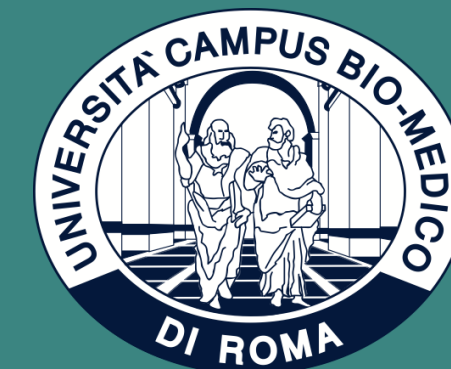


BIOACTIVE LIPIDS: ENDOCANNABINOIDS AND BEYOND

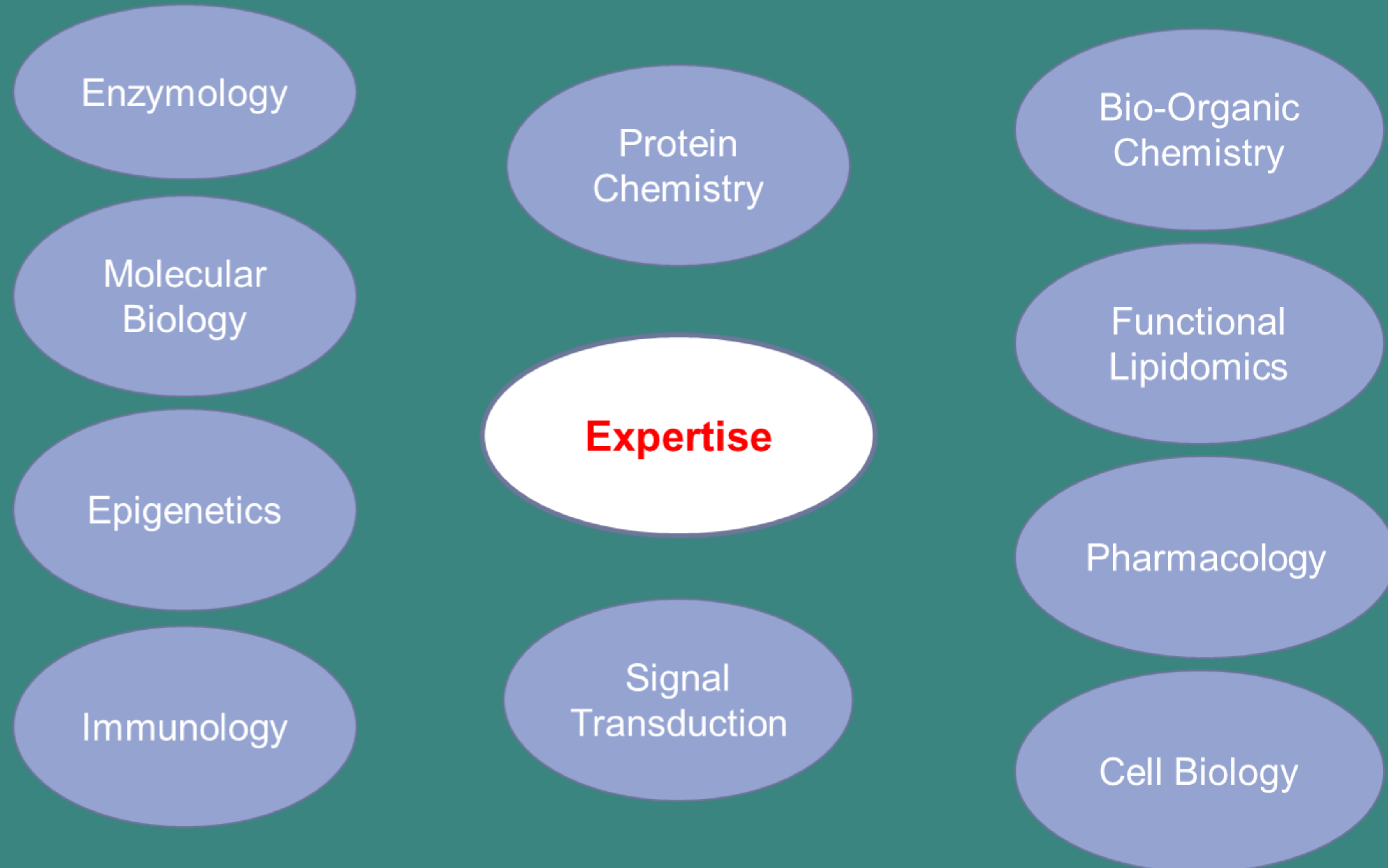
PROF. DR. MAURO MACCARRONE

Unit of Biochemistry and Molecular Biology

The Lipid Connection



The Lipid Connection



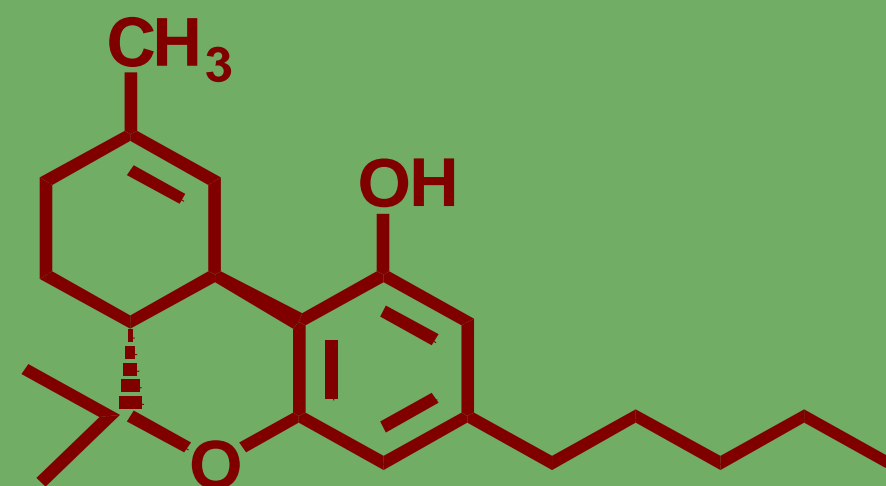
Brief history of (endo)cannabinoids



- 200: The therapeutic properties of cannabis are described in Chinese pharmacopoeia

