

## **CHEMOTHERAPY COMBINED WITH REGIONAL HYPERTHERMIA IN LOCALLY ADVANCED UNRESECTABLE PANCREATIC CANCER: CLINICAL AND ANTHROPOLOGICAL BENEFITS.**

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The latest medical studies on pancreatic adenocarcinoma show a remarkable increase in incidence, prevalently in the western countries.

Recent medical studies have pointed out that there is a correlation between pancreatic tumours and high-fat diets; the influence of coffee intake is debated. In general, obstructive icterus and pain associate to the involvement of retroperitoneal nerve fibres are symptomatic of head and body tumours. Frequently a poor digestion, as a consequence of a bad state of the pancreatic enzymes synthesis, brings about taking off weight. Unfortunately the prognosis of this kind of neoplasia is unfavourable, patients are likely to survive less than 12 months. Many of this kind of neoplasia are inoperable and often a treatment with chemotherapy and radiotherapy is not successful. The treatment with capacitive hyperthermia (HT) shows antitumoral effects associate with a chemotherapy (CHT) treatment consisted of gemcitabine (GEM) alone or in association with oxaliplatin, cisplatin, or 5-FU (1,2,3,4,5). This treatment utilizes a modern and functional apparatus so that the patients have a good ability to tolerate it because of the short incidence of complications.

The aim of this study was to evaluate the action of CHT associated with regional Hyperthermia (HT) tested on a group of 25 patients suffering from locally advance unresectable pancreatic carcinoma (LAPC).

### **MATERIALS AND METHODS**

**APPARATUS:** We used an RH equipment at 13.56 MHz endowed with liquid-cooled flexible antennas.

**PERIOD:** from 02/2001 to 07/2009.

**TREATMENT:** 3 cycles of treatment, every cycle is structured in 8 sessions of 45 minutes each on alternate days.

**PATIENTS:** a group of 25 patients suffering from locally advanced unresectable pancreatic cancer (12 male and 13 female).

## RESULTS

Median overall survival (OS) was 16 months in the group CHT+HT vs 8-11 months as reported in literature. We did not observe adverse effects or increase toxicity in CHT.

1. Survival 12<sup>th</sup> month: 19 pts - 76%
2. Survival 18<sup>th</sup> month: 12 pts - 48%
3. Survival 24<sup>th</sup> month: 11 pts - 44%
4. Survival over 24<sup>th</sup> month: 9 pts - 36%

## CONCLUSIONS

Anticancer nucleoside Gemcitabine, Oxaliplatin and 5 FU have dose limiting toxicities (DLT) Major side effects of Gemcitabine include bone marrow suppression, flu-like syndrome and severe hepatic toxicity.(6,7,8,9,10)

The application of Regional Hyperthermia (HT) on this restricted group of patients has given very interesting results. The HT + CHT can reduce the tumour increase, can increase the survival of the patients and, above all, the HT can improve the general conditions of the patients that have been treated with this kind of combined therapy. The results justified further evaluation in a large number of patients to confirm the benefit.

The Hyperthermotherapy improved the quality of life of all responding patients.

Compared to the severe physical, existential and esthetic impact of the chemotherapy alone, patients with Hyperthermia do not experience particular side effects. As a consequence of that, patients are less anxious in facing the treatment; they establish a fruitful empathic relation with cares and doctors. So, the anguish proceeding the moment of cares (CHT) turns now into a necessary but not threatening and foreboding moment (11,12,13,14,15). Physiotherapy intervention, in its turn, has something to offer throughout the whole cancer journey, also for patients who are not curable and whose life is limited.

## REFERENCES

1. Cunningham D, Chau I, Stocken D, et al. Phase III randomised comparison of gemcitabine (GEM) versus gemcitabine plus capecitabine (GEM-CAP) in patients with advanced pancreatic cancer. European Cancer Conference (ECCO 13), presentation/abstract PS11, Paris, France, 2005 November 2. European Journal of Cancer Supplements 2005;3:4.
2. Reni M, Cordio S, Milandri C, et al. Gemcitabine versus cisplatin, epirubicin, fluorouracil, and gemcitabine in advanced pancreatic cancer: a randomized multicentre phase III trial. Lancet Oncol 2005;6:369-376.

