Annual Report of Research Facility
Column E Explanation

1. Registration Number: 48-R-0111 (Xenometrics LLC)

2. Number of animals reported in Column E: One

3. Species (common name) of animals used in this study: Dog

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

   The affected animal was included in a toxicity study in which allergic-type reactions were possible. If such reactions occurred, the animal(s) was to be treated and/or euthanized. The one affected animal exhibited rapid onset of severe allergic-type clinical signs and died despite rapid administration of treatment.

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 Item 6 below).

   Not applicable, as the protocol included a contingency plan for the treatment and/or euthanasia if pain or distress occurred.

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).

   The study was conducted per 21 CFR 312.23 and ICH Guidance M3 (R2) Nonclinical Safety Studies for the Conduct of Human Clinical Trials and Marketing Authorization for Pharmaceuticals.

Annual Report of Research Facility
Column E Explanation

1. Registration Number: 48-R-0111 (Xenometrics LLC)

2. Number of animals reported in Column E: Ten

3. Species (common name) of animals used in this study: Dog

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

   The affected animals were included in a toxicity study in which pain and/or distress were not expected, however the contingency plan included provision for treatment and/or euthanasia if unexpected reactions resulted in pain and/or distress. The affected animals exhibited no signs of pain or distress prior to being found dead.

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 Item 6 below)

   Not applicable, as the protocol included a contingency plan for the treatment and/or euthanasia of unexpected pain or distress.

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)

   The study was conducted per FDA/CDER and ICH Guidance ICH-M3 M3(R2) Nonclinical Safety Studies for the Conduct of Human Clinical Trials and Marketing Authorization for Pharmaceuticals and ICH S9 Nonclinical Evaluation for Anticancer Pharmaceuticals.