- 1838-1840: The medicinal properties of cannabis are assessed methodically

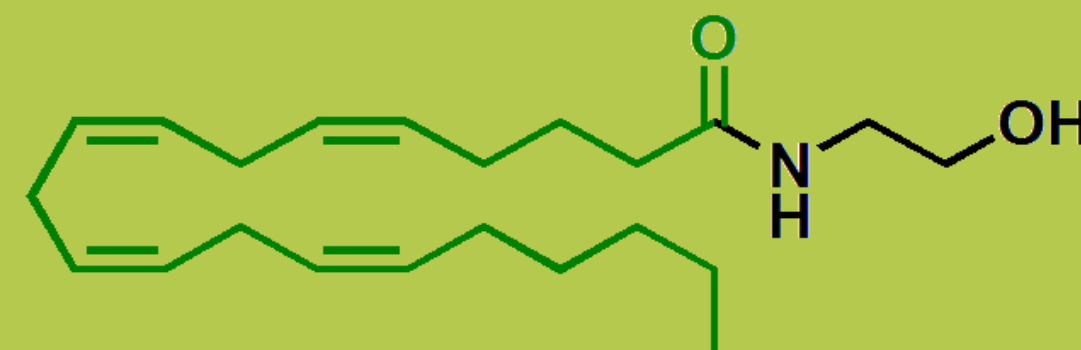
- 1964: Gaoni and Mechoulam elucidate the structure of **Δ^9 -tetrahydrocannabinol (THC)**



- 1990: Matsuda and colleagues clone the CB₁ receptor

- 1992: Mechoulam's group in collaboration with Pertwee's group identifies the first endocannabinoid, *N*-arachidonylethanolamine or **anandamide (AEA)**

