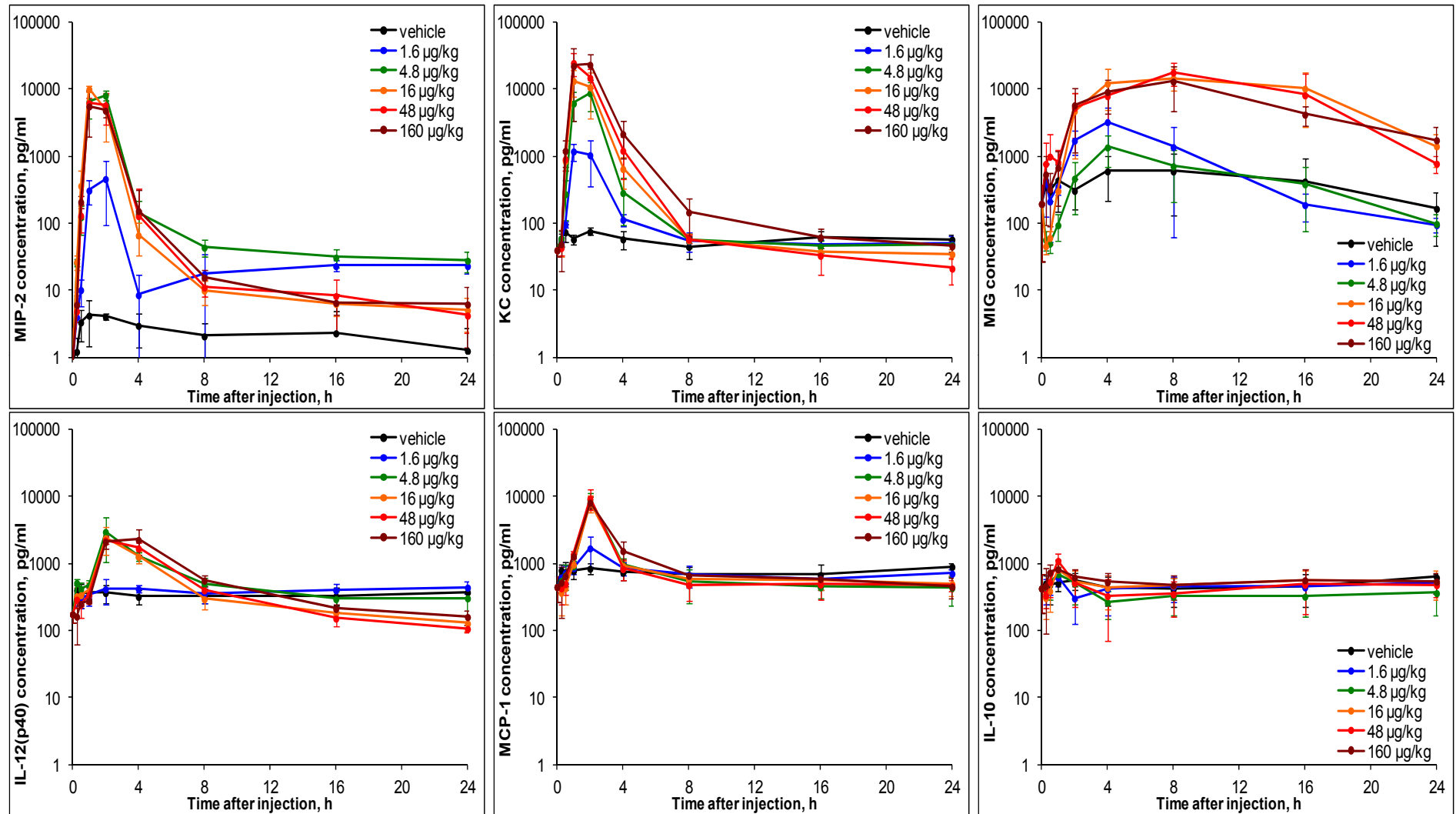


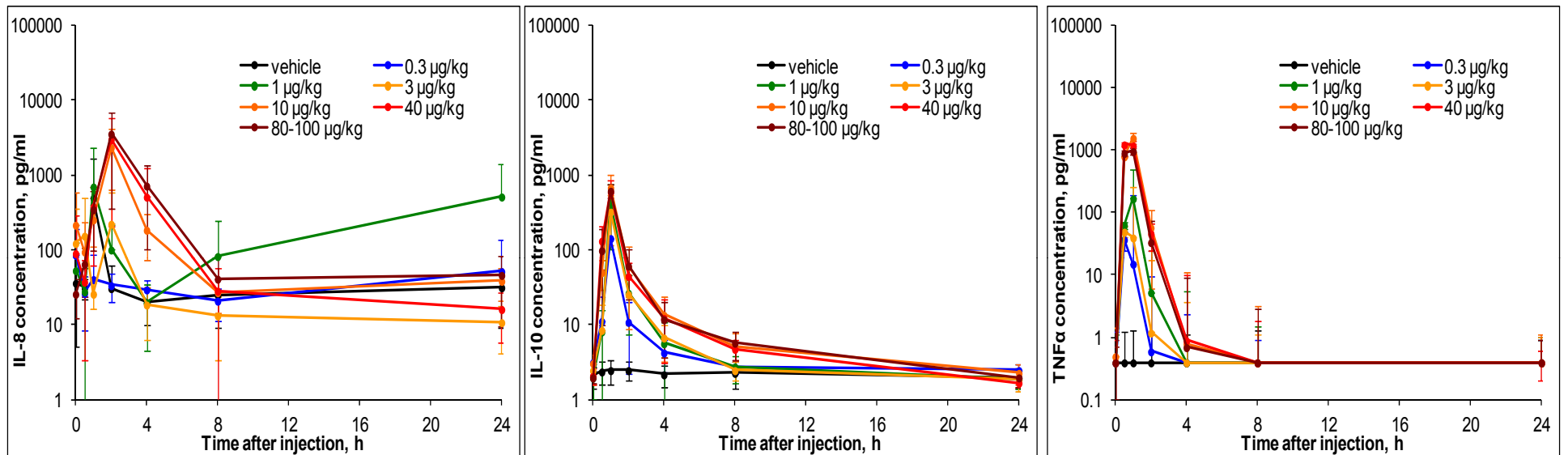
Supplemental Figure S1. Additional cytokines induced by CBLB502 in non-irradiated ICR mice after i.m. injection. Time-dependent changes in plasma cytokine levels in non-irradiated female ICR mice injected i.m. with either the vehicle or with the indicated doses of CBLB502. Each data point represents mean \pm SD of cytokine concentration measured in plasma of 3 mice. G-CSF, IL-6, KC, IP-10, Eotaxin, IL-10, IL-1a, IL-1b, IL12p70, TNF, GM-CSF, TPO, SCF were tested. IL-1a, IL-1b, IL12p70, TNF, GM-CSF, TPO, SCF did not display substantial or consistent response. IL-10 had shown very minor response.



Supplemental Figure S2. Additional cytokines induced by CBLB502 in non-irradiated C57BL/6J mice after s.c. injection. Time-dependent changes in plasma cytokines levels in non-irradiated female C57BL/6J mice injected s.c. with either the vehicle or with the indicated doses of CBLB502. Each data point represents mean \pm SD of cytokine concentration measured in plasma of 6 mice. G-CSF, IL-6, KC, IL-10, IL12p40, TNF, MCP-1, MIP-2, MIG were tested. IL-10 showed a very minor response. TNF did not show a consistent response. One outlier sample (vehicle/16 h, very high levels of MIG and MIP-2) was excluded.

