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Histologically, seven cases were pure ESCCs and one was combined with adenocarcinoma. Immunohistochemical studies were performed in three cases, and in one of them, electron microscopy in addition. Only two patients were women. The ages of the eight patients ranged from 15-70, with a median of 62.5 years. Primary tumor locations were: colorectal (2), upper aerodigestive tract (2), urogenital (2: 1 prostatic), peritoneal (1), presenting one of the patients extensive abdominal disease of unknown primary origin. Five limited (LD) and three extensive diseases (ED) were found. In our series, initial treatments used, were chemotherapy (CHT) alone in five cases, surgery plus CHT in one, surgery plus radiotherapy (RT) in one, and surgery alone in one. The results were six complete remissions (CR) and two progressions (P). Overall survival median (from diagnosis), was 17 months (R: 1-68), being 20 months (R: 1-68) for patients with LD, against 16 months (R: 6-18) for the group with ED. There were two long term survivals (>2 years), which were treated with surgery plus RT and surgery plus CHT respectively.

Because of the low incidence of ESCC, and the responses and survivals observed by us and in other reports, it would be necessary the achievement of multicentric trials to standardize therapy.

PUBLICATION PUBLICATION

## CHEMOTHERAPY VERSUS CONCURRENT RADIO-CHEMOTHERAPY SCHEDULE IN SMALL CELL LUNG CANCER LIMITED DISEASE

S. Smickoska, D. Jovanovski

Institute of Radiotherapy and Oncology, Faculty of Medicine, University St. Kiril & Metodij, Skopje, Macedonia

Out of 72 patients with small cell lung cancer (SCLC), limited disease 33 have been treated with chemotherapy only according to protocol 1 (HEMO) and 39 have been treated with concurrent radio-chemotherapy according to protocol 2 (KORHA). Analyzing according to the therapeutic protocols, the objective response has been achieved with 26 (79%) patients in HEMO group and 37 (95%) patients in KORHA therapeutic group. The objective response rate has been statistically significantly higher in KORHA group compared with HEMO group (P < 0.05). The individual statistical analysis of complete and partial remissions has shown statistical significantly higher rate of complete remissions in KORHA radiotherapeutic group (P < 0.05) compared with the HEMO group. The analysis of survival curves of the therapeutic groups has demonstrated that statistically significant difference is existing in survival among HEMO group on one side and KORHA group on the other side, (P < 0.05) in favour of radiotherapeutic group (median and two-years survival 10 months and 11%, versus 16 months and 32%). Therefore, the inclusion of thoracic irradiation in combined treatment of examined patients with SCLC having limited disease has resulted with statistically significant improvement of survival compared with patients treated with chemotherapy only.

## 98 PUBLICATION PROSPECTIVE STUDY WITH CARBOPLATIN, VINCRISTINE, ETOPOSIDE I.V., AND ETOPOSIDE IN PROLONGED ORALLY DOSAGE, AS FIRST LINE CHEMOTHERAPY IN SMALL CELL LUNG CANCER

J.J. Reina, A. Rueda, J.C. Valenzuela, M. Valladares, J.A. Contreras, L. Iglesias, M. Noguer, P. Pastor, J.A. Moreno

Service of Medical Oncology, Virgen del Rocio Universitary Hospital, M. Siurot Av., Sevilla, Spain

The combination of etoposide and cisplatin now appears to have the best therapeutic index of any combined chemotherapeutic regimen used in small cell lung cancer (SCLC), and it allows the addition of some "thirds drugs" of probed efficacy, e.g. vincristine (V), without prohibitive hematologic toxicity. The use of carboplatin (C) instead of cisplatin is eventually preferable because it has fewer side effects with a similar efficacy. Etoposide (E) is the most active drug as single agent, and has been demonstrated the influence of dosage regimens in therapeutic results in SCLC. We report a prospective study with C, V, intravenous E, and E in prolonged orally dosage, as first line chemotherapy (CHT) in SCLC.

Seventy-seven patients (pt) with histologically confirmed diagnosis and a minimum performance status (PS) value (ECOG) of 2, were included from November 1991 to November 1994. Pretreatment evaluation consisted of complete medical history and physical examination, complete blood counts, liver and kidney function tests, chest roentgenograms, CT scans of chest and abdomen, bone marrow aspiration and biopsy, and fiberoptic bronchoscopy. The planned treatment consisted of C 350 mg/m<sup>2</sup> LV. day 1, E 100 mg/m<sup>2</sup> day 1, V 1.4 mg/m<sup>2</sup>