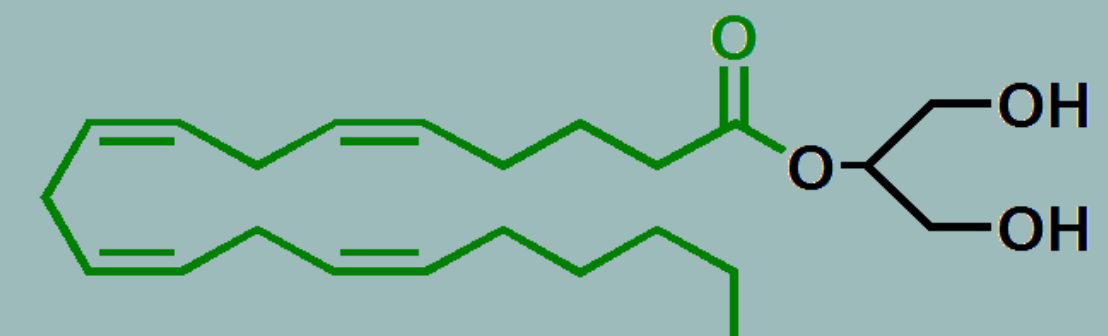
- 1993: Munro and colleagues clone the CB₂ receptor



- 1995: Mechoulam's group and Sugiura's group identify the second endocannabinoid, **2-arachidonoylglycerol (2-AG)**

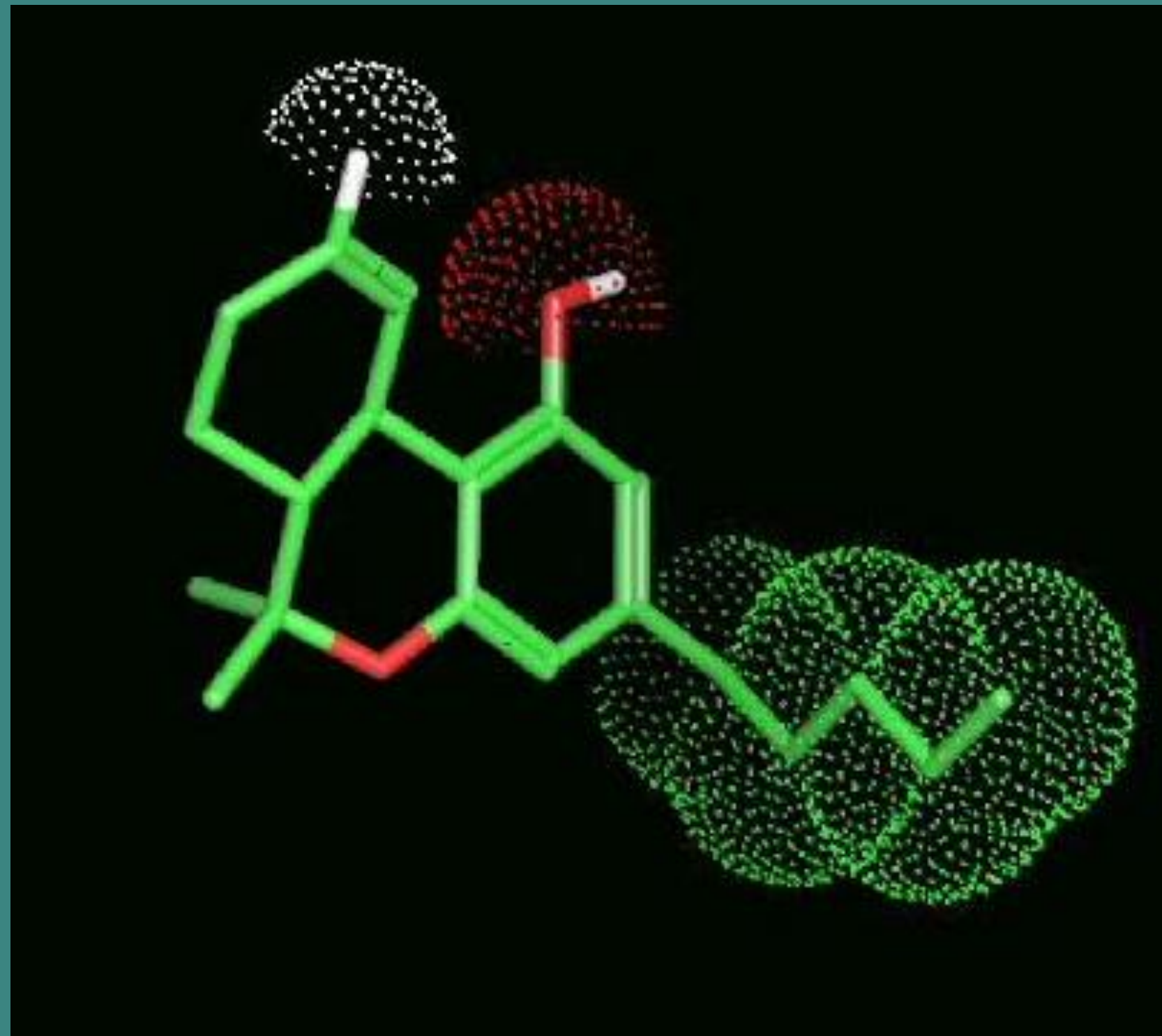
- 1996: Cravatt and colleagues clone the first endocannabinoid-degrading enzyme, FAAH

