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Mechanism of Action and Detection

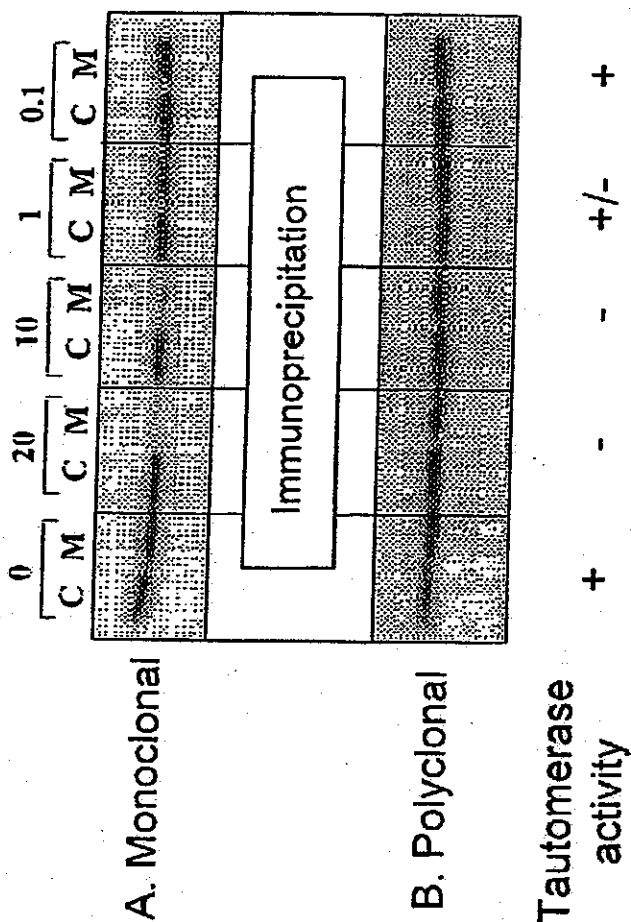


FIG. 1

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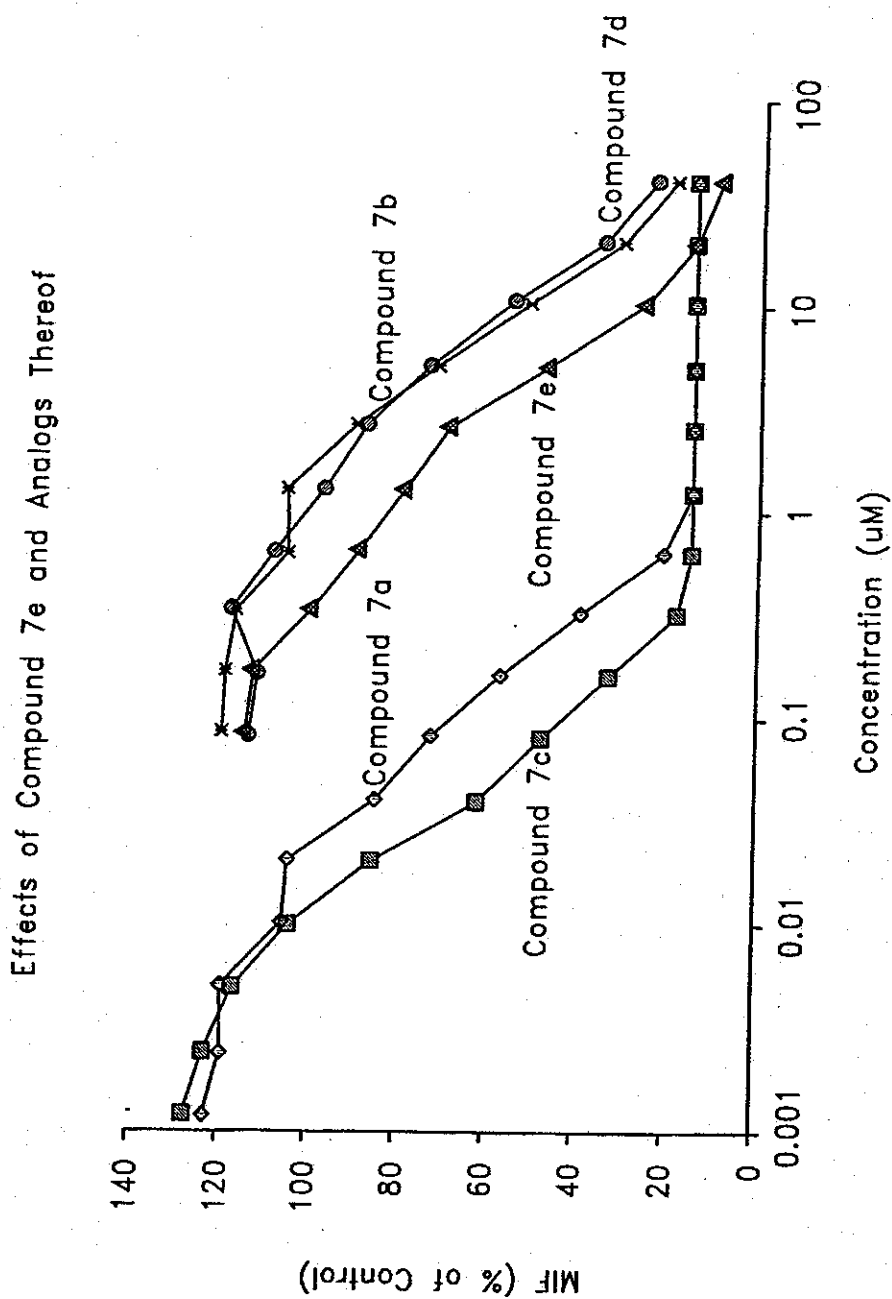


