



Home Office

232300 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	Brian Huntly
PPL number	PP3042348 (Huntly) Investigation of normal haematopoietic stem cell subversion and the evolution, maintenance and targeting of haematological malignancies
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	17/10/2023
2. Species	Mouse
3. Brief details of study and how the limits or constraints have been breached	The specific study uses a murine leukaemia cell line that gives rise to leukaemic transformation once cells have acquired additional spontaneous mutations (also called 'second hits'). As such, mice under this study plan had received a lethal dose of irradiation (on the 19 th of July 2023) followed by intravenous injection of murine cells. Follow up analyses confirmed engraftment (a blood sample on the 27 th of September revealed very low levels of these cells in the peripheral blood (~0.0-0.9%)). The study was comprised of 15 mice. One mouse on study plan SP133512 was found dead on Tuesday 17 th of October 2023.
4. Have you taken advice from the Named Animal Care and Welfare Officer/Named Veterinary Surgeon/others?	One animal was found dead. No advice was deemed necessary and the NACWO was informed that tissues would be analysed and a Condition 18 report would be sent.
5. Cause of Problem, if known	Leukaemic disease progression. Macroscopic examination of the animal demonstrated splenomegaly, (an obligate phenotype for this model system). This animal had been visually inspected on the previous day(s) by both the AMB staff and a member of our team and displayed no clinical symptoms. Despite viability having been compromised due to the state of decomposition, we were able to confirm the presence of leukaemic cells in Spleen and Bone marrow by flow cytometry analysis of cells.

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>	The remaining animals in this cohort were thoroughly inspected for any signs of distress or ill health (including weighting and a scheduled blood sample). One more animal was identified with increased white blood counts and was sacrificed. No weight loss was observed for this animal. However, its blood parameters are in agreement with disease progression (elevated white blood counts) and we have observed splenomegaly and hepatomegaly too. No other animal displayed elevated white blood counts at this stage. All blood samples are currently being analysed to establish the percentage of murine leukaemic cells. These mice will continue to be monitored and an additional blood sampling has been scheduled two weeks from today.
7. Reported by (name)	George Giotopoulos
8. Date	19/10/2023

*****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 19/12/23
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