

ABSTRACT N° 33

RELATIONSHIP BETWEEN APOPTOSIS AND
CASPASE 3 EXPRESSION IN MALIGNANT GLIOMAS.

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Caspase-3 represents the last step in the pathways to apoptosis. Once activated, it cleaves the inhibitory caspase-activated DNase (ICAD) with activation of CAD and subsequent fragmentation of DNA. The activation of caspase-3 is achieved through both transcriptional and receptorial apoptotic pathways. In many systemic tumors, caspase-3 expression correlates with apoptotic nuclei and also with survival. Caspase-3 has been studied in a series of formalin fixed surgical specimens of 30 gliomas (20 glioblastomas and 10 astrocytomas) by "cleaved caspase-3" (D175) polyclonal antibody (Cell Signaling) and compared with apoptotic nuclei demonstrated by TUNEL technique and morphology. There is a correlation of caspase-3 positive nuclei with apoptotic nuclei, more evident in proliferative areas and less in perinecrotic pseudo-palisadings. Usually the staining is nuclear, but it can be also cytoplasmic or in both cell compartments. This observation is important for the interpretation of the mechanism of caspase-3 cleaving in the cell. This staining, together with the incomplete overlapping of positive nuclei with apoptotic nuclei in perinecrotic pseudo-palisadings suggest caution when using caspase-3 in order to improve the recognition of apoptotic nuclei in tissue sections.

ions that result in the inactivation of proto-oncogenes. have been identified in the past with the various types of mutation of the TP53 gene or early gene alterations identified in astrocytomas include 14ARF, RB1 and PTEN, as proto-oncogenes, such as CD2 and CCND3. In consequence they show allelic losses and gains on 1p and 19q in gliomas with a good response to treatment. The current status with respect to the role of aberrant DNA is presented. In addition, we discuss our findings with respect to