

ImmunoTools IT-Box-Cy55M-Award 2013



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Investigation of the role of CD26 in immune regulation

CD26, also known as dipeptidyl peptidase IV (EC 3.4.14.5), is a ubiquitous, multifunctional protein which involved in several biological processes. A great number of observations have linked CD26 to the functions of the immune system and, in particular to the functions of T cells. As a co-stimulator CD26 play an important role in T cell activation (Fleischer, 1987; Dang et al., 1990a). Only CD4⁺ cells that co-express CD26 can provide helper functions to activate cytotoxic T cells (Dang et al., 1990a) and induce immunoglobulin synthesis by B cells (Gruber et al., 1988).

In order to investigate of the role of CD26 in immune response, CD26 knockout mice are used. In our AG it has been found that CD26^{-/-} mice display an apparently normal phenotype, but in their spleen lymphocyte population, the percentage of CD4⁺ T cells are lower and that of NK cells are higher than in CD26^{+/+} mice. In their peripheral blood, CD26^{-/-} mice presented a conspicuously decreased proportion of CD4⁺ NKT lymphocytes. *In vitro*, compared with CD26^{+/+} mice, the IL-4 production stimulated by PWM in CD26^{-/-} mice was decreased by 60 – 80% in the supernatants of spleen lymphocytes, whereas the production of IL-10 and IFN- γ were increased. After immunization of mice with PWM *in vivo*, serum concentrations of total IgG, IgG1, IgG2a and IgE were markedly lower in CD26^{-/-} mice than those in CD26^{+/+} mice, while no difference was found in IgM production. After ovalbumin immunization *in vivo*, both IgM and IgG production were lower in CD26^{-/-} mouse. But IgE production was not affected. Aerosol challenge with OVA resulted in a much more severe eosinophilia and inflammation in the lung of CD26^{-/-} mice, which was caused by the significant increment of the local Th2 cytokines IL-4, IL-5 and IL-13.

For better understanding of the underlying molecular mechanisms of CD26 function lymphocytes will be isolated from both wild type and knockout mice. After stimulation with different cytokines (rm IFN-g, rm IL-2, rm IL-4, IL-6, rm IL-1 β , rm IL-10 and rm GM-CSF) the cell differentiation and immune response of different cells will be analysed. The recombinant proteins from the *IT-Box-Cy55M* are useful as a standard for the analysis of different cytokine secretion during lymphocytes activation, proliferation and differentiation. In addition, a series of cytokines are substrates of DPP-IV, such as SDF-1a, SDF-1b, RANTES, Eotaxin, MCP-1, MCP-2, MCP-3 and IP-10. By treatment of such recombinant cytokines the different responses of CD26^{+/+}- and CD26^{-/-}-lymphocytes for migration and invasion will be studied.

ImmunoTools *IT-Box-Cy55M* for Xiangli Zhao includes 55 recombinant cytokines

rm EGF, rm Eotaxin / CCL11, rm FGF-a / FGF-1, rm FGF-b / FGF-2, rm FGF-8, rm Flt3L / CD135, rm G-CSF, rm GM-CSF, rm GRO-a / CXCL1, rm GRO-b / CXCL2, rm IFN γ , rm IL-1alpha, rm IL-1beta, rm IL-2, rm IL-3, rm IL-4, rm IL-5, rm IL-6, rm IL-7, rm IL-9, rm IL-10, rm IL-11, rm IL-13, rm IL-15, rm IL-16, rm IL-17A, rm IL-17C, rm IL-17F, rm IL-19, rm IL-20, rm IL-21, rm IL-22, rm IL-25 / IL-17E, rm IL-27, rm IL-31, rm IL-33, rm IP-10 / CXCL10, rm LIF, rm MCP1 / CCL2, rm M-CSF, rm MIP-1 α / CCL3, rm MIP-1 β / CCL4, rm MIP3 α / CCL20, rm MIP3 β / CCL19, rm NGF-beta, rm PDGF-AA, rm PDGF-BB, rm RANTES / CCL5, rm sCD40L / CD154, rm SCF, rm SDF-1 α / CXCL12a, rm SDF-1 β / CXCL12b, rm TNF α , rm TPO, rm VEGF

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