



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 The Anne McLaren Building
Establishment Licence number	X81BD37B1
Name of PPL holder	Brian Huntly
PPL number	PP3042348
Protocol no	07
Protocol title	Study of the impact of cell extrinsic factors on normal, pre-malignant and malignant haematopoiesis through transplantation (Mice)

Severity classification	Moderate
1. Date of incident	29/09/2022
2. Species	Mouse
3. Brief details of study and how the limits or constraints have been breached	This study involves the generation of haematopoietic chimaeras in immunodeficient NSG mice by transplantation of highly purified human haematopoietic stem cells. One mouse was found dead by AMB staff (Fallon Miller) without having shown any previous signs of disease or ill health. Mice on this study are visually inspected by AMB staff and members of our team on a regular basis. After irradiation, mice are weighed twice weekly by members of AMB staff for two weeks to monitor any weight loss following irradiation. No significant weight loss had been recorded for this animal post transplantation. This animal had received a sub-lethal dose of irradiation on the 7th of June followed by intravenous injection of normal human haematopoietic stem cells on the 8th of June 2021. The time of death (16 weeks after irradiation) excludes any complications from irradiation, which normally arise within 3 weeks. The use of highly purified human haematopoietic stem cells eliminates the risk of graft vs host disease as no T cells are transplanted in the animals and no reactive T cells are produced by these transplanted stem cells. The proportion of mice that have been found dead after performing transplantation of human haematopoietic stem cells in the hands of our group is very low (7/1292 mice and therefore approximately 0.5%).
4. Have you taken advice from the Named Animal Care and Welfare Officer/Named Veterinary Surgeon/others?	Yes, we have discussed this animal with the staff in charge of the holding room. The animal technician who found the body reported no obvious signs for the cause of death. No clinical/poor health signs had ever been reported for this animal.

5. Cause of Problem, if known	The cause of death remains unknown at the time we are submitting this Condition 18. We request an autopsy but the mouse has been cannibalised.
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>	We will continue to monitor these animals (along with AMB staff) at regular intervals and will include regular weighing. Previous advice given by an Inspector in August 2022 has been to amend the PPL to include a <1% death rate in transplanted mice. This advice has been taken on board and will be included in the list of changes for our prospective PPL amendment.
7. Reported by (name)	Elisa Laurenti
8. Date	29/09/2022

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

Inspector's comments and recommendations:

Comments and recommendations	
Inspector name	
Date	