
suppressor proteins, which exert their effects via common target genes, and members of the recently defined BCL-2 family of proteins. The BCL-2 family can be divided into 2 groups based on the type of cell death they induce. The "classical" BCL-2 proteins (BCL-2, BCL-XL, and MCL-1) promote survival and prevent apoptosis in most cells. By contrast, the BH3-only proteins, which include BID, BAD, BIK, NOXA, PUMA, and HRK, all members of the BCL-2 family implicated in apoptosis, lack the BCL-2 homology region (BH) and, thus, cannot directly bind to classical BCL-2 family members. Rather, they may function by forming heterodimers with the former proteins, thereby freeing the classical BCL-2 family members from their anti-apoptotic interactions. Each BH3-only protein is multifunctional and exerts a context-dependent cellular outcome, presumably by associating with different combinations of BCL-2 family members in different cell types. Although BH3-only proteins and classical BCL-2 family proteins regulate apoptosis in opposing ways, they often have complementary functions. The Bcl-2-interacting killer (BIK) gene was discovered in a search for novel apoptosis regulators (Agarwal et al., 1995 d0c515b9f4

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