



DEPARTMENT OF HEALTH & HUMAN SERVICES

PUBLIC HEALTH SERVICE
NATIONAL INSTITUTES OF HEALTH

FOR US POSTAL SERVICE DELIVERY:

Office of Laboratory Animal Welfare
Rockledge One, Suite 360
6705 Rockledge Drive - MSC 7982
Bethesda, Maryland 20892-7982
Home Page: <http://grants.nih.gov/grants/olaw/olaw.htm>

FOR EXPRESS MAIL:

Office of Laboratory Animal Welfare
Rockledge One, Suite 360
6705 Rockledge Drive
Bethesda, Maryland 20817
Telephone: (301) 496-7163
Facsimile: (301) 402-2803

May 10, 2017

Re: Animal Welfare Assurance
#A3331-01 (OLAW Case M)

Mr. Richard Best
Director of Compliance
North Carolina State University
103 Holladay Hall, Box 7003
Raleigh, NC 27695

Dear Mr. Best,

The Office of Laboratory Animal Welfare (OLAW) acknowledges receipt of your May 9, 2017 letter reporting an adverse event involving research mice at North Carolina State University. According to the information provided, OLAW understands that 10% of a cohort of mice were found dead following an approved procedure involving instillation of cancer cells in polylysine into the bladder. It was determined that a different source for polylysine was used and the solution was inadvertently prepared ten times more concentrated than described in the protocol.

The corrective actions consisted of the Principal Investigator stopping the conduct of additional procedures until the problem was solved. The PI implemented a standardized formula to avoid overdoses and subsequent procedures no longer had high mortality.

Based on its assessment of this explanation, OLAW understands that measures have been implemented to correct and prevent recurrence of this problem. OLAW concurs with the actions taken by the institution to comply with the PHS Policy on Humane Care and Use of Laboratory Animals.

Sincerely,

Axel Wolff, M.S., D.V.M.
Director
Division of Compliance Oversight

cc: IACUC Chair



Office of Research, Innovation
and Economic Development
Sponsored Programs and
Regulatory Compliance Services
research.ncsu.edu/sparcs/

A 3331-M
Campus Box 7514
2701 Sullivan Dr. Suite 240
Raleigh, NC 27695-7514
P: 919-515-2444, F: 919-515-7721
E: sps@ncsu.edu

Dr. Axel Wolff
Director, Division of Compliance Oversight
Office of Laboratory Animal Welfare
National Institutes of Health
Rockledge 1, Suite 360, MSC 7982
6705 Rockledge Drive
Bethesda, MD 20892-7982

RE: Adverse event - Animal Welfare Assurance # D16-00214

Principal Investigator: David A. Zaharoff

NIH Grant #7R01CA172631-04

NCSU IACUC protocol 16-256-B

Dear Dr. Wolff,

This letter serves to report a recent adverse event for the investigator identified above.

On March 13, 2017 mice were anesthetized in preparation for bladder instillation of cancer cells in polylysine. After the procedure, the mice recovered from general anesthesia without incident. On March 14th, approximately 10% of those mice were found dead, without visible signs of disease. Dr. Gabriel McKeon, University Attending Veterinarian, received information from animal care staff regarding the found dead animals on the afternoon of the 14th. He inspected the cages in question, and communicated with the Principal Investigator in order to determine if this death rate fell within the expected range of the procedure. Dr. Zaharoff indicated that it was unusual, and that he would contact the postdoc who had performed the procedure to get more information. He suspected that the polylysine was the cause of mortality. Dr. Zaharoff voluntarily ceased further procedures until the cause of the mortality was determined. Dr. Zaharoff discovered that the postdoc was unable to obtain polylysine from the usual source and purchased it from an alternate vendor. The alternate vendor's formulation was different, and the postdoc mistakenly made a solution of polylysine that was 10-fold more concentrated than the dosage prescribed in the protocol. Dr. Zaharoff reviewed the product preparation with the postdoc, instituting a standardized formula in order to avoid overdose of polylysine. The lab personnel conducted subsequent procedures with the appropriate concentration, and without high mortality.

Dr. McKeon considered the matter appropriately corrected, and that there is no further risk of sudden death from improper polylysine formulation for the mice undergoing the procedure. He reported the event and the corrective follow up to the IACUC Chair, Dr. William Flowers and to me on April 13 2017. He also included it in his monthly report at a convened meeting of the IACUC on April 20th.

I am in agreement with Dr. McKeon that Dr. Zaharoff took appropriate action and that the problem is resolved. In compliance with our Assurance, we are submitting this report of the adverse event and its resolution.

Dr. Axel Wolff
Page 2

If there are questions regarding this report, please contact me, Dr. Gabriel McKeon (gpmckeon@ncsu.edu,
[Redacted Phone Number] or our IACUC Administrator, [Redacted Secondary Individual], [Redacted Secondary Individual]@ncsu.edu or [Redacted Phone Number].

Best regards,



Richard Best, MPA
Director for Research Compliance
Institutional Official

Cc: Gabriel McKeon
[Redacted Secondary Individual]
David Zaharoff

Wolff, Axel (NIH/OD) [E]

From: Wolff, Axel (NIH/OD) [E]
Sent: Wednesday, May 10, 2017 7:32 AM
To: econdary Individu
Subject: RE: Adverse Event Report - Assurance #D16-00214 - North Carolina State University

Thank you for this report, econdary Individu. I will respond shortly. In future, please send all final reports to olawdco@od.nih.gov.

Axel Wolff, M.S., D.V.M.
Director, Division of Compliance Oversight OLAW

-----Original Message-----

From: econdary Individu [mailto:ary In@ncsu.edu]
Sent: Tuesday, May 09, 2017 3:06 PM
To: Wolff, Axel (NIH/OD) [E] <WolffA@OD.NIH.GOV>
Cc: Richard Best <rebest@ncsu.edu>; Gabriel McKeon <gpmckeon@ncsu.edu>; Secondary individual @ncsu.edu; David Zaharoff <dazaharo@ncsu.edu>
Subject: Adverse Event Report - Assurance #D16-00214 - North Carolina State University

Dear Dr. Wolff,

Attached please find an adverse event report submitted by our Institutional Official, Mr. Richard Best.

Best regards,

econdary Individu

Certified Professional IACUC Administrator Regulatory Compliance Administrator Institutional Animal Care and Use Committee Sponsored Programs and Regulatory Compliance North Carolina State University Campus Box 7514 Raleigh, NC 27695-7514

Phone: Phone Number

Fax: 919.515.7721

Email: econdary Individu@ncsu.edu

Web: www.ncsu.edu/iacuc

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