- 2003: Bisogno and colleagues clone the first 2-AG-bio-synthesizing enzyme, DAGL

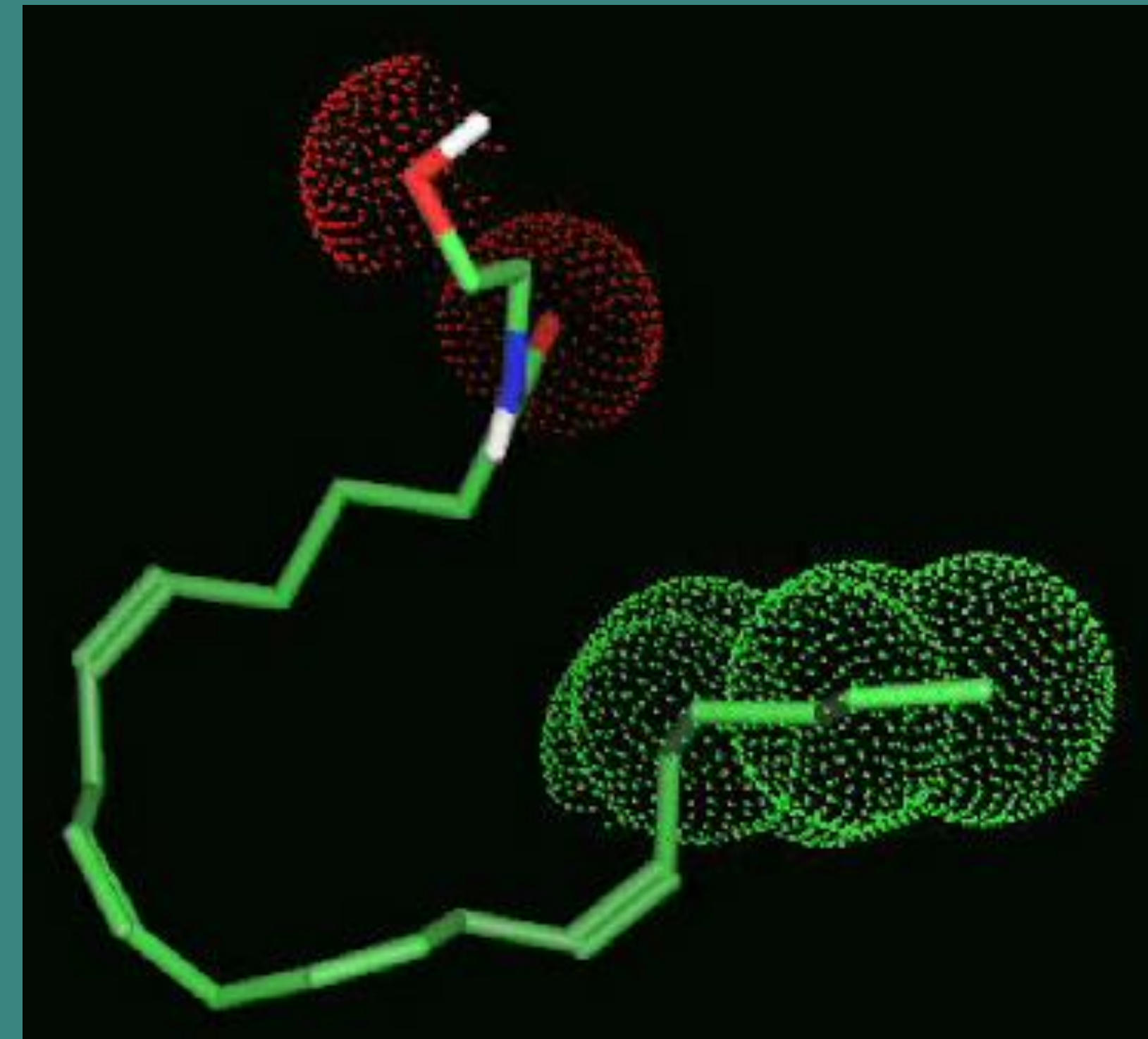


Similarities between:

THC
(a terpeno-phenol compound)