FIG. 2

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Compound 7e Inactivates MIF Produced by THP1 Cells

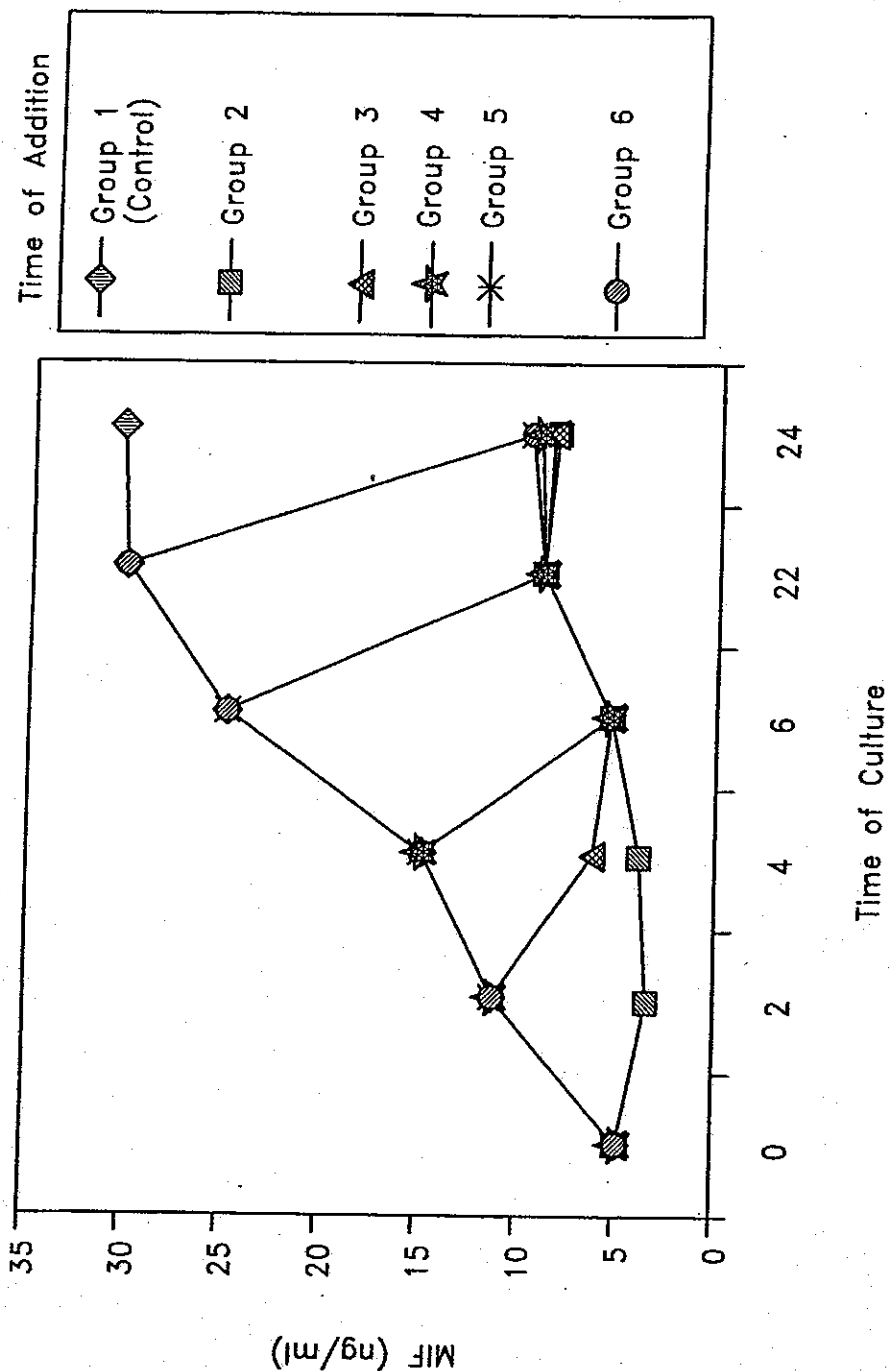


FIG. 3

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Compound 7e inactivates MIF in Conditioned Media

