ABSTRACT

Large Vessel Vasculitis (LVV) is a family of autoimmune diseases characterized by inflammation, smooth muscle cell proliferation, and aneurysm formation in the large vessels, leading to ischemia and/or increased risk of vessel rupture. We previously reported the presence of autoantibodies to 14-3-3ζ in the sera of LVV patients, as well as its increased expression in regions of inflamed vessel tissue in the majority of LVV patients. $14-3-3\zeta$ is one of seven mammalian isoforms of 14-3-3 family of proteins. Like other autoimmune disease, LVV patients have increased levels of IL-17A, therefore, we questioned the role of 14-3-3ζ in the IL-17A signaling and the pathogenesis of LVV. Earlier results show that $14-3-3\zeta$ is required for IL-17A-induced IL-6 production, and that it binds to TNF Receptor Associated Factor (TRAF) 2, 6, and especially 5. Current results indicate 14-3-3ζ role in IL-17A signaling via its influence on TRAF5, its binding partner. 14-3-3ζ was found to promote TRAF5 degradation. We hypothesized that 14-3-3ζ regulates TRAF5 levels in basal and cytokinestimulated environments to influence the outcomes. Our results indicate that the expression of 14-3-3ζ negatively correlates with TRAF5 levels in different cell types. Furthermore short-term stimulation of cells with IL-17A also resulted in degradation of TRAF5, which was found to be reversed after prolonged treatment. Our study provides mechanistic details of 14-3-3ζ role in regulation of IL-17A signaling.



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THESIS
PRESENTATION
by
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The Role of 14-3-3 ζ in IL-17A Signaling

M.S. in Biomedical Sciences

PRESENTATIONS

Peter, C., Chakravarti, R "The Role of 14-3-3ζ in IL-17A Signaling" Poster Presentation at the 46th Annual Physiology and Pharmacology Research Colloquium, Toledo, OH, June 2019

Peter, C., Chakravarti, R "The Role of 14-3-3ζ in IL-17A Signaling" Poster Presentation at the Graduate Research Forum, Toledo, OH, March 2019

Peter, C., Chakravarti, R "The Role of 14-3-3ζ in IL-17A Signaling" Oral Presentation at the Graduate Research Forum, Toledo, OH, March 2018

PUBLICATIONS

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Subramanian, G., Kuzmanovic, T., Zhang, Y., **Peter, C. B.,** Veleeparambil, M., Chakravarti, R., ... Chattopadhyay, S. (2018). A new mechanism of interferon's antiviral action: Induction of autophagy, essential for paramyxovirus replication, is inhibited by the interferon stimulated gene, TDRD7. PLoS pathogens, 14(1), e1006877.

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FUTURE PLANS

Cara plans to return to her home of Seattle, WA and pursue a position in immunology research

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