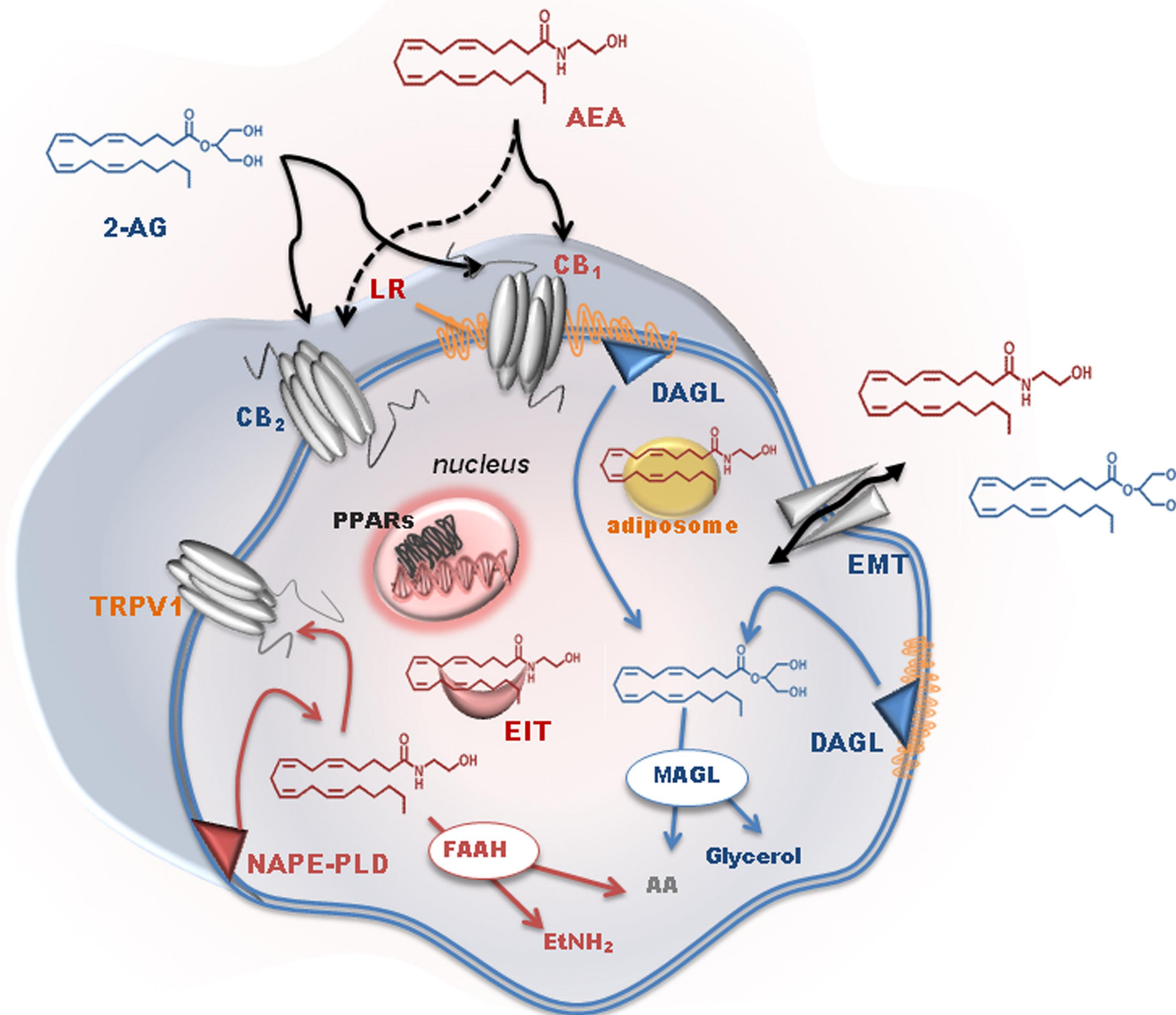


&



Anandamide
(*N*-arachidonylethanolamine)

Endocannabinoid system at a glance



Biological activities of endocannabinoids

Interference with dopaminergic transmission
Inhibition of GABAergic transmission
Inhibition of glutamatergic transmission
Induction of long term depression
Control of pain initiation
Control of wake/sleep cycles
Control of thermogenesis
Control of appetite