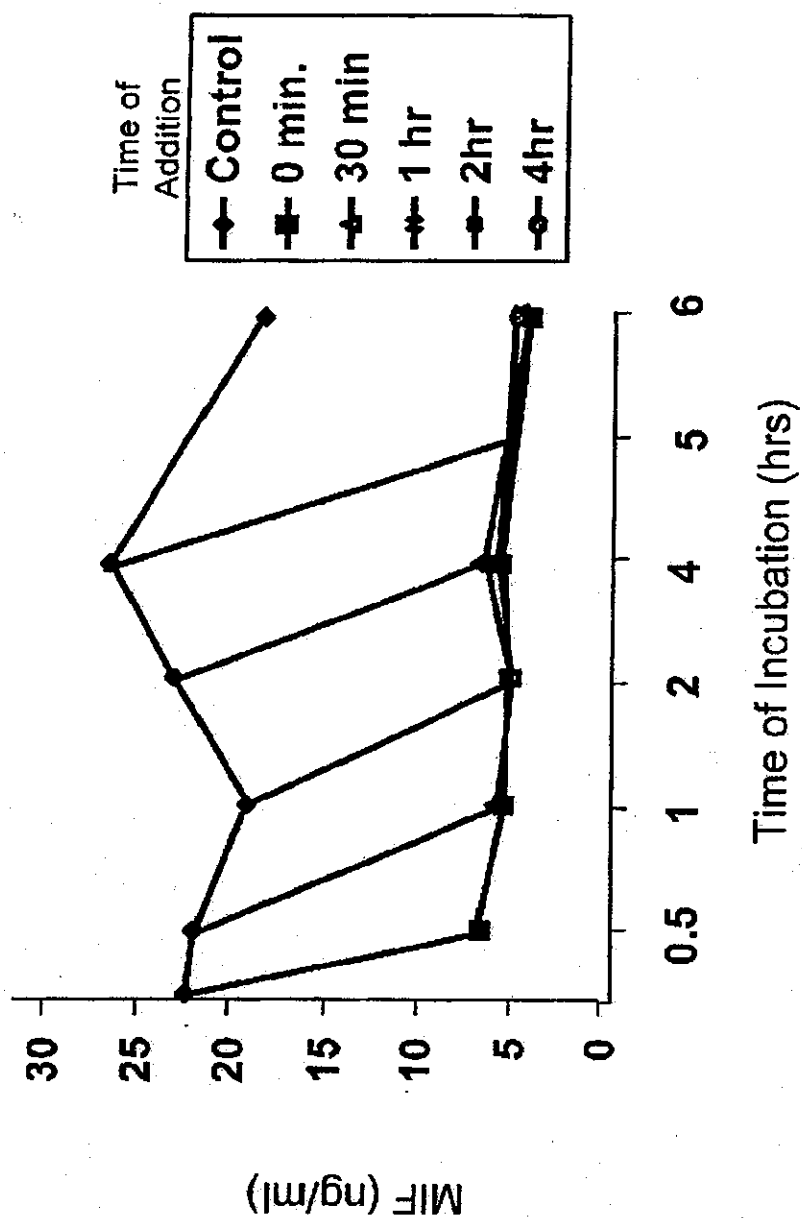


FIG. 4

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Inhibition of LPS-Induced MIF Release from RAW 264.7 Cells by Compound 7e

