

for lymphoma patients with chemotherapy (CHOP, ABVD) and lung cancer patients with radio and/or chemotherapy – 1 injection followed by 1 injection every 5 days (total of 5 injections). Mesothelioma patient received two courses of 2 mg of Liasten every two weeks and 3 capsules Del-Immune b.i.d. Effect was measured on the basis of blood test results and survival rate.

**Results:** Striking decrease were found in incidence of neutropenia complications (60% versus 21%,  $p=0.0032$ ) in those patients who received chemotherapy with liasten. Postoperative cases acute conditions of chronic bronchitis, infectious pneumonia cases occurred in 25% of LC patients versus 40% in control group. Patients that received radiation therapy in combination with muramyl peptides preparations had substantial reduction of toxic side effects. There was no hospital mortality. Increased activity of T-cell immunity and plasma interferon was reported in the process of treatment. Tumor process development was consistent with the control group. There were no cases of malignant lymphoma progression. No relapses and 3.5-fold NK functional activity as well as improved activity of T3 and B-cells were reported by mesothelioma patient more than 3 years after the treatment.

**Conclusions:** It was possible to develop algorithm of using Liasten injections for prevention and treatment of chemotherapy-induced hematotoxic complications in malignant lymphoma and lung cancer. Because of the incurable natures of mesotheliomas, it thus seems warranted to further research the use of Liasten in combination with oral muramyl peptides preparation Del-Immune V® to extend the life expectancy of these patients.

#### 1053 POSTER

##### Development of humanized monoclonal single chain antibodies, against the tumour suppressor interferon regulatory factor 1 (IRF-1), through phage display

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**Background:** Despite a potentially important role in cancer suppression, detailed knowledge of IRF-1 pathways has been hampered by a lack of biochemical tools, and in particular monoclonal antibodies, due to low immunogenicity, poor expression, and toxicity of the IRF-1 protein in *E. coli* and mammalian cells.

**Materials and Methods:** Preformed *in vitro*, phage display circumvents the need for immunogenicity, and was therefore particularly well suited to generating IRF-1 specific antibodies. Additionally, once antibodies capable of binding specifically to IRF-1 were selected, the phage continued to act as a genetically stable source of the antibody which could be stored over a long period. The antibody genes were extracted and moved into a variety of plasmids which allowed for higher level expression in *E. coli*, and *in vivo*, expression in a variety of human cancer cell lines.

**Results:** The single chain antibodies were raised against functional domains of IRF-1 spanning the length of the entire protein to ensure that a range of antigens were targeted. By expressing the antibodies that target functional domains *in vivo*, it may be possible to influence specific activities of IRF-1 within the cell, such as its ability to bind DNA or become transactivated.

**Conclusions:** In this way, single chain antibodies can be used to tease apart the IRF-1 pathway and determine the functional relevance of identified intracellular interactions.

#### 1054 POSTER

##### The existence of humoral immunity to gliadin and cow's milk proteins in patients with prostatic diseases

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**Background:** The goal of this study was to determine the incidence of the presence of serum IgA and IgG antibodies to gliadin and IgA, IgG and IgE antibodies to cow's milk proteins by ELISA test, in patients that are having different diagnosis of malignant or nonmalignant prostatic diseases and in control group of healthy people.

**Patients and Methods:** Twenty-six patients with various diagnosis of prostatic diseases (carcinoma, benign prostatic hyperplasia, adenoma) were included in this research. Nine patients had prostate-specific antigen (PSA) level less than 4 ng/ml, four of them had PSA level in range 4–10 ng/ml and thirteen patients had PSA level more than 10 ng/ml. Fifty healthy people was control group.

Two kinds of antigens were used: skimmed pasteurized cow's milk powder (ICN) and crude gliadin (Sigma). Determination of IgA and IgG serum's

immuno-reactivity to gliadin, or IgA, IgG and IgE to cow's milk proteins (CMP), has been performed by home made ELISA tests. The cut off value, for each test, was evaluated as the mean +2 SD of control group.

**Results:** Statistical analysis of obtained data reveals that the levels of anti-gliadin IgA and anti-CMP IgE were significantly higher in patients with prostatic diseases than that of controls ( $p < 0.008$  and  $p < 0.02$ ). Anti-gliadin IgG and anti-CMP IgA immunoreactivities were not significantly higher in patients, comparing to the control group. The level of anti-CMP IgG immunoreactivities in patients with prostatic diseases comparing to the control group was on the limit of statistical significance ( $p = 0.0543$ ).

**Conclusion:** Results from this study, point to the non-specific association between immunity to food proteins (gliadin and cow's milk proteins) and prostatic diseases.

#### 1055 POSTER

##### Expression of cancer/testis tumor antigens MAGE-A1, MAGE-A3/4 and NY-ESO 1 in medullary breast cancer

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The medullary breast carcinomas (MBC) account for <2% of breast invasive carcinomas. Recent publications on breast cancer classifications based on gene expression profile analyses indicate that MBC may be considered part of the basal-like carcinoma spectrum. As regards this uncommon type of invasive breast cancer we have recently published an article on the clinicopathological features of MBC in 48 patients who were operated on at our two hospitals between 1999 and 2005 (Matkovic B et al. Tumori 2008).

The present study includes immunohistochemical analyses of the expression of cancer/testis (C/T) antigens MAGE-A1, MAGE-A3/4 and NY-ESO 1 in these MBC samples. C/T genes are normally expressed in germ line cells. However, they may also become activated in a wide range of cancer types. Although a study of the expression of these C/T antigens in breast cancer was in part conducted in invasive ductal carcinomas of no special type (NOS), this has not been done with respect to special and/or relatively rare histological types of breast cancers (Hofmann O et al. PNAS USA 2008; Kavalari R et al. Virchows Arch 2001).

In the present study monoclonal antibodies "77B", "57B" and "B9.8.1" (Juretic A et al. The Lancet Oncol 2003) were used to immunohistochemically determine the expressions of, respectively, MAGE-A1, MAGE-A3/4 and NY-ESO-1 C/T antigens in MBC tissues. MAGE-A1, MAGE-A3/4 and NY-ESO-1 antigens were found to be expressed in 33% (16/49), 33% (16/49) and 22% (11/49) of patients, respectively. Immunohistochemical data concerning these C/T antigen expressions were correlated with the following MBC clinicopathological features: patient's age at diagnosis, type of operation, tumor size, axillary lymph node metastasis, adjuvant therapy, ER, PR, HER-2 expression, and patient's survival. No significant correlation between the above-stated clinicopathologic parameters and the antigen expression was identified. The only exception was the patients' survival data which indicate a possible association between the expression of these C/T antigens and decreased overall survival: MAGE-A1  $P = 0.07960$ , MAGE-A3/4  $P = 0.01088$ , NY-ESO-1  $P = 0.11742$ .

The results of our retrospective study suggest that the aforementioned C/T antigens may be used in MBC as tumor markers of potential prognostic relevance. Due to the relative rarity of this type of breast cancer, further tests need to be performed on additional MBC tumor samples with respect to the expression of these C/T antigens before being able to definitely confirm this possibly original observation.

#### 1056 POSTER

##### Cd274/Pdc111 as a genetic modifier controlling aggressiveness of T-cell lymphoblastic lymphoma

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**Background:** The use of consomic and congenic mouse-strains has greatly facilitated the identification of tumour modifier genes. Using sub-congenic interspecific mice generated between SEG/Pas and C57BL/6J strains, we report a critical region on chromosome 19 (*Tlyr1c*) which does