3. Poplin E, Levy DE, Berlin J, et al. Phase III trial of gemcitabine (30-minute infusion) versus gemcitabine (fixed dose-rate infusion [FDR]) versus gemcitabine+oxaliplatin (GEMOX) in patients with advanced pancreatic cancer (E6201). J Clin Oncol (Meeting Abstract) 2006;24:LBA4004.
4. Cartwright TH, Cohn A, Varkey JA, et al. Phase II study of oral capecitabine in patients with advanced or metastatic pancreatic cancer. J Clin Oncol 2002;20:160-164.
5. G. Colucci, R. Labianca, F. Di Costanzo et al: Randomized phase III trial of gemcitabine plus cisplatin compared to single-agent gemcitabine as first-line treatment of patients with advanced pancreatic cancer. The GIP-1 (Gruppo Italiano Pancreas GOIM/GISCAD/GOIRC) study. J Clin Oncol 2010
6. S. Goel, A. Bulgaru, H. Hochster, S. Wadler, W. Zamboni, M. Egorin, P. Ivy L. Leibes, F. Muggia, G. Lockwood, E. Harvey, G. Renshaw & S. Mani; Annals of Oncology 2003; 14: 1682–1687.
7. Stephen A. Welch and Malcolm J. Moore.; Combination Chemotherapy in Advanced Pancreatic Cancer: Time to Raise the White Flag? 2007 Journal of Clinical Oncology by American Society of Clinical Oncology.
8. Richard Herrmann, György Bodoky, Thomas Ruhstaller, Bengt Glimelius, Emilio Bajetta, Johannes Schüller, Piercarlo Saletti, Jean Bauer, Arie Figier, Bernhard Pestalozzi, Claus-Henning Köhne, Walter Mingrone, Salomon M. Stemmer, Karin Tamas, Gabriela V. Kornek, Dieter Koeberle, Susanne Cina, Jürg Bernhard, Daniel Dietrich and Werner Scheithauer; Gemcitabine Plus Capecitabine Compared With Gemcitabine Alone in Advanced Pancreatic Cancer: A Randomized, Multicenter, Phase III Trial of the Swiss Group for Clinical Cancer Research and the Central European Cooperative Oncology Group. 2007 Journal of Clinical Oncology by American Society of Clinical Oncology; Presented in part at the 41st Annual Meeting of the American Society of Clinical Oncology, May 13-17, 2005, Orlando, FL, and the 13th Annual European Cancer Congress, October 31-November 3, 2005, Paris, France
9. Muhammad Wasif Saif, Armin Shahrokni, Daniel Cornfeld; Gemcitabine-Induced Liver Fibrosis in a Patient with Pancreatic Cancer; JOP. J Pancreas (Online) 2007; 8(4):460-467
10. Pantaleoni  
Pharmacology: [http://scholar.google.ch/scholar?hl=it&as\\_sdt=0&as\\_vis=1&q=pantaleoni+pharmacology](http://scholar.google.ch/scholar?hl=it&as_sdt=0&as_vis=1&q=pantaleoni+pharmacology)
11. F. Gabrielli, G. Camurati, M. Iannò, F. Mauro, Patterns of suffering: an Anthropological Reading. New Medicine, XIV, 2, 2010: 63-65.
12. F. Gabrielli, M. Iannò, a cura di, Del limite. Pagine di filosofia e medicina, Ludes University Press, Lugano 2010.

13. M. Cocchi, L. Tonello, F. Gabrielli, M. Pregolato, Depression, Osteoporosis, Serotonin and Cell Membrane Viscosity between Biology and Philosophical Anthropology. *Annals of General Psychiatry* 2011, 10:9doi:10.1186/1744-859X-10-9.
14. M. Cocchi, L. Tonello, F. Gabrielli, Intech Platelet fatty acids membrane viscosity depression and ischemic heart disease biological molecular path with medical anthropology insights, cap. 17, in *Intech – Coronary Angiography. Advances in Noninvasive Imaging Approach for Evaluation of Coronary Artery Disease*, edited by Branislav Baskot, 2012: 315-352.
15. F. Gabrielli, *Philosophy and Psychiatry. The violated body in the era of the invisible man.* *NeuroQuantology*, June 2012, Volume 10, Issue 2, | Pages S1-28: 19-20.