## Category E explanation- Hamsters

Administration of pain/distress-relieving drugs can change the clinical sign monitoring and increase chance for contamination. Syrian hamster (Mesocricetus auratus), develops only moderate signs of clinical disease and lung pathology after infection with SARS-CoV-2 (https://pubmed.ncbi.nlm.nih.gov/32408338/). Roborovski hamsters (Phodopus roborovskii) are highly susceptible COVID-19 model with consistent and fulminant clinical signs within 3 days following infection (https://pubmed.ncbi.nlm.nih.gov/33271063/). Animals will be monitored for pain and distress daily or twice a day based on the scoring. Any animal that reaches a score of 1 will be sacrificed.

## Explanation for Category E- Swine

Seven pigs were born with a CDKL5-deficiency disorder which causes seizures in humans. The effect of this mutation on pigs is unknown so these animals were classified as Category E in case they do have recurring seizures. To date, no pigs have exhibited seizures.

Fifteen pigs were used to study influenza infection. They are categorized as Category E since no pharmacological intervention was to be given. Pigs were monitored for well-being/pain/distress at least 2 times per day and scored for any pain or distress. The protocol states that once a predetermined score is reached the pig will be humanely euthanized. To date, no pigs have reached a pain/distress score to require euthanasia.