

A. Column E Explanation (all studies listed below were approved by the IACUC at their respective site):

One guinea pig developed acute complications while on a toxicology study and died before euthanasia could be accomplished.

Ten cotton rats developed acute complications while on a pharmacokinetic study and died before euthanasia could be accomplished.

Two rabbits involved developed acute complications while on efficacy studies and died before euthanasia could be accomplished.

Three dog developed acute complications while on a toxicology study and died before euthanasia could be accomplished. One dog developed signs of weight loss and inappetence while on a toxicity study. A decision was made to euthanize the dog.

The activities and procedures of conduct for the above-referenced studies were reviewed and approved by the Institutional Animal Care and Use Committee (IACUC). Animals used on product release testing and toxicology studies were required by federal regulations for submission regarding safety and toxicity testing. The regulations that require these procedures include 21 CFR 58 (FDA) and 21 CFR 610.11 (FDA).