



Home Office

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	Ole Paulsen
PPL number	PP0499088
Protocol no	09
Protocol title	Ex vivo preparations preceded by neonatal intracranial injection
Severity classification	Moderate
1. Date of incident	Reported missing/found dead on 08/03-24 (at P1).

2. Species	Mouse
3. Brief details of study and how the limits or constraints have been breached	The pup underwent an intracranial AAV injection at P1 (including the rest of the batch the pup came from). The aim of the study is to express eGFP in a subgroup of GABAergic interneurons (somatostatin positive cells) to specifically target them for electrophysiology recordings. We are interested in understanding changes in their intrinsic properties in a mouse model of Rett syndrome (a neurological disorder mainly affecting girls). These AAV injections are done at P1, so that eGFP can be expressed by P14 at the latest, when we do the recordings. The procedure has been successful and has been done on 8 litters. Though pups are more resistant to isoflurane and take longer time to be fully anaesthetised, the procedure has worked well, and the mother has taken the pups back as long as the injections are done at P1 the earliest. The PIL holder was informed on the 8th of March, that half of the litter had gone missing/found dead at P1. AAV injections were done on three pups, of which two has been reported "missing" on the MCMS database on the 9th of March.
4. Have you taken advice from the Named Animal Care and Welfare Officer/Named Veterinary Surgeon/others?	The incident has been reported to NACWO.

5. Cause of Problem, if known	A total of 9 pups were born on the 7th of March, and half of the batch was found missing on the 8th of March prior to the procedure. Following the procedure on the 8th of March, it was reported that two of the mice that underwent intracranial AAV injections were found 'missing' (9th of March). A post-mortem was not performed, as the pups were not found. It is likely due to the mother being stressed and hence unable to take care of her pups. To reduce stress/disturbance in future litters, electro gel can be provided, as this can help with milk production.
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>	No deviation from the study plan. Next preventive step is to re-assess overall wellbeing of litter prior to procedure and reduce any stress in the mother.
7. Reported by (name)	Abinayah John
8. Date	09/03/2024

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

Inspector's comments and recommendations:

Comments and recommendations	Thank you for submitting this notification and thereby complying with standard condition 18 of your project licence. The content of this SC18 report has been reviewed and this matter is now CLOSED.
Inspector name	Dr Noelia Lopez-Salesansky

Date	02/04/2024
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