## 単球走化活性化因子の免疫学的測定法の確立とその 臨床的意義の検討

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## 1993 Fiscal Year Final Research Report Summary

## Establishment of an immunoassay system for monocyte chemotacticand activating factor (MCAF) and its clinical application

Research Project

Project/Area Number
04671430
Research Category
Grant-in-Aid for General Scientific Research (C)
Allocation Type
Single-year Grants
Research Field
Laboratory medicine
Research Institution
Kanazawa University
Principal Investigator
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Co-Investigator(Kenkyū-buntansha)
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Project Period (FY)
1992 – 1993
Keywords
cytokine / chemokine / enzyme-linked immunosorbent assay / interleukin / monocyte chemotactic and activating factor
Research Abstract

The results of the project is summarized as follows.

- 1. We established MCAF-producing cell line by transfecting MCAF cDNA into a mouse myeloma cell line and purified MCAF from the culture supernatants of this cell line. We observed that the preadministration of purified recombinant MCAF could prevent either normal or granulocytopenic mice from lethal infection caused by pseudomonas aeruginosa o r Salmonella typhymurium.
- 2. We established a specific and sensitive ELASA against human MCAF. Using this ELASA, we observed that the pleural fluid levels of MCAF increased in patients with malignant pleurisy upon the administration of an immunomodulator, OK-432. In addition, most human glioblastoma cell lines could produce MCAF in response to IL-1 or TNFalpha, suggesting a potential role of MCAF in tumor immunity.
- 3. We also examined the role of MCAF-related molecule, interleukin-8 (IL-8) in the acute inflammation by administering a neutralizing antibody against IL-8. these experiments demonstrated that IL-8 plays an essential role in neutrophil infiltration and neutrophi-dependent tissue injuries in lipopolysaccharide-induced dermatitis and lung reperfusion injury.
- 4. The determination of urinary IL-8 lavels using ELISA revealed that urinary IL-8 levels increased in the urines of patients with urinary tract infections and those with several types of glomerulonephritis such as lupus nephritis, acute glomerulonephritis, and IgA nephropathy with acute exacerbations.

These results imply that the determination of urinary IL-8 level can be used for diagnosing and monitoring several diseases in urinary tracts. In addition, we explored the molecular biological aspects of IL-8 gene activation and established that NF\_<-k>B site was essentially involved in the transcription of IL-8 gene in every cell line that we examined.

## Research Products (19 results)

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	All	Publica	ations (	(19 re	sults
[Publications] Ko,YC.,Mukaida,N.,Panyutich,A.et al.: "A sensitive enzyme-linked immunosorbent assay for human interleuk Immunological Methods. 149. 227-235 (1992)	in 8.	." Journa	al of		~
[Publications] Yasumoto,K.,Okamoto,Si.,Mukaida,N.,et al.: "Tumor necrosis factor α and interferon γ synergistically induce a human gastric cancer cell line through acting concurrently on AP-1 and NF-kB like binding sites of the interleukin 8 gene." Chemistry. 267. 22506-22511 (1992)					in 🗸
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[Publications] Sekido,N.,Mukaida,N.,Harada,A.,et al.: "Prevention of lung reperfusion injury in rabbits by a monoclonal antib Nature. 365. 654-657 (1993)	ody	against	interleu	ıkin-8	." 🗸
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[Publications] Nakano,Y.,Kasahara,T.,Mukaida,N.,et al.: "Prevention of fatal bacterial infection in mice by monocyte chemota factor(MCAF)." Infection and Immunity. (印刷中).	ıctic	and acti	ivating		~
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