION SERVICE

1. CERTIFICATE NUMBER: 14-R-0101  
CUSTOMER NUMBER: 147

FORM APPROVED  
OMB NO. 0579-0036

**ANNUAL REPORT OF RESEARCH FACILITY**  
(TYPE OR PRINT)

Toxikon Corporation  
15 Wiggins Ave  
Bedford, MA 01730

Telephone: (781)-275-3330

**3. REPORTING FACILITY** ( List all locations where animals were housed or used in actual research, testing, or experimentation, or held for these purposes. Attach additional sheets if necessary )

FACILITY LOCATIONS ( Sites ) - See Attached Listing

**REPORT OF ANIMALS USED BY OR UNDER CONTROL OF RESEARCH FACILITY** ( Attach additional sheets if necessary or use APHIS Form 7023A )

A. Animals Covered By The Animal Welfare Regulations	B. Number of animals being bred, conditioned, or held for use in teaching, testing, experiments, research, or surgery but not yet used for such purposes.	C. Number of animals upon which teaching, research, experiments, or tests were conducted involving no pain, distress, or use of pain-relieving drugs.	D. Number of animals upon which experiments, teaching, research, surgery, or tests were conducted involving accompanying pain or distress to the animals and for which appropriate anesthetic, analgesic, or tranquilizing drugs were used.	E. Number of animals upon which teaching, experiments, research, surgery or tests were conducted involving accompanying pain or distress to the animals and for which the use of appropriate anesthetic, analgesic, or tranquilizing drugs would have adversely affected the procedures, results, or interpretation of the teaching, research, experiments, surgery, or tests. ( An explanation of the procedures producing pain or distress in these animals and the reasons such drugs were not used must be attached to this report ).	F. TOTAL NUMBER OF ANIMALS ( COLUMNS C + D + E )
4. Dogs	0	48	100	2	150
5. Cats	0	0	0	0	0
6. Guinea Pigs	0	13967	0	21	13988
7. Hamsters	0	99	0	0	99
8. Rabbits	0	846	1208	5	2059
9. Non-human Primates	0	0	0	0	0
10. Sheep	0	0	0	0	0
11. Pigs	0	31	36	1	1
12. Other Farm Animals	0	0	0	0	0
13. Other Animals	0	0	0	0	0

**ASSURANCE STATEMENTS**

- 1) Professionally acceptable standards governing the care, treatment, and use of animals, including appropriate use of anesthetic, analgesic, and tranquilizing drugs, prior to, during, and following actual research, teaching, testing, surgery, or experimentation were followed by this research facility.
- 2) Each principal investigator has considered alternatives to painful procedures.
- 3) This facility is adhering to the standards and regulations under the Act, and it has required that exceptions to the standards and regulations be specified and explained by the principal investigator and approved by the Institutional Animal Care and Use Committee (IACUC). A summary of all such exceptions is attached to this annual report, in addition to identifying the IACUC-approved exceptions, this summary in brief explanation of the exceptions, as well as the species and number of animals affected.
- 4) The attending veterinarian for this research facility has appropriate authority to ensure the provision of adequate veterinary care and to oversee the adequacy of other aspects of animal care and use.

**CERTIFICATION BY HEADQUARTERS RESEARCH FACILITY OFFICIAL**  
( Chief Executive Officer or Legally Responsible Institutional Official )

NAME & TITLE OF C.E.O. OR INSTITUTIONAL OFFICIAL ( Type or Print )  
DATE SIGNED  
1/30/03

*C. K. ...*

## Optional Column E Explanation Form

This form is intended as an aid to completing the Column E explanation. It is not an official form and its use is voluntary. Names, addresses, protocols, veterinary care programs, and the like, are not required as part of an explanation. A Column E explanation must be written so as to be understood by laypersons as well as scientists.

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**1. Registration Number:** 14-R-101

**2. Number of animals used in the study (s).** 2 Dogs, 21 Guinea Pigs, 5 Rabbits, 1 Pig

**3. Specie (common name) of animals used in this study (s).**

Dogs, Guinea Pigs, Rabbits, Pigs

**4. Explain the procedure producing pain and/or distress.**

Exposure to a test article (drug /chemical compounds).

**5. Provide scientific justification why pain and/or distress could not be relieved. State methods or means used to determine that pain and/or distress relief would interfere with test results.** (For Federally mandated testing, see question 6 below)

Studies conducted at Toxikon are performed for sponsors to obtain toxicity information on experimental materials, drugs or chemicals, or to ensure the safety of a new lot of material. Regulatory guidelines do not permit the use of analgesic or anesthetics during toxicity determination studies. However, Toxikon does employ a step approach, exposing one or two animals at a time, thus minimizing the total number of animals needed. Toxikon's IACUC approves and monitors all animal use protocols.

**6. What, if any, federal regulations require this procedure? Cite the agency, the Code of Federal Regulations (CFR) title number and the specific section number (e.g., APHIS, 9 CFR 113.102):**

Based upon the following standards:

Dog:

Organization for Economic Co-operation and Development (OECD), Guidelines for Testing of Chemicals, "Repeated Dose Oral Toxicity - Rodent: 28-day or 14-day Study," Section 407 (adopted 27 July 1995).

ICH Harmonized Tripartite Guideline. Note for Guidance on Toxicokinetics: The Assessment of Systemic Exposure in Toxicity Studies, S3A, 1994. FDA: Published in the Federal Register, Vol. 60, No. 40, March 1, 1995, pages 11264-11268.

ICH Harmonized Tripartite Guideline. Dose Selection for Carcinogenicity Studies of Pharmaceuticals, 1997 (revised). FDA: First published in the Federal Register, Vol. 60, No.40, March 1, 1995, pages 11278- 11281, Revision published in the FR, Vol. 62, No. 233, December 4, 1997, page 64260.

