

51F0006

12/21/99

Protocol 6

1. A total of 24 column "E" non-human primates were utilized in this study.
2. Painful procedure: All monkeys receive a convulsive dose of the nerve agent soman, which is though to cause some pain and/or distress due to the intense physical activity caused by the seizure.
3. Justification: The pain may be relieved to some extent by administration of one of the test drugs that successfully terminates the seizure. Anesthetics and analgesics are known to have profound effects on brain function that can interact with the drugs of interest and/or the toxicity of the nerve agent and thus complicate interpretation of the results. This is especially so since the purpose of the study is to model the pharmacological intervention a human nerve agent casualty would receive. Since the goal of the study is to evaluate the ability of novel anticonvulsant drugs to treat nerve agent-induced seizures, subjecting the animals to nerve agent intoxication that elicits seizures is essential.
4. No federal regulations mandate this procedure.

DEC 21 1999

51F0006

12/21/99

Protocol 7

1. A total of 12 column "E" non-human primates were utilized in this study.
2. Painful procedure: Non-human primates may experience pain or distress due to exposure to cyanide.
3. Justification: Untreated cyanide administration induces rapid unconsciousness and death. Several authors have argued that this is painless and does not induce stress, and may actually produce analgesia (Tadic, 1992), while others have suggested that it does induce pain or distress (Potter, 1950). Recovery from cyanide intoxication may induce stress. The investigators believe that pretreatment with WR242,511 will prevent death. The efficacy of WR242,511 cannot be determined unless these animals are challenged with a potentially fatal dose of cyanide. However, it is not certain that these animals will not experience pain or distress as a result of the non-fatal cyanide exposure. The investigators have no other choice but to expect that cyanide administration may be painful/distressful.
4. No federal regulations mandate this procedure.

DEC 21 1999