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France, Germany, Italy and Netherlands, and is supported by experts in information retrieval and analysis, evaluation and software development.

The consortium is developing a tool (COLLATE), based on emerging information and communication technologies. COLLATE has been designed to achieve two main goals: (a) to support networks of clinicians collaborating on clinical guideline development (involving activities such as the collection, analysis and systematic review of evidence, and consensus forming); (b) to support the dissemination of guidelines and promote their use in clinical practice by making them available to clinicians over the internet.

The impact of COLLATE on guideline development and dissemination is being assessed initially in the diagnosis and treatment of breast cancer. Evaluation is concerning on such aspects as ease of use, the time taken to complete each step of the process, resources used and the quality of guidelines produced. COLLATE is to be demonstrated at the conference, and will be available for comment and discussion by attending clinicians.

Use of COLLATE is expected to produce benefits for clinicians and patients including reducing variations in clinical practice for similar conditions; promoting the appropriate use of medical interventions; speeding up the transfer of state-of-the-art clinical research results into daily practice; improving quality, consistency in clinical care.

362 PUBLICATION

Angiosarcoma of the breast after lumpectomy and radiotherapy for early breast cancer: The key to an early diagnosis

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Secondary angiosarcoma is a rare disease. In recent years, post-treatment angiosarcomas of the breast are described more often. Most reports deal with therapeutic options and bad prognosis. The prognosis is determined by early diagnosis which is often very difficult.

The cases reported in literature and 4 cases in our department are reviewed with special emphasis on diagnosis. The tumor characteristically presents as a painless mass in the breast often accompanied by blueish, reddish, purple or even black skin discoloration. This may be confused with mastitis carcinomatosa in an irradiated breast or with post radiotherapy sequeliae. Mammography, ultrasound, fine needle aspiration cytology and MRI have a low diagnostic sensitivity. Incisional biopsy, including discolored skin and underlying tumor is the most accurate way for diagnosis.

Since the only curative treatment seems to be radical surgery, no valuable time should be lost with repeated and confusing diagnostic procedures. When the clinical picture is present, even though imaging does not provide further information, an incisional biopsy provides the fastest way to early diagnosis and treatment.

363 PUBLICATION

The effect of the interval between surgery and radiotherapy on local control and overall survival in patients with breast cancer

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Purpose: We analyzed the effect of the interval between surgery and radiotherapy (RT) on local control and overall survival in patients (pts.) with breast cancer.

Material and Method: 672 pts. who were treated with postoperative RT after mastectomy and axillary dissection were analyzed retrospectively. Median age was 48 (21–80) years. Postoperative RT was given to all of the pts. (46–65 Gy, Co60-Linak.) through peripheric lymphatic and chest wall portals. 56% of the pts. received RT and 44% RT+chemotherapy (CT). In 68 pts. FAC and in 230 pts. CMF regimens were used. In 196 pts CT was done after irradiation, and in 102 pts. sandwich method (CT + RT + CT) was used. Median follow up was 60 (36–196) months. RT was started in % 61 of the pts. within 8 weeks (group A), and in 39% of the pts. between 8–20 weeks (group B).

Results: Group A and B were similar according to menopausal status, tymph node metastasis and CT apply. T3 tumors were more in group B (p < 0.05). Five and ten year local disease free survivals were 89%, 84% in group A and 84%, 84% in group B. Five and ten year overall survivals were 67%, 51% in group A and 64%, 54% in group B. There were no significant differences in locoregional and distant recurrence incidence, locoregional disease free survival and overall survival between two groups.

Conclusion: The results suggest that administering RT. within 8-20 weeks after mastectomy may not effect local control and overall survival:

Proffered Papers

364 PUBLICATION

Modulation of tamoxifen adjuvant hormonal treatment according to first generation prognostic factors in 702 pT1−2/pN0−1(≤3)/M0 breast cancer, treated by surgery and external Irradiation

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From 1982 to 1992, 702 women under 75 years, without any previous carcinoma, suffering from a unilateral invasive breast cancer classified as pT1-2/pN0-1/M0 were enrolled: 84.3% underwent a breast conservative treatment and 15.7% a modified radical mastectomy. There were 71.7% pT1, 28.3% pT2; 62.4% of the patients were menopaused. Histological axillary lymph node status, Scarff-Bloom and/or cytological grade, ER content were used to set up three groups of patients: 400 were pN0, grade 1-2, ER+ (group 1), 113 were pN0, grade 3, ER+ (group 2), 189 were pN+ ≤ 3, grade 1-2, ER+. Patients from the latter 2 groups received tamoxifen 20 mg/day/2 years, with a pelvic irradiation (12 Gy), should they be not menopaused. In the three groups, the median age is ranging from 51 to 59 (28-75) (p. = 0.07) and the median participation time from 55 to 60 months (1-178) (p = 0.47). 5-year outcomes were assessed with Kaplan Meier method and log-rank test. Overall survival: 97/98/97% (p = 0.39), disease free survival: 85/90/91% (p = 0.08), local-regional free survival: 93/94/98% (p = 0.01), metastatic free survival: 90/93/92 (0.91). As compared to group 1, tamoxifen modifies significantly the outcomes of patients grade 3 or pN1 ≤ 3. Other results are detailed according to age, pT stages, menopausal status, treatment with a Cox multivariate analysis.

## Biotherapy-gene therapy-vaccination

66 ORAL

Interleukin 12 supports Immune responses towards human breast and ovarian carcinomas following initial CD80 mediated T cell activation

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Introduction/Purpose: One explanation for the poor immunogenicity of tumors is the induction of peripheral tolerance by tumor cells that fail to deliver costimulatory signals. Furthermore, T cells stimulated with wild type tumor cells often fail to secrete cytokines. The present study has been undertaken to identify cytokines that cooperate with CD80 in T cell activation towards human breast (MaCa) and ovarian carcinoma (OvCa) cell lines.

Methods/Results: We developed culture conditions enabling us to modulate immune responses towards CD80 transfected MaCa and OvCa by IL-7 and IL-12. IL-7 amplified the proliferative response towards CD80 transfected MaCa and OvCa but stimulated predominantly CD4\* T lymphocytes. IL-12 represses the proliferative response of naive T cells in primary activations. However, during rechallenge stimulations IL-12 cooperates with CD80 mediated activation. In long-term T cell cultures IL-12 synergizes with CD80 expression in stimulation of CD8\* T cell lines which recognize the parental MaCa line in a HLA-restricted manner.

Conclusion: We showed that CD80 expression is neccessary for tumor cells to function as (allo)antigen presenting cells. Following initial CD80 mediated activation IL-12 supports the propagation of CD8\*, HLA-restricted T lymphocytes. Tumor-reactive T cell lines would improve efforts aimed at identifying tumor specific antigens. In clinical approaches immunogenic tumor variants could be directly used for the vaccination of HLA-matched MaCa and OvCa patients.