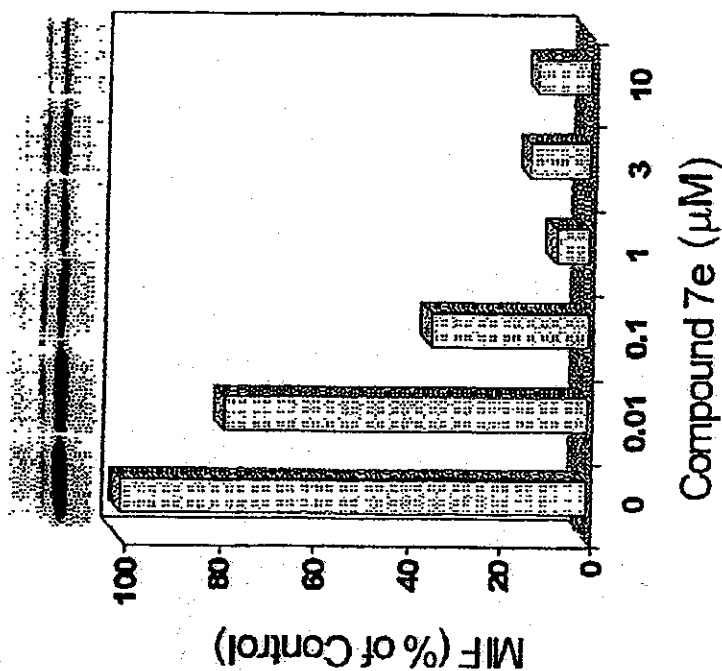


FIG. 5

Inhibition of TSST-1-Induced MIF Release from RAW 264.7 Cells by Compound 7e

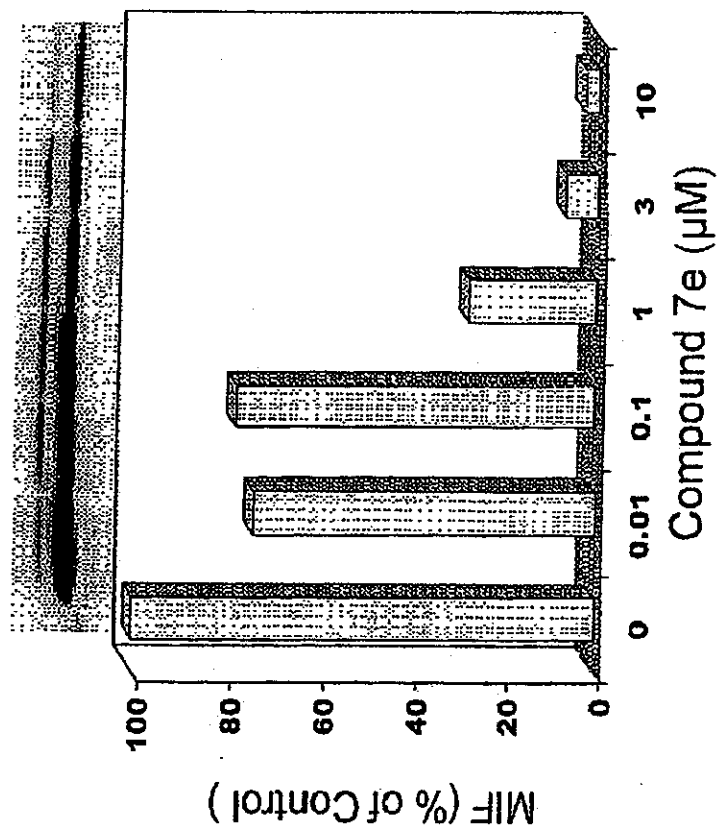


FIG. 6

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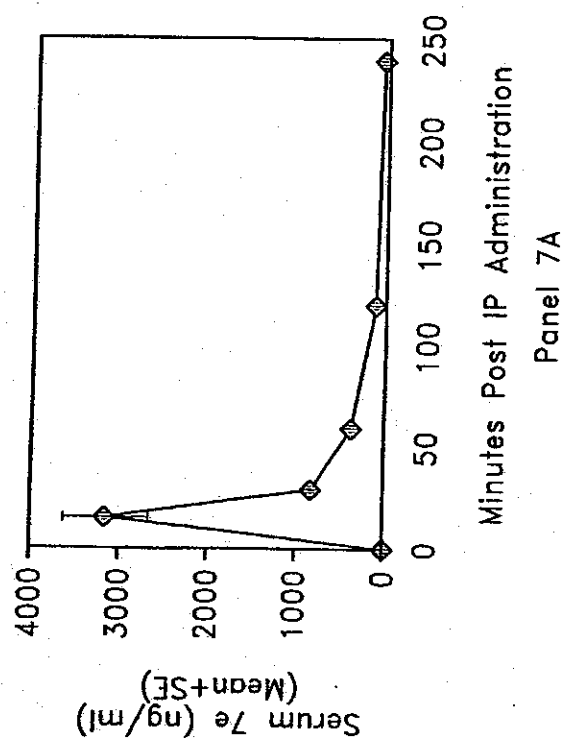
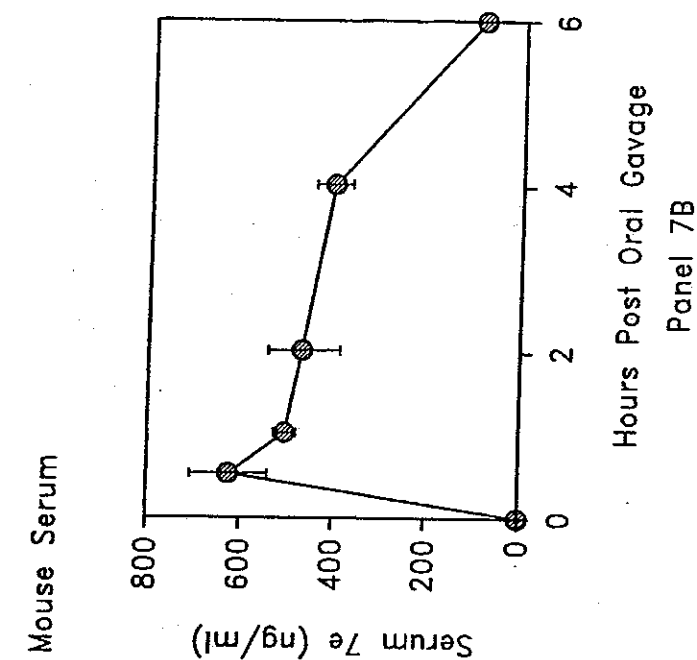
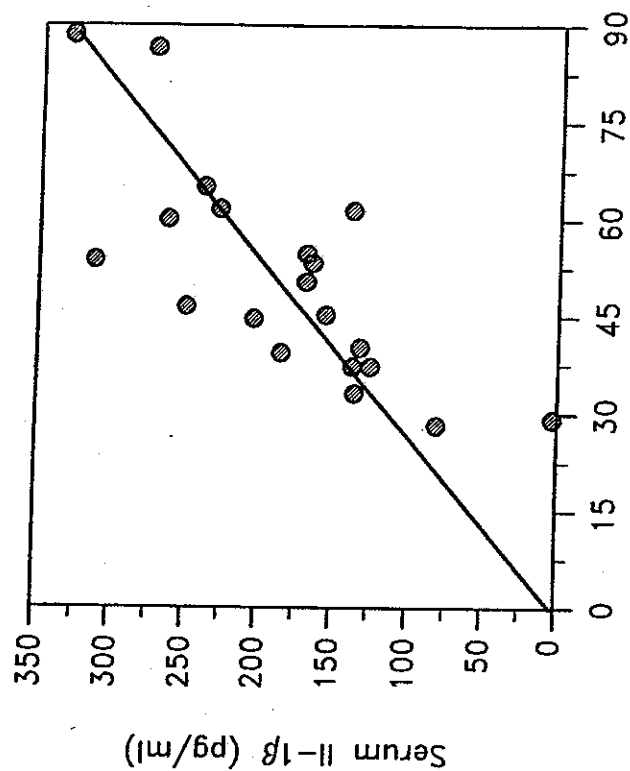


FIG. 7

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Correlation of Serum IL-1 β and Serum MIF