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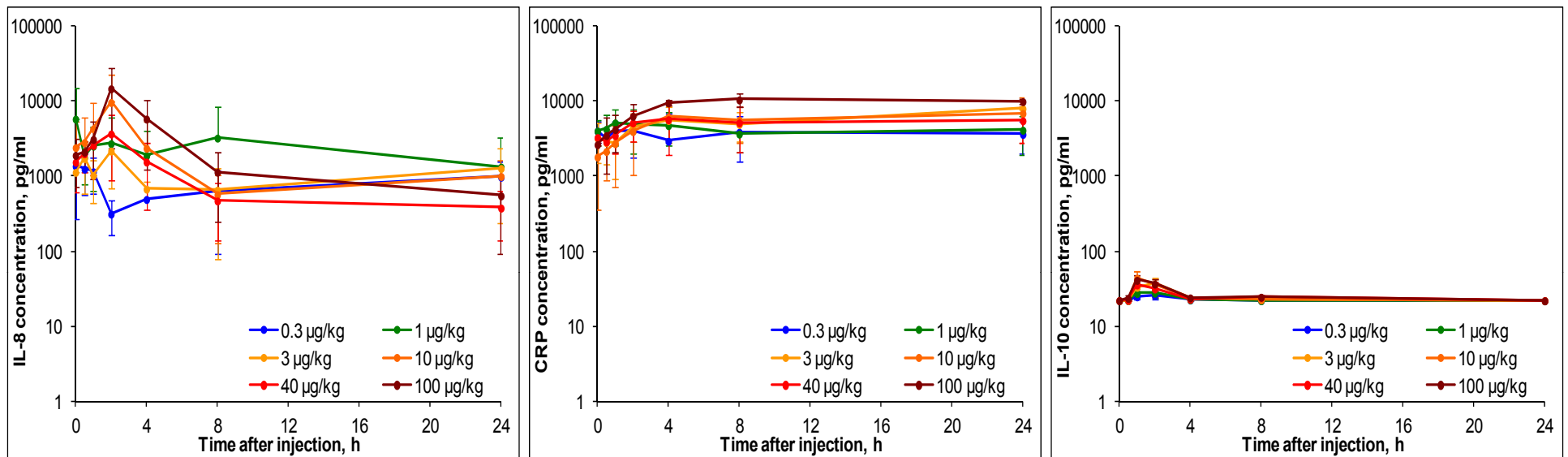


Supplemental Figure S3. Additional cytokines induced by CBLB502 in non-irradiated beagle dogs after i.m. injection. Time-dependent changes in plasma cytokine levels in non-irradiated male and female beagle dogs (n=6, 1:1 sex ratio) injected i.m. with either the vehicle or with the indicated doses of CBLB502. Each data point represents mean \pm SD. IL-6, IL-8, IL-10, and TNF were tested; all showed a CBLB502-dependent response.

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Supplemental Figure S4. Additional cytokines and other factors induced by CBLB502 in non-irradiated rhesus macaques after i.m. injection. Time-dependent changes in levels of plasma cytokines and other factors in non-irradiated male and female rhesus macaques (n=6, 1:1 sex ratio) injected i.m. with either the vehicle or with the indicated doses of CBLB502. Each data point represents mean \pm SD. G-CSF, IL-6, IL-8, IL-10, C-reactive protein, IL-1b, IL12p70, IP-10, GM-CSF were tested. IL-1b, IL12p70, IP-10, GM-CSF did not display substantial or consistent responses. IL-10 showed a very minor response. However, this was later found to be assay-related in later studies, with larger-scale IL-10 responses observed with newer assays.

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Supplemental Table S1. Effect of 60 µg/kg CBLB502 treatment administered s.c. or i.v. 24 hours after 8.5-9.5 Gy TBI on survival of irradiated C57BL/6J mice.

Route	TBI dose (Gy)	30-day survival (% and fraction)		P-value vs. control	Absolute survival increase, %	Odds ratio vs. control
		vehicle	CBLB502			
s.c.	8.5	52.5% (21/40)	85.0% (34/40)	0.003	+32.5%	5.1
	9	22.5% (9/40)	45.0% (18/40)	0.058	+22.5%	2.8
	9.5	7.5% (3/40)	27.5% (11/40)	0.037	+20.0%	4.7
i.v.	8.5	82.5% (33/40)	97.5% (39/40)	0.015	+15.0%	8.3
	9	55.0% (22/40)	80.0% (32/40)	0.031	+25.0%	3.3
	9.5	10.0% (4/40)	45.0% (18/40)	0.0009	+35.0%	7.4