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Anti-PD-1/CTLA-4 Bispecific Antibody AK104

National Cancer Institute

Source

National Cancer Institute. *Anti-PD-1/CTLA-4 Bispecific Antibody AK104*. NCI Thesaurus. Code C153379.

A bispecific antibody directed against the human negative immunoregulatory checkpoint receptors programmed cell death protein 1 (PD-1; PDCD1; CD279) and cytotoxic T-lymphocyte-associated antigen 4 (CTLA4; CTLA-4), with potential immune checkpoint inhibitory and antineoplastic activities. Upon administration, anti-PD1/CTLA4 bispecific antibody AK104 targets and binds to both PD-1 and CTLA4 expressed on tumor-infiltrating T lymphocytes (TILs), and inhibits the PD-1- and CTLA4-mediated downregulation of T-cell activation and proliferation. This restores immune function and activates a sustained cytotoxic T-lymphocyte (CTL)-mediated immune response against tumor cells. Both PD-1 and CTLA4 are selectively expressed on TILs in the tumor microenvironment (TME) and negatively regulate the activation and effector functions of T-cells. They play key roles in the downregulation of the immune system and tumor evasion from host immunity. Dual checkpoint blockade of PD1 and CTLA4 with AK104 may enhance T-cell activation and proliferation more than the blockade of either immune checkpoint receptor alone.