Serum MIF (ng/ml)
 $Y = .112 + 3.713 * X; R^2 = .589$

FIG. 10

LPS-Induced MIF Release in Mice

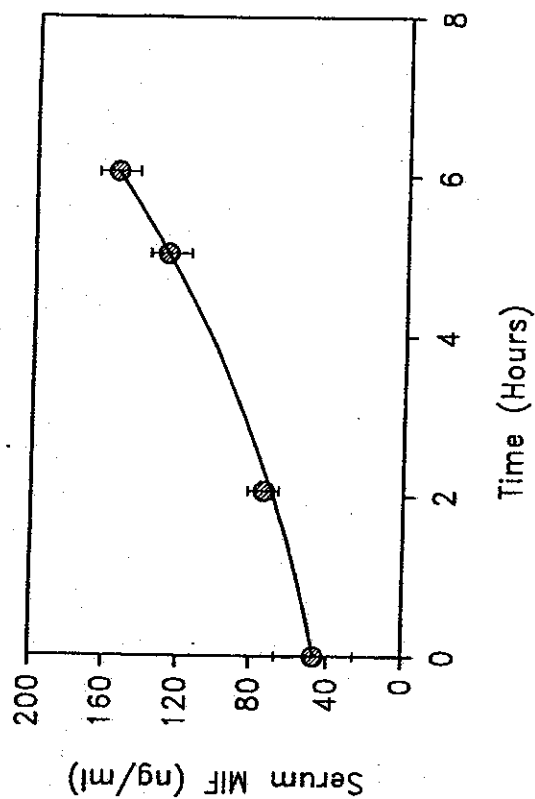


FIG. 8

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Compound 7e Inhibits LPS Induced MIF in Mice

