Paediatric oncology Tuesday 23 September 2003 S195

**Conclusion.** The use of aggressive multimodal therapy (surgery/chemotherapy) and evaluation of prognostic factors are necessary for successful treatment in patients with osteosarcoma relapse. Type of chemotherapy regimen should be assessed in randomized multicentric trials.

650 POSTER

## Childhood cancer incidence in the Kyrgyz Republic 1985-1999

E. Makimbetov. Kyrgyz-Russian Slavic University, Special Clinical Disciplines. Bishkek. Kyrgyzstan

**Background:** The study analyses geographical variations in cancer incidence, trends in survival, health status of long-term survivors of childhood cancer.

Material and methods: The incidence of cancer in children aged less than 15 years between 1985 and 1999. There were 1755 registered cases with new diagnoses of cancer. Male-female proportion 1.3. The study collected data from forms submitted along with histological or cytological findings, and deaths certificates. Histologically verified were 84.7% cases.

The population relative risk was estimated for main tumours in urban and rural areas. Detailed population figures from census have been available at the Kyrgyz National Centre of Statistics. The official estimates have been available for intercensus years. These are based on the census and data on natural population change. The population figures and cancer incidence rates for this report have been provided in for age-groups (0-4, 5-9, 10-14), ethnic groups, for each sex and calendar years of the study period. The crude and age-standardised rates per 1 million have been counted. The classification scheme was used according to the International Classification of Childhood Cancer.

**Results:** Total age-standardised (ASR) annual incidence rate was 74.8 per million. The most frequent diagnostic groups were leukaemias (ASR-20.8), brain and spinal tumours (7.3), lymphomas (5.4), sympathetic nervous system tumours (4.3), kidney tumours (5.0), soft-tissue sarcomas (4.4), retinoblastomas (3.9), germ cell tumours (3.3), hepatic tumours (1.2), carcinomas and non classified tumours (4.6).

Incidence was significantly higher in the Russians, with an ASR of 114.7 cases per million per year, compared with 77.6 for Uzbeks and 65.6 Kyrgyz children. Assigned risk was higher among Russians in lymphomas (RR= 3.5), neuroblastomas (RR=2.5), than Kyrgyz. Kyrgyz and Uzbeks had an increased RR in tumours of eye and testis, but this was not statistically significant. Part of this apparent increase may have been due to the improved registration of cases in the 1980s, particularly of certain tumour types, and of tumours in older children who may have been treated outside the major paediatric treatment centres. Between the 2 periods, 5-year survival increased from 29% to 47%, with improvements in almost all tumour types. In some tumours (Wilm's, soft tissue sarcoma, leukaemias, lymphomas and others) 5-year survival increased from 45% to 64%. There has been an evidence that with new treatment regimens introduced during the 1980s survival in the 1990s has shown the increased gains.

**Conclusions:** In Kyrgyzstan, the childhood cancer incidence is low and similar to those reported from Asian developing countries. The data could be used for a wide range of epidemiological and other studies.

651 POSTER

## Blood and bone marrow transcripts of tyrosine hydroxylase, dopa decarboxylase, and GD2 synthase in neuroblastoma

A. Kullman<sup>1</sup>, C. Träger<sup>1</sup>, P. Kogner<sup>2</sup>, B. Kågedal<sup>1</sup>. <sup>1</sup> Department of Biomedicine and Surgery, Division of Clinical Chemistry, Linköping, Sweden; <sup>2</sup> Karolinska Institutet, Childhood Cancer Research Unit, Department of Woman and Child Health, Stockholm, Sweden

Background: Neuroblastoma is the most common extra cranial tumor in childhood. In spite that multimodal therapy is used the outcome for children with severe neuroblastoma is still poor. The mRNA transcripts of tyrosine hydroxylase (TH) and GD2 synthase (GD2S) are two well-known candidates suitable for identification and monitoring neuroblastoma cells in blood and bone marrow. Dopa decarboxylase (DDC) mRNA also seems to be useful, but has not been systematically investigated. The utility of these transcripts for tracing minimal residual disease should therefore be compared in clinical

Material and methods: Real-time reverse transcription (RT)-polymerase chain reaction (PCR) methods were developed to quantify mRNA of TH, DDC and GD2S in blood and bone marrow in children with neuroblastoma. The calibrators used were obtained by amplification of segments of cDNA from the mRNAs of the enzymes, which included the targets.

