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 low dose-rate (LDR) radiation have not been well studied. We have estimated the incidence of copy number variations (CNVs) in the progeny born from 65 week-old male C57BL/6J mice after continuous exposure to gamma-rays at several LDRs (0.05, 1 or 20 mGy/day for 400 days to total doses of 20, 4000 or 8000 mGy) or from 18 week-old male C57BL/6J mice after exposure to a HDR of 770 mGy/min (total dose 3 Gy). This year, using array CGH to screen for CNVs, we analyzed 44 progenies born from 65 week-old male C57BL/6J mice after exposure to a HDR of 770 mGy/min (total dose 3 Gy). To identify "real new mutations," we tested all candidate CNVs found using quantitative PCR and estimated that the frequency of F1 mice containing CNVs in this group is 9.1%.

Table 1. The number of the (candidate) CNVs detected by array CGH and confirmed by qPCR

Number of affected probes	Number of affected probes in 2nd CGH	Non-irradiated Mating: 65 weeks			20 mGy/day (8000 mGy) Mating: 65 weeks			70 mGy/min (3000 mGy) Mating: 18 weeks			70 mGy/min (3000 mGy) Mating: 65 weeks		
in 1st CGH		Detected by CGH	Confir by PC		Detected by CGH	Confirmed by PCR		Detected by CGH	Confirmed by PCR		Detected by CGH	Confirmed by PCR	
≧2 (Type L)		9	9	[1]	23	23 [1]		5	5		1	1	
1 (Type S)	≧5	1	1	[1]	5	5		0	0		0	0	
	4	1	0		2	1		0	0		0	0	
	3	3	1	[1]	2	1		1	0		1	0	
	2	21	3		3	2		8	3		4	2 [1]	
	(Total)	26	5	[2]	12	9		9	3		5	2 [1]	
(Type L+S)		35	14	[3]	35	32 [1]		15	8		6	3 [1]	
(No. of mice analyzed)		156		142			59		44				

The numbers in brackets indicate those of duplications (internal numbers).

Table 2. The number of the F1 mice with CNVs

	No. of mice				No. of mic					
		analyzed	Type L			Ty	Type L+S			
770 mGy/min (3000 mGy) Mating: 65 weeks	Female	25	1	(4.0)		1	(4.0)	2	(8.0)	_
	Male	19	0	(0.0)		2	(10.5) [1]	2	(10.5)	[1]
	Total	44	1	(2.3)		3	(6.8) [1]	4	(9.1)	[1]
770 mGy/min	Female	28	2	(7.1)		0	(0.0)	2	(7.1)	
(3000 mGy)	Male	31	3	(9.7)		2	(6.5)	5	(16.1)	
Mating: 18 weeks	Total	59	5	(8.5)		2	(3.4)	7	(11.9)	
20 mGy/min	Female	67	8	(11.9)	[1]	9	(13.4)	17	(25.4)	*[1]
(8000 mGy)	Male	75	13	(17.3)		2	(2.7)	15	(20.0)	*
Mating: 65 weeks	Total	142	21	(14.8)	**[1]	11	(7.7)	32	(22.5)	**[1]
NT 1 1 1	Female	81	4	(4.9)		6	(7.4) [2]	10	(12.3)	[2]
Non- irradiated Mating: 65 weeks	Male	75	4	(5.3)	[1]	1	(1.3)	5	(6.7)	[1]
Triumg. 05 Weeks	Total	156	8	(5.1)	[1]	7	(4.5) [2]	15	(9.6)	[3]

The numbers in brackets indicate those of duplications (internal numbers). *P<0.05, **P<0.01