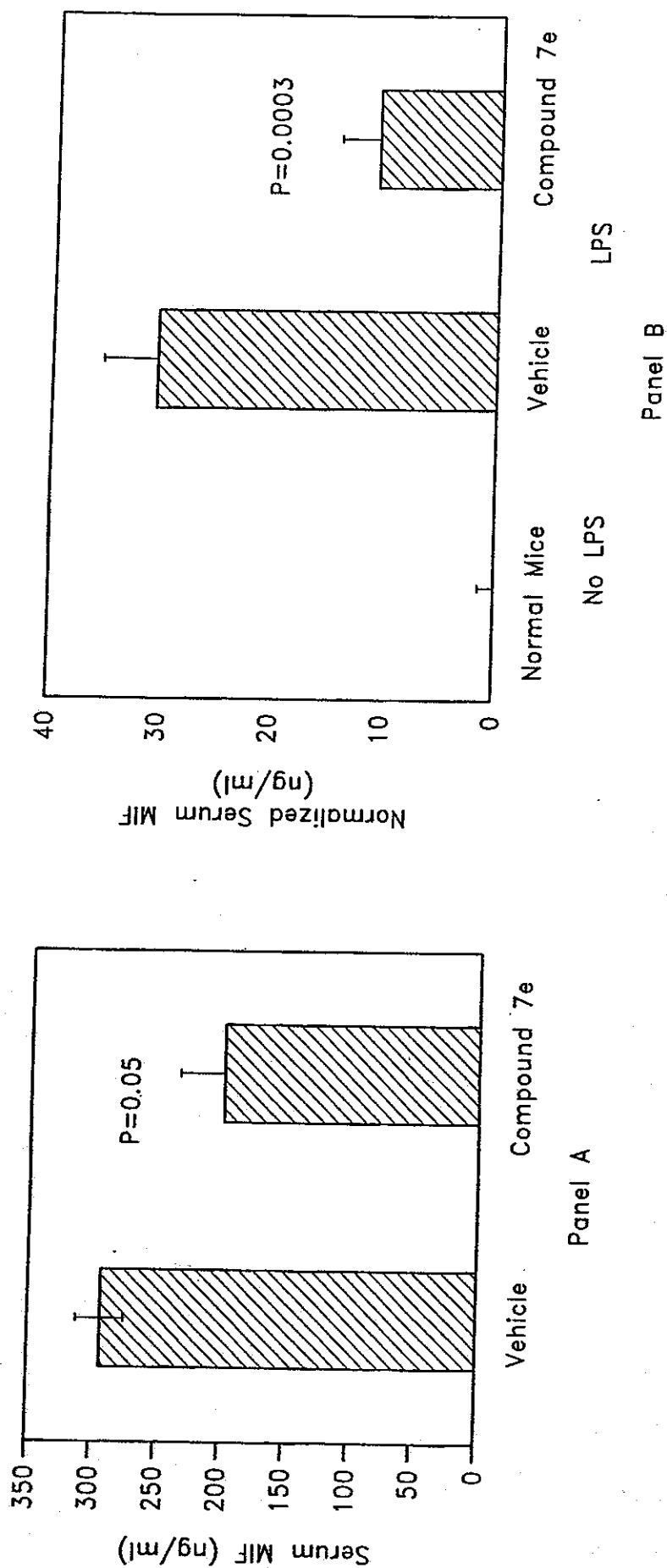


FIG. 9

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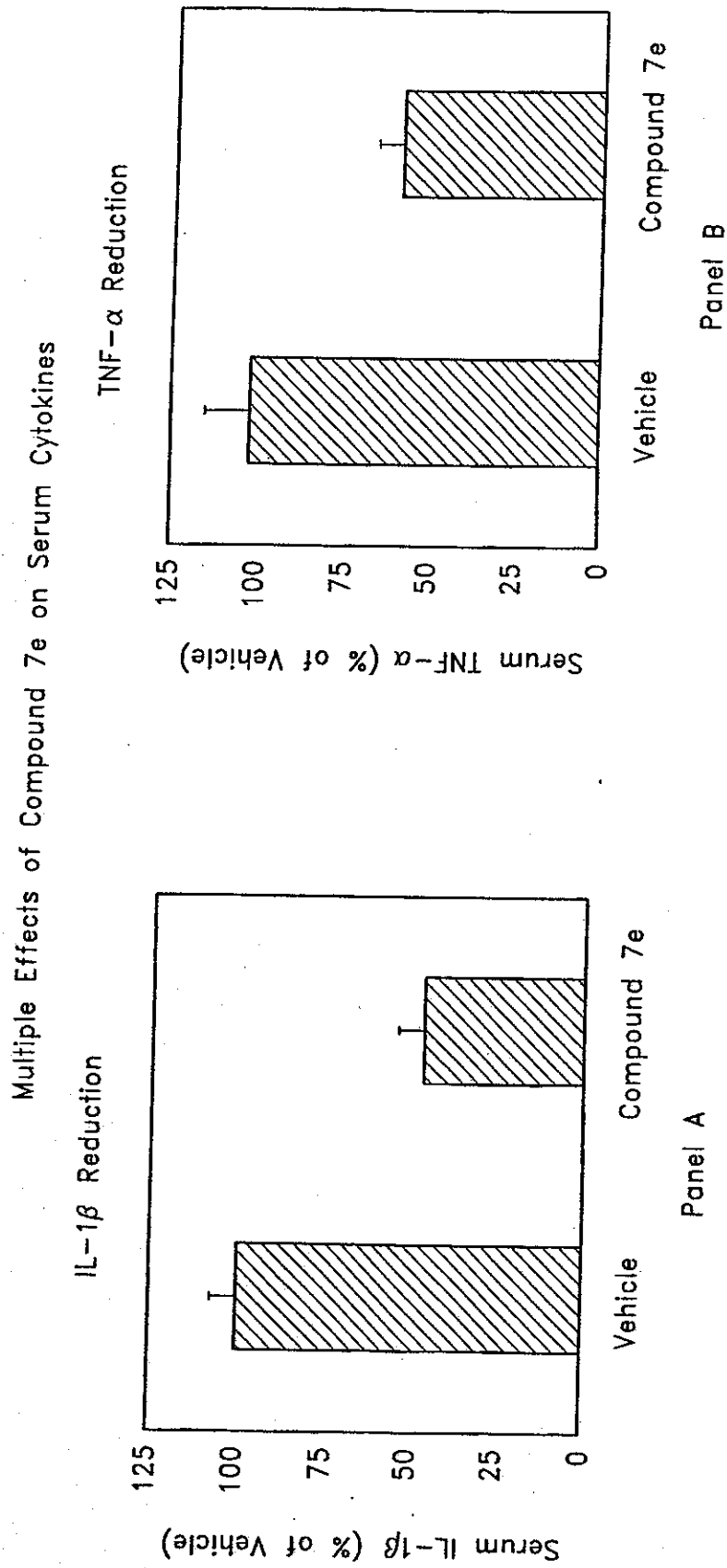


FIG. 11

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Effect of Compound 7e LPS Treated Mice
Kaplan-Meier Assess of Survival