A total of 229 blood samples from 47 patients and 106 bone marrow samples from 41 patients with neuroblastoma of different stages were analyzed. Cord blood from 52 babies and blood from 26 healthy children, 4 months to 16 years of age, were used as controls.

**Results:** Blood samples were obtained at diagnosis from 11 children with a stage 1-3 disease. Increased concentrations were found in 4 children regarding TH mRNA, 1 child regarding DDC mRNA and 3 children regarding GD2S mRNA.

Bone marrow samples were also received from 9 patients with stage 1-3 neuroblastoma. Two of them had TH mRNA values above cut off level, one had increased DDC mRNA and five had increased GD2S mRNA.

We received blood samples from 14 children with stage 4 disease at diagnosis. TH mRNA was increased in 13 of the cases, DDC mRNA in 7 cases, and GD2S mRNA was elevated in 10 of the cases.

In 15 children with stage 4 neuroblastoma there were increased bone marrow concentrations of TH mRNA in 14 cases, of DDC mRNA in 12 cases and of GD2S mRNA in 13 cases.

There was a high correlation between TH mRNA and DDC mRNA in blood taking all samples into account. This was not the case between GD2S and TH. The correlations were lower in bone marrow between DDC and TH but still higher than the correlation between GD2S and TH mRNA.

**Conclusions:** The data indicate that at diagnosis of neuroblastoma sensitivity for detection of stage 4 disease is higher with TH mRNA as compared with both DDC mRNA and GD2 mRNA. It also seems that GD2S is less discriminating of stage 4 from stage 1-3 in the bone marrow since 5 of 9 children with stage 1-3 disease had elevated levels in the bone marrow.

S52 POSTER

## Severe sepsis in children treated for leukemia in a single unit in Romania

C.L. Blag<sup>1</sup>, G. Popa<sup>2</sup>, N. Miu<sup>3</sup>, M. Giurgiulescu<sup>4</sup>. <sup>1</sup> Pediatric Clinic no 2, Pediatric Oncology, Cluj-Napoca, Romania; <sup>2</sup> Pediatric Clinic no 2, Pediatric Oncology, Cluj-Napoca, Romania; <sup>3</sup> Pediatric Clinic no 2, Head of Clinic, Cluj-Napoca, Romania; <sup>4</sup> Pediatric Clinic no 2, Pediatric Oncology, Cluj-Napoca, Romania

**Purpose:** This is a retrospective study aimed to determine the etiology and leading causes of death in severe sepsis related to leukemia in a country with limited financial resources.

Patients and methods: The records of 51 children treated for acute leukemia (ALL 44 pts and AML 7 pts) between 1991-2003 were retrospectively reviewed for severe sepsis. Data concerning clinical, microbiologic and other investigative documentation of infection and outcome were analyzed using the SPSS software programme.

Results: Forty five episodes of severe sepsis were documented in 27 patients. All were under chemotherapy: induction phase (6 episodes), consolidation (34 episodes) and maintenance (5 episodes). Etiology was precised in 58% of episodes: 77% bacterial and 11,5% viral and fungal, respectively. The commonest bacteria was Pseudomonas aeruginosa (23% episodes). Of three fungal infections, 2 were systemic aspergillosis and 1 candidiasis. Severe viral infections were varicella (2 pts) and CMV reactivation (1pt).

All severe septic episodes were determined by neutropenia and its degree of severity significantly correlated with mortality (p 0,05).

The addition of G-CSF in therapy had no significant contribution to reducing fatalities (p 0,58).

Mortality rate by sepsis was 19% and documented bacterial infections were 6: Shigella (2 cases) and Pseudomonas, Acinetobacter, Haemophilus and Staphylococcus aureus (1 case each).

Conclusions: Gram negative infection was the leading cause of morbidity and mortality during treatment of leukemia. Severity of neutropenia was the most important factor predicting fatality.

653 POSTER

## Epidemiology of childhood cancer in Bihor and Timis counties during the years 1981-2000

G. Miculschi<sup>1</sup>, M. Serban<sup>2</sup>, L. Ritli<sup>1</sup>, R. Spineanu<sup>3</sup>. <sup>1</sup> Clinical Children Hospital, Oncohematology, Oradea, Romania; <sup>2</sup> Paediatric Clinic III, Oncohematology, Timisoara, Romania; <sup>3</sup> Clinical Children Hospital, Paediatrics, Oradea, Romania

**Background:** the objective of this study was the epidemiologic study of childhood cancer in Bihor and Timis counties in order to describe the extend and nature of the cancer burden and to assess the effectiveness of cancer control activities in this part of the country.