Guinea Pig:

Organization for Economic Co-operation and Development (OECD), Guidelines for Testing of Chemicals, " Skin Sensitization" section 406, July 1992.

The Office of Prevention, Pesticides and Toxic Substances (OPPTS), United States Environmental Protection Agency, Health Effects Test Guidelines. OPPTS 870.2600, Skin Sensitization, March, 2003

United States Pharmacopoeia 26, National Formulary 21, 2003, <88> Biological Reactivity Tests, *In Vivo*.

21 CFR, Part 610.11, 2002.

Rabbit:

Organization for Economic Co-operation and Development (OECD), Guideline for Testing of Chemicals, Section 402, 1987.

Procedure and facilities complied with the requirements of Commission Directive 86/609/EEC concerning the protection of animals used for experimental and other scientific purposes. National legislation, harmonising with this Directive, is defined in Decreto Legislativo No. 116 of 27th January 1992.

Swine:

Toxicokinetics: Guidance on the Assessment of Systemic Exposure in Toxicity Studies, International Conference on Harmonization/FDA/CDER/CBER.

Specifications for the Conduct of Studies to Evaluate the Toxic and Carcinogenic Potential of Chemical, Biological and Physical Agents for the National Toxicology Program, 8/1992; Revised 10/1996.

CRC Handbook of Toxicology (eds: Derelanko MJ, Hollinger MA) CRC Press, 1995.

Ninth Addendum to the Organization for Economic Cooperation and Development Guidelines for the Testing of Chemicals, Neurotoxicity Study in Rodents, Section 424, adopted July 21, 1997.

Handbook of Pre-clinical Continuous Intravenous Infusion (eds. Healing, G., & Smith, D.) Taylor and Francis, 2000.

	<b>Date</b> 11/30/03
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DEC 02 2003

This report is required by law (7 USC 2143). Failure to result in an order to cease and desist and to be subject to penalties as provided for in Section 2150.

See according to the regulations can addi side for information.

See side for information.

Interagency Report Control No 0180-DOA-AN

UNITED STATES DEPARTMENT OF AGRICULTURE  
ANIMAL AND PLANT HEALTH INSPECTION SERVICE

1. REGISTRATION NO.

14-V-0001

657 NO  
652

FORM APPROVED  
OMB NO. 0579-0036

**ANNUAL REPORT OF RESEARCH FACILITY  
(TYPE OR PRINT)**

2. HEADQUARTERS RESEARCH FACILITY (Name and Address, as registered with USDA, include Zip Code)

Dept. of Veterans Affairs Central Office  
810 Vermont Avenue NW  
Washington DC 20420

3. REPORTING FACILITY (List all locations where animals were housed or used in actual research, testing, teaching, or experimentation, or held for these purposes. Attach additional sheets if necessary.)

**FACILITY LOCATIONS (Sites)**

518 VA Medical Center  
200 Springs Rd.  
Bedford, MA 01730

**REPORT OF ANIMALS USED BY OR UNDER CONTROL OF RESEARCH FACILITY (Attach additional sheets if necessary or use APHIS FORM 7023A)**

A. Animals Covered By The Animal Welfare Regulations	B. Number of animals being bred, conditioned, or held for use in teaching, testing, experiments, research, or surgery but not yet used for such purposes.	C. Number of animals upon which teaching, research, experiments, or tests were conducted involving no pain, distress, or use of pain-relieving drugs.	D. Number of animals upon which experiments, teaching, research, surgery, or tests were conducted involving accompanying pain or distress to the animals and for which appropriate anesthetic, analgesic, or tranquilizing drugs were used.	E. Number of animals upon which teaching, experiments, research, surgery or tests were conducted involving accompanying pain or distress to the animals and for which the use of appropriate anesthetic, analgesic, or tranquilizing drugs would have adversely affected the procedures, results, or interpretation of the teaching, research, experiments, surgery, or tests. (An explanation of the procedures producing pain or distress in these animals and the reasons such drugs were not used must be attached to this report).	F. TOTAL NO. OF ANIMALS (Cols. C + D + E)
4. Dogs	0				
5. Cats	0				
6. Guinea Pigs	0	0	14	0	14
7. Hamsters	0				
8. Rabbits	0				
9. Non-human Primates	0				
10. Sheep	0				
11. Pigs	0				
12. Other Farm Animals	0				
13. Other Animals	0				
14. Mice	1112	5356	68	0	6424

**ASSURANCE STATEMENTS**

- 1) Professionally acceptable standards governing the care, treatment, and use of animals, including appropriate use of anesthetic, analgesic, and tranquilizing drugs, prior to, during, and following actual research, teaching, testing, surgery, or experimentation were followed by this research facility.
- 2) Each principal investigator has considered alternatives to painful procedures.
- 3) This facility is adhering to the standards and regulations under the AIA, and it has required that exceptions to the standards and regulations be specified and explained by the principal investigator and approved by the Institutional Animal Care and Use Committee (IACUC). A summary of all such exceptions is attached to this annual report. In addition to identifying the IACUC-approved exceptions, this summary includes a brief explanation of the exceptions, as well as the species and number of animals affected.
- 4) The attending veterinarian for this research facility has appropriate authority to ensure the provision of adequate veterinary care and to oversee the adequacy of other aspects of animal care and use.

**CERTIFICATION BY HEADQUARTERS RESEARCH FACILITY OFFICIAL  
(Chief Executive Officer or Legally Responsible Institutional Official)**

I certify that the above is true.

SIGNATURE OF C.E.O. OR INSTITUTIONAL OFFICIAL

DATE SIGNED

11/3/03

APHIS FORM 7023  
(AUG 91)

(Replaces VS FORM 18-23 (OCT 88), which is obsolete)