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Myc as a Disease Target Beyond Cancer Jonathan R. Whitfield,Mariano F. Zacarias-Fluck,John Sedivy,Yulia A. Nevzorova,2024-09-02 Myc is a pleiotropic transcription factor fundamental for many cellular processes It is an infamous and highly studied oncogene with around 10 000 articles published in the last decade The majority of these link it to cancer The focus on Myc as a hot target in cancer has led to immense efforts from academia through to pharmaceutical companies being directed towards finding a Myc inhibitor Not surprisingly given its fundamental nature in many physiological processes Myc has also been linked to a large variety of pathologies other than cancer The growing efforts to develop Myc inhibitors as cancer therapies will have important implications for the treatment of these diseases too whether they have a clearly demonstrated aetiology linked with Myc or so far are only at early stages of association with it Interestingly the activation of Myc may also prove helpful in the field of regenerative medicine and reversal of ageing The aim of this Research Topic is therefore to describe non oncological pathologies and conditions to which Myc modulation could be applied These will include conditions that have strong links to Myc and where Myc modulation has already been modelled as well as conditions that so far have more limited data linking them to Myc but where it seems to have a key role Authors will be expected to briefly review the particular disease and what is known to link Myc to it plus the possibility of any clinical trials in the near future

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Regulatory T Cells Ling Lu,Bruce Blazar,Fadi Issa,2022-08-26

Evolution of NK-mediated target recognition under the pressure of physiologic or pathologic stimuli Massimo Vitale, Simona Sivori,Miguel López-Botet,Daniel Olive,2015-03-23

Since their discovery NK cells have come out as potential tools to fight cancer and viruses This finding early urged different groups to study the mechanisms governing NK cell function The identification of the MHC I specific inhibitory receptors i e KIRs NKG2A and certain Ly49 molecules allowed defining rather rapidly how NK cells could avoid self aggression and how they could be directed towards targets that were forced by viral infection or tumor transformation to down regulate MHC I expression In a second time also the repertoire of surface activating receptors addressing NK cytotoxicity towards tumors and pathogens was mostly defined In spite of the first findings however most recent studies may suggest that NK cells and their receptors might not have been evolved to kill tumor targets and perhaps they might have been only partially influenced in their evolution by the need of recognizing viruses Indeed certain NK receptors known to activate NK cell cytotoxicity NKp30 DNAM 1 NKp80 can also participate at regulatory interactions occurring between NK and myeloid cells In addition a peculiar NK cell subset which intensively populate decidua during the first trimester of pregnancy through the engagement of specific receptors and the interaction with decidual DC produce chemokines and pro angiogenic cytokines and induce Tregs Thus in this context NK cells favor decidua vascularization and development of the semiallogeneic foetus in a tolerant environment Viruses have nevertheless played an important role in shaping the NK cell receptor repertoire Several studies have unveiled clues of the evolutionary struggle between these pathogens and NK cells Different NK receptors including NKp46 NKp30 NKp44 NKG2D NKG2C Ly49 and certain KIRs have been demonstrated to recognize virus encoded or virus induced ligands The expression of TLR specifically recognizing microbial products together with the unexpected role of KIR3DL2 in shuttling these products to TLR containing endosomes have also been documented in NK cells On the other side different viral immune evasion molecules have been shown to interfere with the expression of ligands for T or NK cell activating receptors In addition viral

infections can occur in the reproductive stage of life cycle and may represent a serious threat for the species propagation. Thus the control of viruses together with the maintenance of foetus during pregnancy should represent major evolutionary forces in shaping NK receptors. Along this line the NK mediated control of tumors should not be under the same evolutionary pressure as tumors mostly appear later in the life cycle and the recognition of tumor encoded ligands may be less efficient as the NK cell receptors might have not been selected for such aim. This may be the reason why although displaying strong antitumor activity in vitro NK cells could hardly contain tumor burden in vivo. In addition the pathogen driven evolution of NK cell function may also favor the role of NK cells in the insurgence of immune mediated diseases. This research topic will collect contributions that may clarify the relationships between the evolution of the NK receptors and their role in an efficient recognition of viruses and tumor cells or in immune mediated diseases.

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web his paintings were shown along with theirs in major exhibitions of pop art both in this country and in europe and were reproduced often with irony in books catalogs and periodicals throughout the world though primarily a figurative painter mel ramos has experimented freely with realist and abstract art forms for the past twenty years

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