I.V. day 1, and E 50 mg/24 hr days 2-2, repeated every 28 days for six courses. Twelve pt were considered not evaluable. Six pt received less than two courses. Three pt were lost to follow-up. Two pt were wrongly included. One pt was wrongly diagnosed. Sixty five pt were evaluable, sixty three men and two women. The ages ranged from 38-75, with a media of 60.2 years (median: 62). Staging was performed according to VALg criteria: 40 (61.5%) cases of limited disease (LD) and 25 (38.5%) extensive disease (ED). We obtained an overall remissions rate of 78.5%, with 21 (32.3%) complete remissions (CR) and 30 (46.2%) partial remissions (PR). The median survival time was 300 days in patients with LD and 240 in those with ED, with an overall value of 251.5 days (R: 30-1020). Forty-five percent of the pt with LD got CR, and 27.5% PR. Twelve percent of pt with ED got CR, and 76% PR. There were five long-term survivals (>2 years). There was 1 toxicity secondary death (febril neutropenia). Observed grades 3 and 4 toxicities (WHO) were: anemia 21 pt (32.3%), leucopenia 10 pt (15.4%), thrombocytopenia 6 pt (9.2%), nausea-vomiting 2 pt (3.1%), alopecia 8 pt (12.3%), and neurological 1 pt (1.5%)

In conclusion, in our experience, this is an effective regimen in SCLC treatment, and also with an acceptable toxicity (especially alopecia and nausea-vomiting).

POSTER

## THE EFFECT OF CONTINUING SMOKING ON LATE RELAPSES AND SECOND PRIMARY CANCERS IN LONG-TERM SURVIVORS WITH SMALL-CELL LUNG CANCER (SCLS)

F. Cardenal, M. Garcia, X. Castellsagué, X. Garcia del Muro, A. Montes, J.R. Germà

Departments of Medical Oncology and Cancer Epidemiology and Control, Hospital Duran i Reynals, L'Hospitalet, Barcelona, Spain

From 1981 to 1992, 338 consecutive patients with untreated SCLC received standard therapy at our Institution. Of the 338 patients, 23 (6.8%) were alive and cancer-free two or more years from initiation of therapy. The other 315 patients died or relapsed with SCLC before two years. Follow-up time from initiation of therapy is 2.1 to 1.1 years (median 4.6) for the 23 patients. Six of the 23 patients (26%) developed recurrent SCLC 2.23 to 4.6 years (median 3.4) after beginning therapy. Four of the 23 patients (17%) developed one second primary cancer (SPC) 2.5 to 7.7 years (median 3.9) after initiating therapy. SPC diagnoses included: non-small cell lung cancer, gastric adenocarcinoma, squamous cell carcinoma of the floor of the mouth and carcinoma metastatic, to the neck of occult primary site. The cumulative actuarial risk of developing recurrent SCLC at five years is 39%. The cumulative actuarial risk of developing SPC at 5 and 8 years is 20% and 40% respectively.

Five patients continued to smoke after initiating therapy for SCLC, and 18 did not. Four of those who continued to smoke redeveloped cancer: recurrent SCLC (Three patients) and carcinoma of the floor of the mouth (one patient). As compared to patients who stopped smoking the Cox relative risk for developing recurrent SCLC or a SPC was 2.2 (95% CI, 0.6–8.4) in patients who continued to smoke (log rank test, P = 0.22). The corresponding relative risk for SCLC recurrence was 2.9 (95% CI, 0.6–14.5; log rank test, P = 0.17).

These data are suggestive of an increased risk of late relapses and SPC in 2-year cancer-free survivors with SCLC who continued to smoke after initiating therapy, although relative risks do not reach statistical significance. A study with larger number of cases is warranted.

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## RANDOMIZED COMPARISON OF ETOPOSIDE-CISPLATIN VS ETOPOSIDE-CARBOPLATIN AND IRRADIATION IN SMALL CELL LUNG CANCER (SCLC): EVALUATION OF LONG TERM SURVIVAL

E. Samantas, M. Milonakis, G. Klouvas, A. Panoussaki, M. Agelidou, F. Palamidas, E. Boleti, E. Kosmas, E. Papadakis, N. Kiamouris, T. Vardoulakis, P. Kosmidis, N. Pavlidis, G. Foutzilas, D.V. Skarios Hellenic Cooperative Oncology Group (HeCOG), Greece

Between 2/88 and 5/91, 143 patients (pts) with SCLC (82 extensive and 61 limited disease) were randomized to receive Cisplatin and Etoposide (PE) or Carboplatin and Etoposide (CE). Responders, received concurrently with the 3rd cycle of chemotherapy chest irradiation, while CR's received additionally prophylactic cranial irradiation. No statistically significant difference was observed in the response and survival between

the two groups (Sem. Oncol. 21:23-30, 1994, suppl.6). A recent analysis of long term survival pts, showed that 10 pts of the PE group and 9 of the CE group lived more than 2 years. Patients were similarly stratified according to the stage of the disease, PS, sodium, alkaline phosphatase and albumin. Results:

 Response
 CR
 Survival (months)
 TTI (months)

 PE
 90%
 80%
 45.7 (28.1–69.0)
 43.90 (4–69)

 CE
 100%
 67%
 48.4 (27.6–76.7)
 23.48 (3–77)

None of the above differences was found to be statistically significant. Eleven pts have died so far from SCLC and 1 pt from NSCLC respectively. Seven pts (5 PE, 2 CE) are still alive from 42+ to 77+ months.

In conclusion it appears that the less toxic Carboplatin is at least equally effective to Cisplatin as far as it concerns response, duration of responses and long term survival, in pts with SCLC.