

Transgenerational Effects in the Progeny of Mice Exposed to Acute High  
and Chronic Low Dose-rate Gamma-rays  
– Germ Cell Mutation Analyses–

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Abstract

Transgenerational effects of continuous low dose-rate (LDR)  $\gamma$ -ray irradiation of male mice have not been well studied. We have been analyzing the incidence of copy number aberrations (CNAs) in the progeny of male C57BL/6J mice continuously exposed to LDR (20 mGy/day, 1mGy/day, 0.05mGy/day)  $\gamma$ -rays for 400 days (total dose: 8000 mGy, 400 mGy, 20 mGy) and to high dose-rate (HDR, 770 mGy/min)  $\gamma$ -rays (total dose: 3000 mGy). This year, we analyzed 24 progenies from 4 pairs of parents in the 3000 mGy HDR irradiated group using oligo-microarray CGH (Agilent Technologies), and found two mice with Type L genomic aberrations. We also verified by the TaqMan® Copy Number Assay the aberrations in the 20 mGy/day irradiated group and non-irradiated group as "real new mutations".

Table 1 Mice with aberrations detected by oligo-microarray CGH

|                                      |        | No. of mice<br>examined using<br>oligo-microarray assay | No. of mice<br>with type L<br>aberrations |         |
|--------------------------------------|--------|---|---|---------|
| 20 mGy/day<br>(Total dose 8000 mGy)  | Female | 48  | 3   | 6.3 (%) |
|                                      | Male   | 63  | 8   | 12.7(%) |
|                                      | TOTAL  | 111   | 11  | 9.9 (%) |
| 1 mGy/day<br>(Total dose 400 mGy)    | Female | 24  | 2   | 8.3 (%) |
|                                      | Male   | 21  | 3   | 14.3(%) |
|                                      | TOTAL  | 45  | 5   | 11.1(%) |
| 0.05 mGy/day<br>(Total dose 20 mGy)  | Female | 21  | 1   | 4.7 (%) |
|                                      | Male   | 25  | 0   | 0.0 (%) |
|                                      | TOTAL  | 46  | 1   | 2.1 (%) |
| 770 mGy/day<br>(Total dose 3000 mGy) | Female | 14  | 0   | 0.0 (%) |
|                                      | Male   | 10  | 2   | 20.0(%) |
|                                      | TOTAL  | 24  | 2   | 8.3 (%) |
| Non-irradiated                       | Female | 73  | 2   | 2.7 (%) |
|                                      | Male   | 67  | 2   | 3.0 (%) |
|                                      | TOTAL  | 140   | 4   | 2.8 (%) |

Table 2 Results of aberration verification by TaqMan® Copy Number Assay

| No. of probes<br>with aberrations | No. of aberrations<br>detected by<br>oligo-microarray CGH |            | No. of aberrations<br>verified by TaqMan®<br>Copy Number Assay |            |
|-----------------------------------|---|------------|--|------------|
|                                   | Non-irradiated  | 20 mGy/day | Non-irradiated   | 20 mGy/day |
| Type L                            | 5   | 11         | 5  | 11         |
| Type S                            |   |            |  |            |
| ≥ 5                               | 2   | 3          | 2  | 3          |
| 4                                 | 1   | 2          | 0  | 0          |
| 3                                 | 2   | 2          | 1  | 1          |
| 2                                 | 12  | 3          | 1  | 0          |
| Total                             | 17  | 10         | 4  | 4          |
| TOTAL (Type L + S)                | 22  | 21         | 9  | 15         |