Letter to the Editor

Improved Subjective Tolerability of Interferon by Combination with Prednisolone

S.D. FOSSA

The Norwegian Radium Hospital, Montebello, 0310 Oslo 3, Norway

During the last year we treated five patients with metastatic renal cell carcinoma with a combination of interferon alpha-2a (Roferon®-A) and various doses of prednisolone per os (Table 1).

In three patients (patients Nos. 1-3) the reason for this combined treatment was intolerable flulike symptoms during prior interferon treatment. Patient No. 4 was treated with prednisolone due to repeated asthma-like dyspnoe episodes lasting for several hours after each interferon injection. The fifth patient had used 5 mg prednisolone per os for many years prior to interferon treatment due to peripheral polyneuropathia of unknown aetiology. This treatment was continued during interferon treatment.

All three patients experienced a remarkable reduction of the flu-like side-effects with an improvement of their general condition. The asthma-like episodes ceased completely in the 4th patient. Patient No. 5 did not at all complain of subjective toxicity.

One partial remission (patient No. 1) and a 45% tumour reduction (patient No. 4) were observed during combined interferon-prednisolone treatment.

The mode of action of interferon in solid tumours is not clear. A direct antiproliferative activity has been suggested together with immunomodulation of the host [1]. If the latter mode of action plays a major role in interferon's activity, the combination

Table 1. Subjective tolerability and objective response during combined interferon-prednisolone treatment in patients with metastatic renal cell carcinoma

Patient No.	Interferon toxicity	Before prednisolone Dose of oral prednisolone (mg/day)	After prednisolone	
			Interferon toxicity	Objective response
l	+++†	20–10	+†	PR‡
2	+++	20-10	+	PD
3	+++	10	+	PD
4	+++	10	No	45% reduction
5	Not evaluable	5	No	NC

^{*}Flu-like symptoms, muscle pain/weakness (patients 1-3); asthma-like episoded (patient 4).

^{†+++:}severe and treatment-resistant; +:mild.

[‡]PR: Partial remission; NC: no change; PD: progressive disease.

with prednisolone should be avoided from a theoretical point of view.

The observations described, however, indicate that tumour reduction can be obtained if the combination of interferon and prednisolone is used. This is partly in agreement with *in vitro* studies

showing a synergestic effect of prednisolone and interferon in haematological malignancies [2]. The benefit of this combination treatment is that the application of prednisolone seems to improve the subjective tolerability to intermediate/high doses of interferon.

REFERENCES

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- 2. Takahashi I, Oda Y, Lai M et al. Intraction between human lymphoblastoid interferon and chemotherapeutic agents in vitro. Acta Med Okayama 1984, 38, 501-504.