Letters to the Editor

CPI⁶⁰ total activity of cysteine pep-tidase inhibitors can be used as markers of the total defense of 'own' organism. I would also propose determination of the difference between CPI⁶⁰ and CPI³⁶ (Δ CPI) to find out the amount of the complex of inhibitors with cysteine endopeptidases. The parameters, taken as a ratio of the levels of 'tumor aggressivity' and 'defense of organism', might be useful in the diagnosis of a cancer, as well as in new efficient therapy.^{7,8}

References

- Hiwasa T. Possible involvement of proteinase inhibitor ras proteins in malignant transformation. Biol Chem Hoppe-Seylers 1988; 369: 239–41.
- 2. Baici A, Knopfel M, Keist R. Tumor-host interactions in the rabbit V2 carcinoma stimulation of cathepsin B in host

- fibroblasts by tumor-derived cytokine. *Invas Metast* 1988; **8**: 143–58.
- 3. Sloane BF, Honn KV. Cysteine proteinases and metastasis. *Cancer Metast Rev* 1984; **3**: 249–63.
- Lah TT, Buck MR, Honn KV, et al. Degradation of laminin by human tumor cathepsin B. Clin Exp Metast 1989; 7: 461-8.
- 5. Katunuma N. Mechanism and regulation of lysosomal proteolysis. *Cell Biol Rev* 1989; **20**: 35–61.
- Siewiński M. Method of purification of thio proteinase inhibitors from human urine. *Cancer Biochem Biophys* 1991; 12: 33-44 (P. 28976).
- 7. Siewiński M, Kręcicki T, Rak J, et al. Thiol proteinase inhibitors in saliva of patients with laryngological cancers. *Diagnost Oncol* 1992; 2: 141-5.
- 8. Kręcicki T, Siewiński M. Serum cathepsin B-like activity as a potential marker of laryngeal carcinoma. Eur Arch Otorhinolaryngol 1992; **249**: 293–5.

(Received 5 October 1992; revised version received 23 November 1992; accepted 2 December 1992)

Intrapleural and intraperitoneal palliative treatment of malignant effusions with mitoxantrone

Özgür Özyılkan, Ayşe Kars, Nilüfer Güler, Gülten Tekuzman, Eşmen Baltalı and Dinçer Fırat

Hacettepe University Institute of Oncology, 06100 Ankara, Turkey. Tel: 90-4-3117972 or 2353798. Fax: 90-4-3242009

In a recent article, Torsten *et al.*¹ investigated the effect of intrapleural mitoxantrone. They found that malignant pleural effusion could be stopped for a mean period of 3.2 months in 11 of 12 patients.

Mitoxantrone has several characteristics that make it an optimal drug for local administration, i.e. high local advantage of its pharmacokinetics, extensive tissue binding, steep dose-dependent cytotoxicity and good tolerability by the tissue.^{2,3} Since 1991 we treated seven patients with malignant effusions (three pleural and four peritoneal) with intracavitary administration of mitoxantrone. All patients were refractory to systemic chemotherapy. Pleural or peritoneal effusions were drained as completely as possible by simple needle aspiration. Mitoxantrone, dissolved in 100 ml of physiologic saline (20 mg for pleural effusion, 40 mg for

peritoneal effusion), was instilled into the cavity via a catheter. Response was defined as follows. Complete response (CR) meant that there was no reaccumulation of fluid or no progression of the residual small effusion within the first 30 days. Partial response (PR) was defined as the relapse of effusion up to 50% of the pretreatment condition. Progressive disease (P) was accepted as the relapse of the effusion of more than 50% of the initial fluid. While four of the patients with malignant peritoneal effusion had PR, three patients with pleural effusion showed progression after 2 weeks of mitoxantrone instillation. The durations of responses with peritoneal effusion were 7, 8, 9 and 16 weeks. All patients with pleural effusion died within 2 months. Two patients with peritoneal effusion died within 4 months and two were alive at 4 months with relapse of the effusion (Table 1).

The results of our study do not support the idea^{1,4} that local therapy with mitoxantrone is useful in the

Correspondence to: Ö Özyılkan

Table 1. Primary site of the tumor and response

Age/ sex	Primary site	Site of effusion	Response	Response duration (weeks)
67/F	stomach	peritoneal	PR	8
51/M	stomach	peritoneal	PR	7
35/F	breast	peritoneal	PR	9
60/F	mesothelioma	peritoneal	PR	16
39/M	rectum	pleural	Р	
50/F	lung	pleural	Р	
29/F	breast	pleural	P	_

palliation of malignant pleural effusion. We think local administration of mitoxantrone deserves further investigation.

References

- Torsten U, Opri F, Weitzel H. Local therapy of malignant pleural effusion with mitoxantrone. *Anti-Cancer Drugs* 1992; 3: 17–8.
- Haskell CM. Drugs used in cancer chemotherapy. In: Haskell CM, ed. Cancer treatment. Philadelphia: WB Saunders 1990: 44–102.
- Alberts DS, Peng YM, Bowden GT, et al. Pharmacology of mitoxantrone: mode of action and pharmacokinetics. Invest New Drugs 1985; 3: 101–7.
- Musch E, Loos U, Mackes KG, et al. Intrapleural mitoxantrone in the treatment of malignant pleural effusion. In: Kreidler H, Link KH, Aigner RB, eds. Advances in regional cancer therapy. Basel: Karger 1988: 184-9.

(Received 18 August 1992; accepted 5 November 1992)