Analysis of the immunological role of spleen in gastric cancer patients: from the viewpoint of cytokine producing ability of splenocytes

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## 1999 Fiscal Year Final Research Report Summary

## Analysis of the immunological role of spleen in gastric cancer patients - from the viewpoint of cytokine producing ability of splenocytes -

**Research Project** 

Project/Area Number
10671167
Research Category
Grant-in-Aid for Scientific Research (C)
Allocation Type
Single-year Grants
Section
Research Field
Digestive surgery
Research Institution
Kanazawa University
Principal Investigator
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Project Period (FY)
1998 – 1999
Keywords
gastric carcinoma / splenocyte / Th1 / Th2 / IL-12 / IFN-γ / OK-432 / tumor bearing state / cytokine

## **Research Abstract**

Cytokine production from splenocytes of healthy mouse induced with OK-432 differs depending their genetic background. In C57BL/6 mice with B16 melanoma, splenocytes derived from the middle stage of tumor bearing produced IL-2, IL-6, IL-10 and IFN- $\gamma$ . There was little spontaneous IL-12 production from tumor bearing mice splenocytes. All cytokine production were reduced in their later stage of tumor bearing. Splenocytes derived from the middle stage of tumor bearing mice could produce IL-12 when activated with OK-432 in vitro. Furthermore, in vivo treatment with OK-432 tend to inhibit tumor growth of B16 melanoma, which was accompanied by IL-12 and IFN- $\gamma$  production of splenocytes. Regarding the splenocytes of gastric cancer patients, production of IL-4, IL-6 and IFN- $\gamma$  was observed but we need further

investigation because there exist individual differences between the patients.

In the ascitic mouse model, OK-432 could induce Th1 cytokine production such as IL-12 and IFN- $\gamma$  in the ascites. In human treatment of gastric cancer, we have tried to administer MMC and OK-432 to control peritoneal carcinomatosis, and there were responder cases with increasing I L-2, IL-12, ILI5 and IFN- $\gamma$  in their cancerous ascites. There was a correlation between their favorable prognosis and their Th1 cytokine production after the OK-432 treatment.

## Research Products (10 results)

lia	Other
All Publica	ations
[Publications] 藤本敏博、他: "癌性胸腹膜炎に対するOK-432局所投与の抗腫瘍効果と作用機序について"Biotherapy. 12巻11号. 1479-1485 (1998)	~
[Publications] 藤本敏博、他: "Th1 inducerとしてのOK-432の役割"Therapeutic Rssearch. 19巻7号. 113-119 (1998)	~
[Publications] Fujimoto, T. et al.: "The effect of OK-432 on the cytokine production of the tumor bearing mouse splenocytes"Biotherapy. (in press). (2000)	~
[Publications] 藤本敏博、他: "新しい癌免疫化学療法の指針、QOLを重視した癌薬物療法"医薬ジャーナル社. 291 (1999)	~
[Publications] 藤本敏博、他: "肝転移、メカニズムと臨床"医学書院. 228 (2000)	~
[Publications] Fujimoto T, Hirano K, Nomura H, Ohta T, Takahashi Y, Mai M and Minami S: "Antitumor effect and immune mechanism of local OK-432 administration agains pleuro-peritoneal carcinomatosis."Biotherapy. 12(11). 1479-1485 (1998)	<sup>;t</sup> 🗸
[Publications] Fujimoto T, Mai M and O'Donnell MA: "Streptococcal preparation OK-432 works as a Th1 inducer."Therapeutic Research. 19(7). 2187-2193 (1998)	~
[Publications] Fujimoto T and Mai M: "Local administration of OK-432 against peritoneal carcinomatosis. New therapeutic guideline of immunochemotherapy for cancer." (Saji S and Toge T ed.) Iyaku-journal, Tokyo. 81-84 (1999)	~
[Publications] Wang X, Fujimoto T, Zhang B, Shimizu A and Mai M: "The effect of OK-432 on the cytokine production of the tumor bearing mouse splenocytes."Biotherapy. (in press).	~
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