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# Molecular Mechanism underlying Tissue FOXP3+ Treg Plasticity and Function

How inflammation is initiated in local tissue microenvironment is largely unknown, especially when the balance of tissue homeostasis is challenged under pathogenic conditions. We proposed that tissue FOXP3+Treg cells may play a critical role in the initiation and regulation of tissue inflammation by secreting key inflammatory cytokines such as IFN- $\gamma$ , IL-17 or IL1- $\beta$ . During the talk, I will present our recent findings on the molecular mechanism underlying tissue FOXP3+Treg plasticity and function, together with new data from FOXP3-cre specific conditional mice models to support our hypothesis and engage further discussions.

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### 銀屑病的免疫學機制和治療新靶點

銀屑病,俗稱"牛皮癬",是慢性炎症性皮膚疾病,被世界衛生組織(WHO)列為人類目前不可治癒的十大頑症之一。銀屑病在我國發病率為 0.5%,患者約 1000 萬。迄今為止,銀屑病發病原因不清且無法根治。因此,明晰銀屑病發病機理和發現功能性治療新靶點是銀屑病領域兩個重要的科學問題。申請人長期從事銀屑病研究,在銀屑病病因學和治療新靶點領域取得了系列原創性研究成果。1、申請人團隊發現 RNA 病毒感染而非鏈球菌感染誘發銀屑病的發生發展,這是銀屑病病因學研究的重要突破,為銀屑病的積極預防和治療提供了全新思路;2、申請人團隊研發出選擇性抑制 Th17 細胞分化的天然候選藥物小分子,該小分子擁有全球自主智慧財產權,獲得國家發明專利 2 項,PCT專利 1 項。該小分子擬于近期申報中國 IND,將來有望成為我國自主研發、自主智慧財產權、First-in-Class 治療銀屑病的國家 I 類新藥。申請人團隊相關工作發表在 Nature Medicine、Nature Communications、Immunity 等 SCI 刊物上。團隊自 2009年組建以來,獲得國家自然科學基金傑出青年基金、國家重點基礎研究發展計畫(973)、國家自然科學基金重點專案、國家自然科學基金重大研究計畫、國家自然科學基金面上專案等多項資助,總經費達 1700 萬。依託上海交通大學醫學院上海市免疫學研究所建立了 15 人的研究團隊和皮膚免疫學實驗室(Lab of Skin Immunology),團隊培養博士生、博士後 9 人。