Letters to the Editor

Value of factor VIII related antigen as a means of demonstrating extramedullary haemopoiesis

We have noted the comments of Dr AL Bloom and Dr JC Giddings concerning our use of the localisation of factor VIII related antigen (F VIII RAg) in platelets and megakaryocytes. These are, indeed, valid points and it is accepted that contaminants such as fibrinogen may be responsible for weak activity of their corresponding antibodies in commercial antisera. It should be appreciated, however, that the staining obtained with such preparations is not only intense but appears to be found specifically in megakaryocytes and vascular endothelium. Moreover, histiocytes, which contain, for example, fibrinogen, are not, in our hands, stained for F VIII RAg in tissue sections. Of additional value is the negative staining of Reed-Sternberg cells. Thus, regardless of impurities or minor contaminants, the value of staining for megakaryocytes with these commercial antisera is still considerable; the negative staining of other cell types serves as a practical negative control.

Dr Bloom and Dr Giddings also suggest that the use of monoclonal antibodies could overcome the problems suggested. Notwithstanding the current popularity of such preparations, however, the occurrence of shared epitopes often renders such antisera highly non-specific with regard to the staining of certain cell types and large molecules. For example, numerous and diverse unrelated cells or tissues have recently been shown to stain with monoclonal antisera to granulocyte and mast cell determinant molecules. Haematologists and histopathologists should, indeed, recognise this problem. Particular care should be taken to ensure that monoclonal antisera to F VIII RAg itself do not react with fragments of the molecule shared by other molecules. If megakaryocytes or endothelial cells are to be stained in tissue sections, commercial preparations of polyclonal antisera to F VIII RAg should, we believe, still be quite adequate for routine work.

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References

Pathological examination of cutaneous malignant melanomas

This laboratory deals with large numbers of cutaneous malignant melanomas and new primary cases are currently running at about 100 a year. In the past it has been our practice to assess the histological parameters required for microstaging by evaluating a single central transverse block through the melanoma. This technique shows maximum tumour thickness in 95% of melanomas.

This protocol, while almost certainly adequate for the assessment of thickness, has serious flaws with regard to the assessment of other features. This is particularly true of any regression which may be present in the lesion, since such regression is likely to be associated with areas significantly thinner than found maximally in the lesion. In consequence it is now our practice to block melanomas completely, using numerous transverse sections, generally about 2 mm in thickness. This technique, advocated by Kopf et al, shows striking variations within individual melanomas. Figs. 1 and 2 show significant differences in thickness and levels seen in two of four blocks, the central one showing maximum thickness, but not deepest extent. One of the two also shows an area of ulceration not seen in the other. The presence of areas of significant regression around the edges of nodular melanomas occurs quite regularly and areas of pronounced regression may be seen in one part of the tumour edge and not elsewhere. This regression, if involving more than three rete ridges from the edge of the nodule, indicates that the melanoma must have been originally a superficial spreading variant in which the development of the central nodule and its subsequent stimulation of the host immune response has caused the loss of the superficial spreading element.

The microstaging of melanomas is a complex process requiring the assessment of many parameters. We currently assess: histological type, thickness, level, pigment production, cellular type, number of...