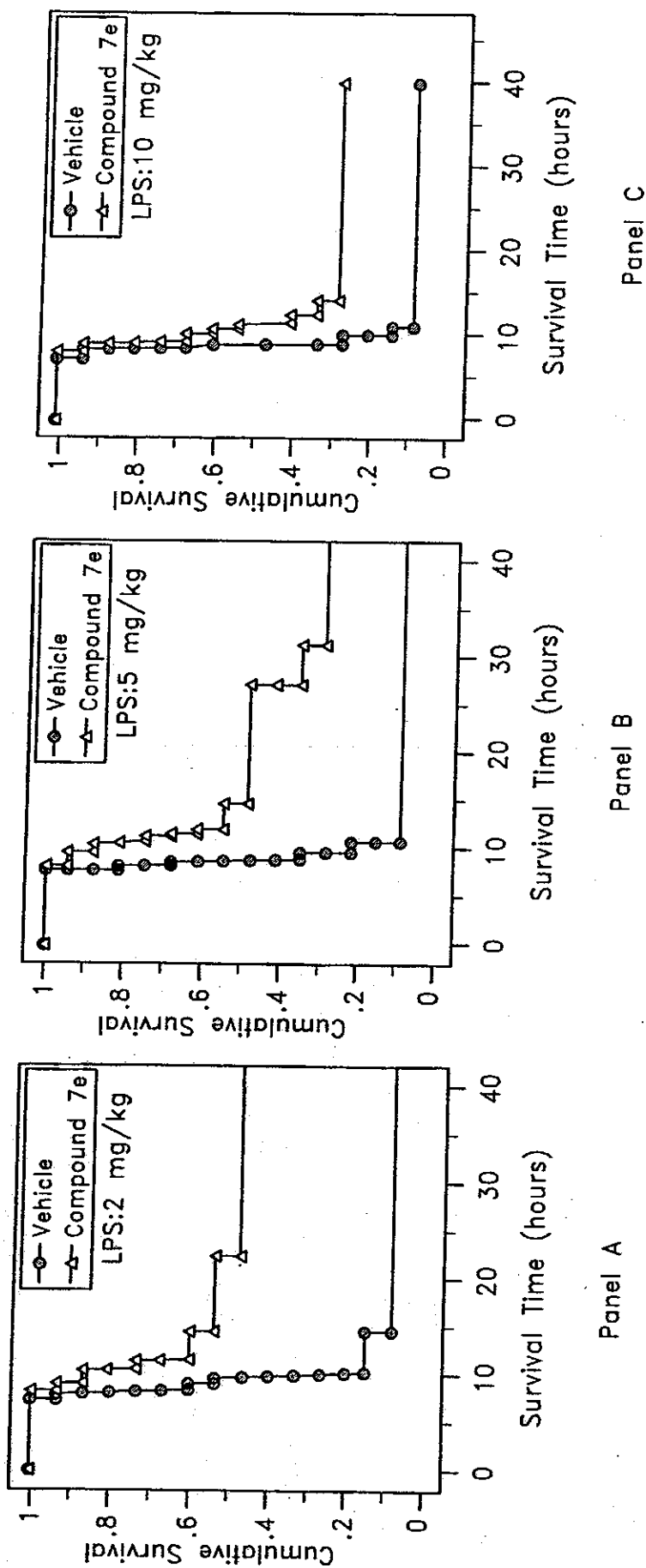
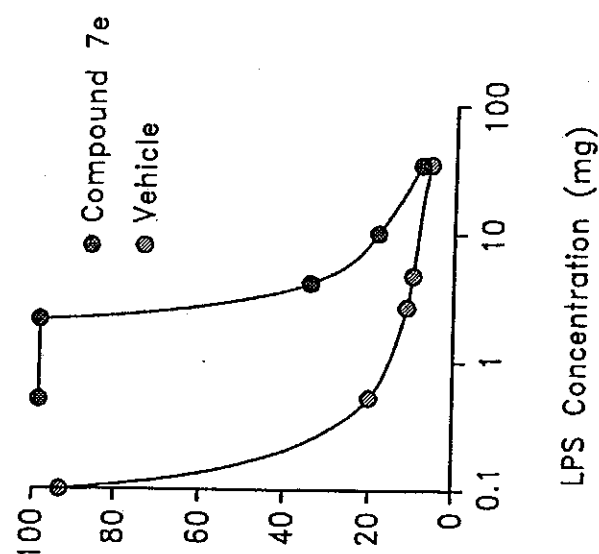


FIG. 12

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Protective Effect of Compound 7e on LPS Induced Toxicity

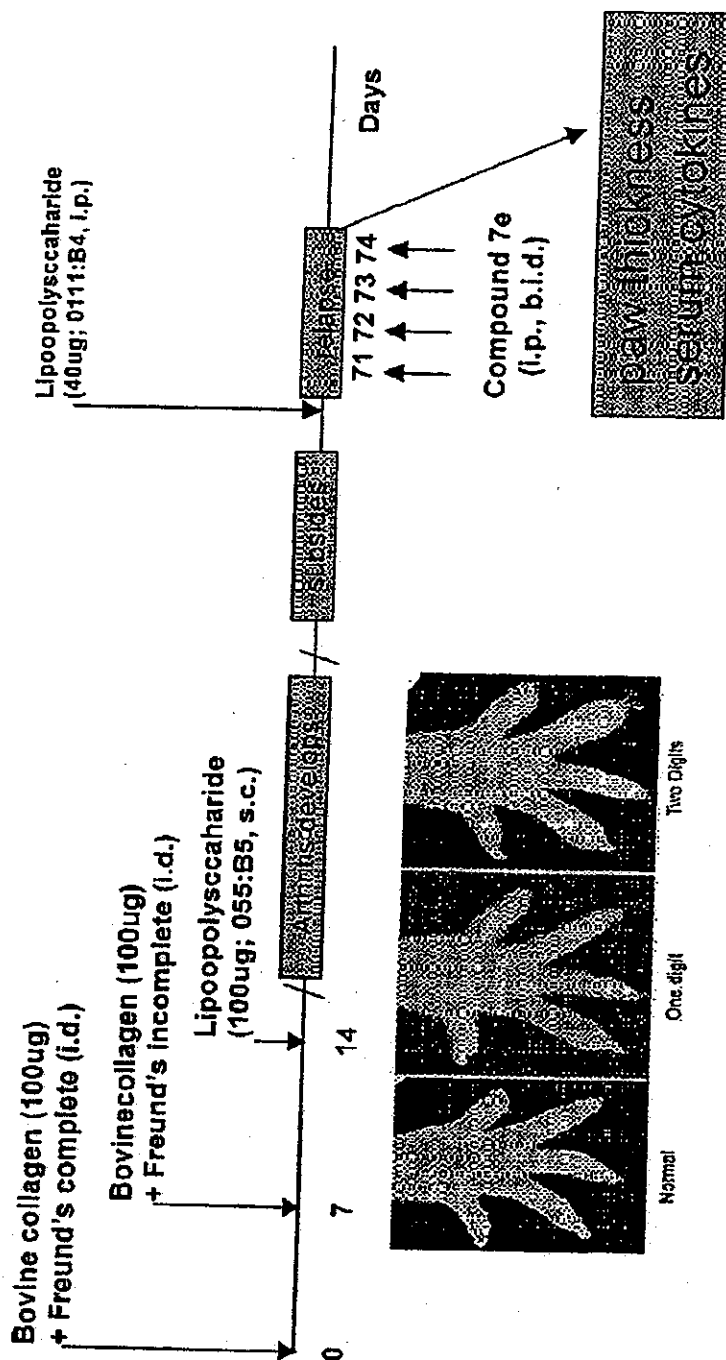


LPS Dosage (mg/kg)	75% Animal Death (hours)	
	Vehicle	Compound 7e
40	10.2	11.60
10	9.9	18.0
5	10.0	32.0
2	10.2	>100
0.5	22.0	>100
0.1	>100	>100

