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170-174.抑制吲哚胺2,3-双加氧酶活性促进慢性粒细胞白血病源树突状细胞的功能[J].许思娟,张连生,吴重阳,柴 晔,宋飞雪,岳玲玲,刘 瑛.中华血液学杂志,2009,16(2)

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摘要:

目的: 研究吲哚胺2,3-双加氧酶(indoleamine 2,3-dioxygenase,IDO)在慢性粒细胞性白血病源性树突状细胞(myeloid leukemia, CML DCs)中的表达,及抑制IDO活性对CML DCs免疫刺激功能的影响。方法: RT-PCR检测17例患者外周血单核细胞中IDO mRNA表达。在有无IDO抑制剂1-甲基色氨酸(1-methyltryptophan,1-MT)作用下,分别以不成熟(mDCs)为刺激细胞,完全缓解期(complete remission,CR)CML患者外周T淋巴细胞为反应细胞建立混合淋巴细胞IL-12水平,MTT法检测CML DCs刺激自体T淋巴细胞的增殖能力。结果:随着CML DCs的诱导分化和成熟,IDO mRNA表达除CD1a外,CD80、CD86、CD83、HLA-DR的表达均明显上调( $P < 0.05$ ),且上述分子的表达不受1-MT的影响。高CML DCs的IL-12分泌水平,增强其对自体T细胞增殖的刺激能力,IDO对DCs的负性调节为白血病生物治疗提供了新的思路。

关键词: [慢性粒细胞性白血病](#); [吲哚胺2,3-双加氧酶](#); [树突状细胞](#); [1-甲基色氨酸](#)

Inhibition of indoleamine 2,3-dioxygenase activity promotes function of dendritic cells derived from leukemia [Download Fulltext](#)

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Abstract:

Objective: To investigate the expression of indoleamine 2,3-dioxygenase (IDO) in dendritic cells derived from chronic myeloid leukemia (CML DCs) and to study the influence of IDO inhibition on the function of CML DCs. Methods: The expression of IDO mRNA in peripheral blood mononuclear cells derived from 17 patients with chronic myeloid leukemia was detected by RT-PCR. The phenotypes of CML DCs were analyzed by flow cytometry. The immature CML DCs (imDCs) and the mature CML DCs (mDCs) were used as stimulating cells, and peripheral T lymphocytes of CR patients were used as reactive cells for a mixed lymphocyte reaction system. IL-12 concentration was detected by MTT assay. Results: It was demonstrated that DCs derived from bone marrow mononuclear cells had a more mature morphology. Expressions of costimulatory molecules on DCs, such as CD80, CD86, CD83 and HLA-DR, were obviously higher after maturation ( $P < 0.05$ ) and were not influenced by 1-methyltryptophan (1-MT, an IDO inhibitor). Activity in mature and immature DCs by 1-MT significantly enhanced their abilities to activate T cells ( $P < 0.05$ ,  $P < 0.01$ ). Conclusion: Inhibition of IDO activity in CML DCs can increase their abilities to activate T cells. Negative regulation of DCs by IDO paves a way for DC-based leukemia immunotherapy.

Keywords: [chronic myeloid leukemia](#); [indoleamine 2,3-dioxygenase](#); [dendritic cell](#); [1-methyltryptophan](#)

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