

Column E Explanation

This form is intended as an aid to completing the Column E explanation. It is not an official form and its use is voluntary. Names, addresses, protocols, veterinary care programs, and the like, are not required as part of an explanation. A Column E explanation must be written so as to be understood by lay persons as well as scientists.

1. Registration Number: **52-R-0124**
2. Number of animals used in this study: **3**
3. Species (common name) of animals used in the study: **Rhesus macaques**
4. Explain the procedure producing pain and/or distress.

Dependence is produced by chronically treating animals with an opioid or allowing animals to self-administer opioids and withdrawal is produced by briefly terminating opioid treatment or self-administration access conditions.

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 the test results. (For federally mandated testing, see Item 6 below).

The distress produced by this procedure is the scientific end-point being measured. The use of anesthetics, sedatives, and analgesics would interfere with scientific dependent measure, as these compound classes would obscure our evaluation of the opioid withdrawal signs. In fact, one goal of our studies is to test compounds for their ability to reduce overt withdrawal signs and withdrawal-associated changes in behavior.

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)

Not Applicable

21 NOV 2019

~~26 NOV 2019~~

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1. Registration Number: **52-R-0124**
2. Number of animals used in this study: **5**
3. Species (common name) of animals used in the study: **Rhesus macaques**
4. Explain the procedure producing pain and/or distress.

Withdrawal from novel opiate formulations (novel drug or known opiate combined with adjunctive therapy) in order to assess the propensity to produce physical dependence.

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 the test results. (For federally mandated testing, see Item 6 below).

The use of anesthetics, sedatives or analgesics would all interfere with evaluation of the withdrawal signs that characterize physical dependence.

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 results from our testing may be used by the FDA.

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21 NOV 2019