



Home Office

232058 PPL Standard Condition 18

Notification Form

Please use this form to notify the Animals in Science Regulation Unit (ASRU) of procedural-related adverse effects that have exceeded or are likely to exceed the severity limitations or controls described in the project licence. ASRU will use this information to determine whether any further action is required.

Do not use to report:

- adverse effects that are due to non-procedural issues;
- potential non-compliance (report by other means);
- issues without adverse welfare consequences.

If you are in doubt whether or not you need to fill in this form, please contact your assigned Inspector.

Please clearly title your document: **'ASRU_establishment name_PPL number_PPLh surname_date_sc18'** and send to your Inspector (e.g. using cjsm) with the subject heading 'PPL SC 18 Notification PPL [Number] – **Check Home Office's Updated Bridging Ways of Working for updated advice on email addresses and subject headings. Cjsm no longer necessary.**

Establishment Licence name	University of Cambridge
Establishment Licence number	X81BD37B1
Name of PPL holder	Darren Schofield
PPL number	PCBA9AEC1
Protocol no	1
Protocol title	Production of Potent Monoclonal Antibodies
Severity classification	Mild
1. Date of incident	19 th September 2023
2. Species	Mouse

3. Brief details of study and how the limits or constraints have been breached	Animals under this study receive five rounds of immunisation following a modified 28 day RIMMs model. This consists of four sets of subcutaneous (s.c.) injections each targeting the left and right auxiliary lymph nodes in the upper thoracic area. The animals are anaesthetised to allow accurate placement of injections. The final boost is an IP dose without anaesthetic. This was the final s.c. dose of the study. The three mice in this cage were put in an induction chamber at the same time. One at a time, each animal was transferred to a nose cone and the injections performed. Following injection, animals are placed in their home cage and recovery monitored. Mouse 8 was second to be injected. Typically, by the time the next mouse is placed in the home cage, the previous mouse has recovered. The PIL noticed this was not the case. The PIL picked up mouse 8 and noticed the eyes were darker than normal and her breathing was very slow, breathing quickly changed to agonal breathing and then stopped. As the other animals in the cage recovered normally, the PIL continued dosing the remaining animals.
4. Have you taken advice from the Named Animal Care and Welfare Officer/Named Veterinary Surgeon/others?	This animal died without recovering from anaesthetic. NACWO was informed immediately. The NACWO was present for two further dosing procedures and felt everything looked in order and was a good procedure. It was the NACWO's opinion that this was a one off reaction to the anaesthetised procedure.
5. Cause of Problem, if known	A necropsy was performed, and samples will be sent for pathological assessment. Upon opening the animal, it was noted that the lungs were very dark compared to normal healthy lungs.

6. Action that has been or will be taken <i>(For example, formulation of drug changed, supportive treatment given, study design modified). If no action needed, indicate why (for example, none required as study has been terminated and will not be repeated)</i>	Since none of the mice were affected in previous dosing timepoints nor were any other mice impacted at this time point, it would appear to be an isolated incident for this particular mouse. Discussion with the NACWO suggests that CD-1 mice (strain used in this study) can be susceptible to issues with repeat isoflurane administration. However, this is not something we have encountered on this licence previously with any mouse strain. We will continue to monitor on future studies and review whether to reduce or omit the use of anaesthesia if recurring on future studies.
7. Reported by (name)	Darren Schofield
8. Date	19 th September 2023

*****For official use*****

Inspector's comments and recommendations:

Comments and recommendations	Thank you for the PPL SC18 Report. You are compliant with the licence conditions by reporting this under PPL Standard Condition 18. The content of this SC18 has been read and noted and can now be considered CLOSED. Please do inform us of any other related incidents should they occur. Date 22/11/23
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