

In this issue

Increased survival rates for osteosarcoma of the extremities

Adjuvant and neoadjuvant chemotherapy, introduced in the early 1970s, have significantly improved the long-term survival rate (from 20% to 60%) for patients with high-grade osteosarcoma of the extremities. Simultaneously, the frequency of limb salvage surgery has increased with a corresponding decrease in amputation rates. However, according to Bacci and colleagues, the 5-year event-free survival (EFS) rates calculated in many studies are based on cases where follow-up was often less than three years. Also most studies have not reported on either the post-relapse treatments with patient outcome after developing metastases/local recurrence; or details of the surgical treatment, margins and complications. In this issue of *EJC*, Bacci and co-workers rectify this deficit by reporting the long-term results achieved in a large series of patients in a single institution. Patients (1148) with non-metastatic osteosarcoma of the extremity were treated with 4 different protocols of adjuvant and 7 different protocols of neoadjuvant chemotherapy. In the period from 1972 to 1999 the rate of limb salvage increased from the 20% to 71% and the 5-year EFS and overall survival (OS) were 57% and 66%, respectively. The 10-year EFS and OS were 52% and 57%, respectively. These results significantly correlated with serum alkaline phosphatase levels; the type of chemotherapy (adjuvant *vs.* neoadjuvant); and with histologic response to pre-operative treatment.

Childhood cancers: a common aetiology?

The aetiology of most childhood tumours remains unclear. Genetic predisposition is reported to be directly associated with about 5% of cases, whilst environmental exposure or host response to such exposure (genetically determined) is proposed for the majority. If environmental factors are involved, childhood cancer cases might be expected to exhibit a non-random geographical distribution. Space-time clustering, as seen in infectious diseases, occurs from irregular distribution of cases simultaneously both in time and space and is indicative of a common environmental aetiology. Space-time clustering has previously been shown in cases of childhood leukaemia, central nervous system (CNS) tumours, soft tissue sarcoma and Wilms' tumour. In this issue of *EJC*, McNally and colleagues continue this work to show significant cross-clustering between cases of leukaemia and CNS tumour and in particular between cases of acute lymphoblastic leukaemia (ALL) and astrocytoma. No cross-clustering was seen with Wilms' tumour and soft tissue sarcomas with any other malignancy. The authors highlight that their results are consistent with a common, possibly infectious, aetiological mechanism for ALL and astrocytoma.

Oestrogen receptor expression in breast cancer

Oestrogens have a key role in breast cancer genesis and progression. The effects of estrogens are transduced by the activation of oestrogen receptors alpha and beta (ER α , ER β). In normal mammary tissues ER β is the predominant receptor. However, in breast tumours a shift to ER α overexpression is normally seen in 70% of cases. The molecular mechanism underlying the upregulation of ER α , a significant initial event in breast cancer biology, is not clear. In this issue of *EJC*, Jarzabek and colleagues have analysed ER α and ER β mRNA and protein expression in 41 primary breast cancers and surrounding control tissues. ER α mRNA levels varied more among samples than ER β ; 70% of tumour samples expressed full-length ER α ; ER α was localised exclusively to the nucleus, whereas ER β was observed to have cytoplasmic and perinuclear expression in both control and tumour samples; unlike ER β , the expression of ER α increased with age; and ER α expression correlated positively with progesterone receptor and negatively with proliferation marker Ki-67. The authors conclude that ER α and ER β are differently regulated in breast cancer where ER α is more deregulated than ER β .