FIG. 13

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Effect of Compound 7e on Collagen-Induced Arthritis



Experimental protocol for Inhibition of MIF in RA models

FIG. 14

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Effect of Compound 7e in Collagen-Induced Model (CIA)
of Arthritis in Mice

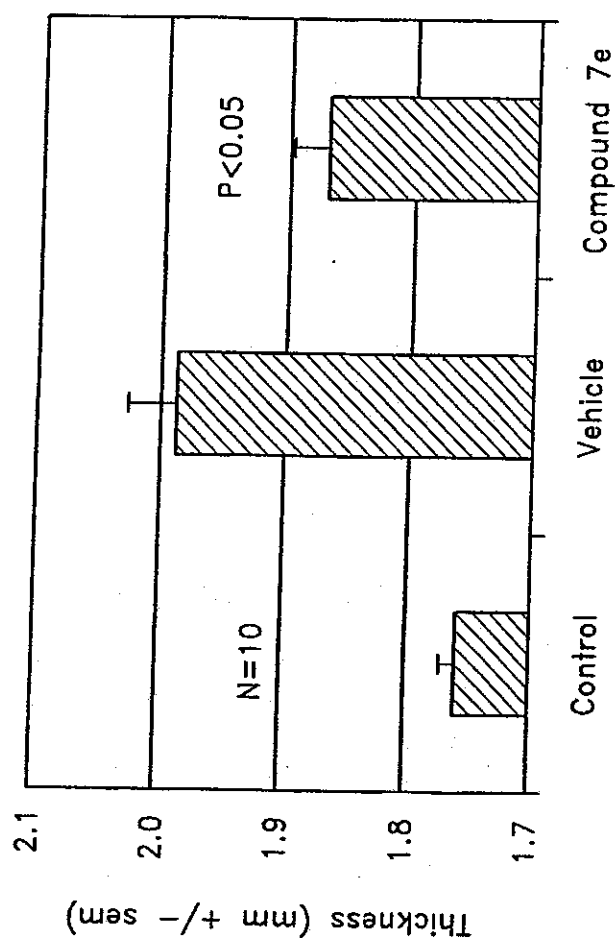


FIG. 15

Cytokine Levels in CIA Mice

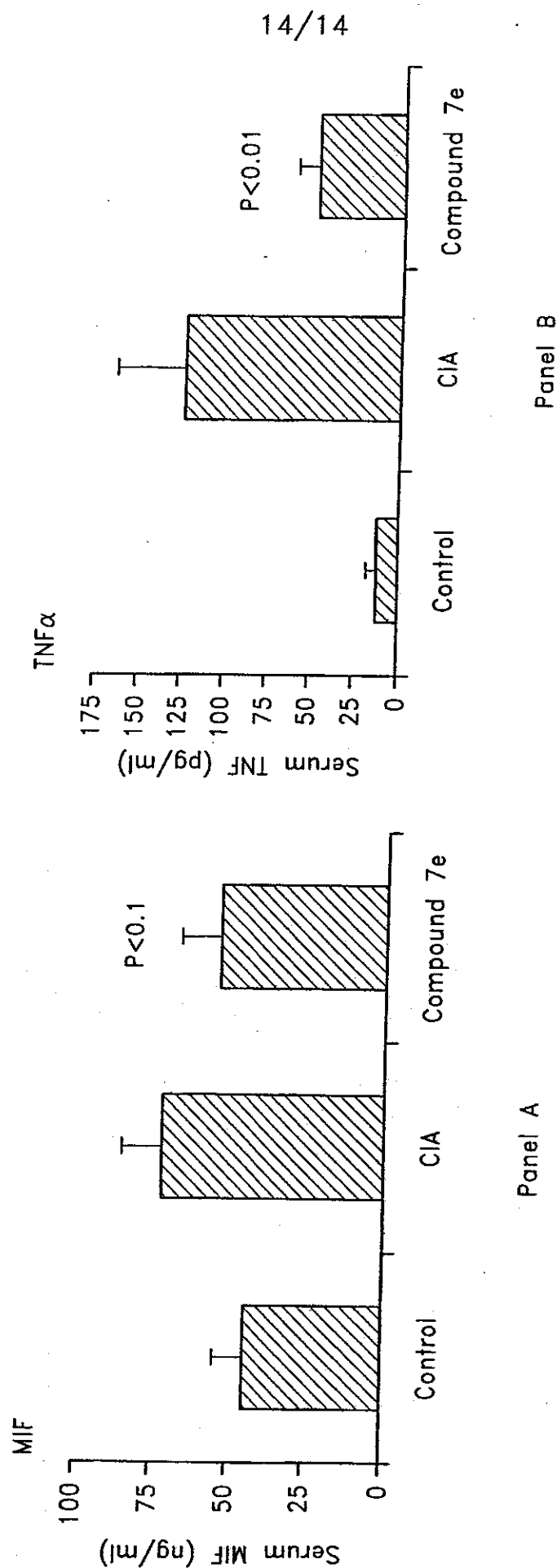


FIG. 16