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]

Establishment Licence name	University of Cambridge
Establishment Licence number	X81BD37B1
Name of PPL holder	Maria P. Alcolea
PPL number	PP7037913
Protocol no	7
Protocol title	Changes in epithelial cell behaviour in response to tissue perturbations
Severity classification	Moderate

1. Date of incident	Sunday 16th January 2022
2. Species	Mouse (MABX47.1e)

3. Brief details of study and how the limits or constraints have been breached	This study plan (SP125688, MA_GS_052_06) includes in total 16 animals that are receiving low doses (40mg/L) of DEN carcinogen via drinking water for 2 months. Towards the end of the DEN treatment Cre transgene expression is activated by Tamoxifen in epithelial cells (using keratinocyte specific promoter) to conditionally knock out Sox9 expression in recombined cells. On the 10th of January the MABX47.1e animal has started DEN carcinogen treatment and received 3 low doses of it (out of 28) via drinking water prior to death. The animal weight went down by 3.8% from baseline, four days after the first DEN bottle was provided, which did not cause any health concerns as the animal was healthy and the weight loss did not breach 10%. If animal reaches 10% weight loss the technicians notify user for health concern. Animals are not expected to show any adverse clinical signs as a result of DEN treatment, except for weight loss and incidental tumours such as lipomas as they age. Sporadic liver damage was occasionally observed resulting in early animal collection and is a potential reason for this animal being found dead. Animals receiving DEN or after DEN treatment are monitored by weighing daily for the first week, weekly for the next month and then once every two months. Animals are also checked every day for any clinical signs or abnormal behaviour as done by the facility technicians on a daily basis. Any concerning animals go back to weighing every other day or daily, if necessary, until the weight is stable again. If the concerning signs do not resolve within 24 hours they are culled. In the event of worsening clinical signs animals will be immediately culled. If vital processes such as eating, defecating and breathing are compromised, as identified in regular inspections, animals are culled immediately. Any animal showing weight loss of 10% will be moved to a mash diet and any weight loss of 15% or more for 2 consecutive days are to be culled immediately. Animals receiving DEN or after DEN treatment are monitored by weighing daily for the first

	week, weekly for the next month and then once every two months. Animals are also checked every day for any clinical signs or abnormal behaviour.
4. Have you taken advice from the Named Animal Care and Welfare Officer/Named Veterinary Surgeon/others?	NACWO and NVS have been notified.
5. Cause of Problem, if known	Upon dissection liver and lungs showed haemorrhage, likely to be liver failure due to adverse reaction to DEN carcinogen.
6. Action that has been or will be taken	The animal weight went down by 3.8% from baseline, four days after the first DEN bottle was provided, which did not cause any health concerns as the animal was healthy and the weight loss did not breach 10%. In addition none of the other 15 animals receiving the same batch of drug did not show any signs of concerned health. Therefore, the animal likely died due to adverse reaction to DEN.
7. Reported by (name)	Greta Skrupskelyte, animal found by Greta Ulyte
8. Date	16/01/2022

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

Inspector's comments and recommendations:

Comments and recommendations	
Inspector name	
Date	