

PublisherInfo		
PublisherName	:	BioMed Central
PublisherLocation	:	London
PublisherImprintName	:	BioMed Central

IFN maintains anergic cell survival

ArticleInfo		
ArticleID	:	139
ArticleDOI	:	10.1186/ar-2000-66857
ArticleCitationID	:	66857
ArticleSequenceNumber	:	96
ArticleCategory	:	Paper Report
ArticleFirstPage	:	1
ArticleLastPage	:	3
ArticleHistory	:	RegistrationDate : 2000-10-13 OnlineDate : 2000-10-13
ArticleCopyright	:	Current Science Ltd2000
ArticleGrants	:	
ArticleContext	:	130753311

Keywords

Anergy, apoptosis, IFN

Context

Following suboptimal stimulation through the T cell receptor, cells either undergo apoptosis or become anergic. Anergic cells have been shown to survive for extended periods *in vivo*, suggesting that anergic cells must be able to escape apoptosis. Previous studies have demonstrated that interferon (IFN)- γ can prevent T cell apoptosis induced by cytokine deprivation. This study aimed to show that IFN- γ is the major survival factor for anergic T cells.

Significant findings

T cell clones were incubated with specific peptide in the absence of antigen-presenting cells. This T:T presentation led to cell death mediated by CD95. When cells were co-cultured with fibroblasts or fibroblast-conditioned media, cell death was significantly reduced. Addition of anti-IFN- γ antibodies to the media significantly reduced the anti-apoptotic activity. IFN- γ prevented apoptosis by inducing STAT-1 activity and blocking protein kinase C- δ translocation to the nucleus. Importantly this study demonstrated that the cells rescued from cell death were all anergic. The authors conclude that IFN- γ rescues T cells from CD95-mediated apoptosis; this mechanism of rescue may account for the persistence of anergic cells *in vivo*.

Comments

There is currently great interest in discovering what influences the cell fate decisions made by T cells following stimulation through their T cell receptor. This study suggests that the micro-environment may determine whether cells become anergic or apoptotic. It focuses on IFN as the main factor for T cell survival; however, it is important to note that other soluble factors may be implicated by the present data. The model used, namely T:T presentation, could occur *in vivo* and the authors postulate that it may

be a mechanism of producing anergic immunoregulatory cells. Previously, the authors demonstrated that IFN- γ maintains T cell survival in the rheumatoid arthritis synovium. They suggest that local secretion of IFN may also promote survival of anergic cells. Such anergic cells may contribute to the maintenance of tolerance by active regulation.

Methods

Co-culture, anergy induction, STAT analysis, PKC staining

References

1. Lombardi G, Dunne PJ, Scheel-Toellner D, Scheel-Toellner D, Pilling D, Taams LS, Life P, Lord JM, Salmon M, Akbar AN: Type 1 IFN maintains the survival of anergic CD4⁺T Cells. *J Immunol.* 2000, 165: 3782-3789.