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Multi-targeted Tyrosine Kinase Inhibitors in the Treatment of Cancer and Neurodegenerative Disorders

Halilibrahim Ciftci,Belgin Sever,Luciano Saso,Rumiana Tzoneva,Valentina Onnis,2024-08-07 Tyrosine kinases which catalyze the transfer of a phosphate from ATP to a hydroxyl group of a tyrosine residue play a plethora of roles in the regulation of diverse functions in normal cells including cell growth motility differentiation and metabolism Moreover they are also associated with oncogenesis Tyrosine kinases are mainly classified as receptor tyrosine kinases and non receptor tyrosine kinases including crucial members Epidermal growth factor receptor EGFR belongs to the ERBB family of receptor tyrosine kinases along with three other closely related receptors namely HER 2 HER 3 and HER 4 EGFR and HER 2 lead to autophosphorylation of the intracellular domain through tyrosine kinase activity and subsequent stimulation of downstream cascade that may result in proliferation suppression of apoptosis metastasis and angiogenesis On the other hand c Abl Abl 1 is a non receptor tyrosine kinase which is also essential in the regulation of several antiapoptotic and proliferative signal transduction pathways They have mainly been identified as important targets for several types of cancer such as EGFR for non small cell lung cancer glioma and colorectal cancer HER 2 for breast and colorectal cancers and Abl for chronic myeloid leukemia One of the major platforms that they have participated in is neurodegenerative disorders such as Parkinson s disease Alzheimer s disease and amyotrophic lateral sclerosis Aberrant activity of tyrosine kinases in particular EGFR and Abl have been reported to induce apoptosis and cell cycle arrest in response to a wide range of stimuli resulting in neurodegeneration and neuroinflammation The main goal of this Research Topic is to identify new and potent effective tyrosine kinase inhibitors to be effective in cancer and neurodegenerative disorders

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