

SIgN SEMINAR

hosted by Dr Ng Lai Guan



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Macrophage Immunometabolic Repair of Heart

During wound injury, efferocytosis fills the macrophage with a metabolite load nearly equal to the phagocyte itself. A timely question pertains to how metabolic phagocytic signaling regulates the signature anti-inflammatory macrophage response. Here we report the metabolome of activated macrophages during efferocytosis to reveal an interleukin-10 (IL-10) cytokine escalation that was independent of glycolysis yet bolstered by apoptotic cell fatty acids and mitochondrial β -oxidation, the electron transport chain, and heightened coenzyme NAD^+ . Loss of IL-10 due to mitochondrial complex III defects was remarkably rescued by adding NAD^+ precursors. This activated a SIRTUIN1 signaling cascade, largely independent of ATP, that culminated in activation of IL-10 transcription factor PBX1. IL-10 activation by the respiratory chain was also important in vivo, as efferocyte mitochondrial dysfunction led to cardiac rupture after myocardial injury. These findings highlight a new paradigm whereby macrophages leverage efferocytic metabolites and electron transport for anti-inflammatory reprogramming that culminates in organ repair.



11 September 2019 (Wednesday)

11:00am – 12:00pm

SIgN Seminar Room, Immunos Level 4

*Seminar is
open for all
to attend.*

*Registration
is not
required.*

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