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Immunological Variation in the 'Common Region' of the T Antigens of Polyoma Virus

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SUMMARY

Characterization of a series of mouse monoclonal antibodies (designated α Py C21 to 26) against the region common to the large, middle and small T antigens of polyoma virus has revealed immunological diversities among the N termini of these antigens. Four of the antibodies (α Py C21 to 24) appeared to recognize all the T antigens present in lytically infected cells, but two of them (α Py C25, 26) failed to immunoprecipitate small T antigen either from ³⁵S- or ³²P-labelled cells and recognized only a subset of middle and large T antigens. None of the antibodies recognized the protein kinase activity normally associated with middle T antigen or the 60K mol. wt. antigen-related protein observed in polyoma virus lytically infected or transformed cells. The possible immunodominance of the N termini of the early gene products (as well as VP1) in the natural host of polyoma virus and the observed antigenic heterogeneity are considered.

INTRODUCTION

Previous studies with rat monoclonal antibodies to the polyoma virus early regions (small, middle and large T antigens: ST, MT and LT, respectively) have shown that distinct antigenic species exist *in vivo*, at least in the case of MT antigen (Dilworth, 1982). The experiments described in this report were carried out to investigate the immunogenicity of the viral antigens in their natural host, the mouse, and to produce mouse monoclonal antibodies to use as specific probes in further studies of the viral early proteins, particularly since such a wide diversity of functions and cellular locations have been associated with these viral gene products (Türler, 1980; Dilworth *et al.*, 1986). As a corollary to this work, it could be anticipated that the number of available monoclonal antibodies for assessing the biological roles of polyoma T antigens would be increased and that the reagents would have better properties than the existing rat monoclonal antibodies, particularly with regard to binding to Protein A (Ey *et al.*, 1978).

Two different methods were used for isolating monoclonal antibodies from mice to polyoma viral functions. One involved immunizing mice with SDS-PAGE-purified viral proteins, and the other immunization with a mixed antigen complex obtained by immunoprecipitation with the rat monoclonal antibody α Py C4 (Dilworth & Griffin, 1982). Analysis of the specificity of the antibodies obtained from these experiments showed them to be directed mainly against either the N-terminal 'common' region of the three T antigens of polyoma virus or the major viral capsid protein, VP1. Six common region antibodies, two antibodies to the major capsid antigen VP1 and a mouse anti-rat antibody were cloned. Here we present data on the characterization of the common region antibodies and their use in elaborating diversities among the N termini of the three viral T antigens.

METHODS

Cell cultures. 3T6 Swiss albino mouse cells (ATCC CCL 96) were obtained from the American Type Culture Collection and propagated as monolayers in Dulbecco's modified Eagle's medium (DMEM) with 5% foetal calf

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