

Appendix 1
USDA ANNUAL REPORT (2013-2014)
Registration #: 22-R-0138

Species	Cat E usage	Protocol Procedure/Justification
Dogs	31	<p>Procedure: Single and Repeat Oral Dosing of test compounds to dogs for the conduct of Pharmacokinetic/Toxicology studies.</p> <p><i>After being orally dosed with test compound these dogs experienced two or more episodes of vomiting and diarrhea. One dog underwent tonic convulsions and died immediately before it could be euthanized.</i></p> <p>Pain and/or distress could not be relieved with tranquilizing drugs, anti-emetics or anti-diarrheals as they will interfere with the goals/aims of the intended studies, primarily study of the physiologic and pathologic processes produced by toxicity of dosed investigational agents.</p> <p>These studies are performed in accord with "Harmonization of Guidelines for Toxicity testing of Pharmaceuticals by 1992," regulatory Toxicology and Pharmacology, Vol 12, 179-211 (1990).</p> <p><i>(In all cases, this met the IACUC approved study endpoint criteria)</i></p>
Rabbits	16	<p>Procedure: IV & IM dosing of rabbits to conduct Bacteremia, Immunogenicity and Safety studies.</p> <p><i>After being intravenously inoculated with bacteria or receiving a vaccine intramuscularly rabbits developed a fever lasting greater than 2 hours.</i></p> <p>Pain and/or distress could not be relieved with the administration of analgesics or antipyretics as this will diminish the pathogenicity of the bacteria being evaluated. In addition they will interfere with vaccine efficacy, immunogenicity and pyrogenicity.</p> <p><i>(In all cases, this met the IACUC approved study endpoint criteria)</i></p>