Impairment of working memory
Impairment of memory consolidation
Inhibition of long term potentiation
Control of psychomotor disorders
Control of energy homeostasis

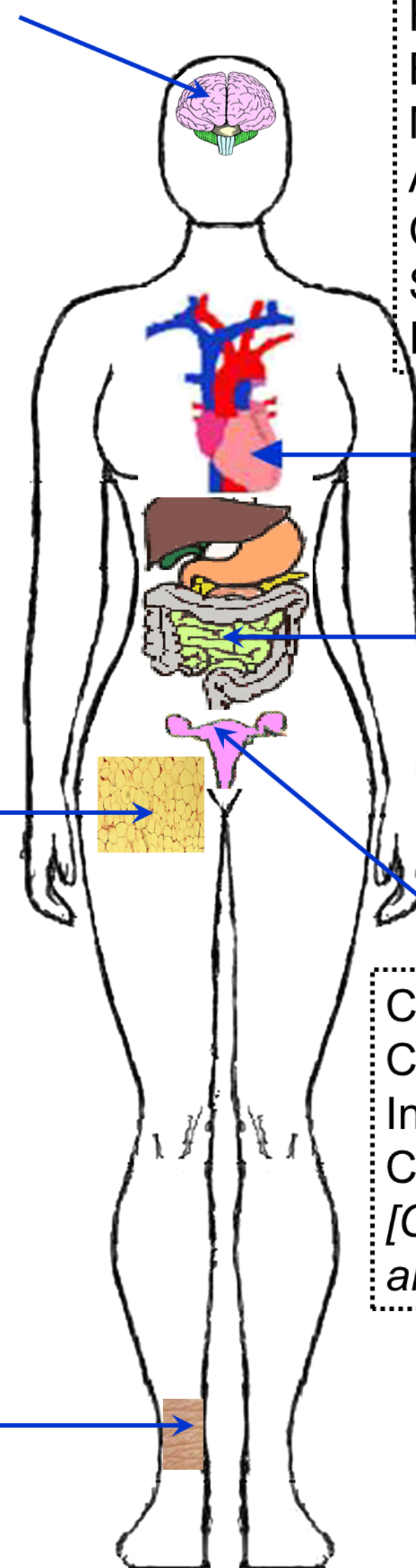
Hypotension
Bradichardia
Modulation of inflammation
Activation of platelet aggregation
Control of cytokine production
Stimulation of haematopoietic stem cell growth
Inhibition of chemotaxis

Inhibition of peristalsis

Regulation of lipid metabolism

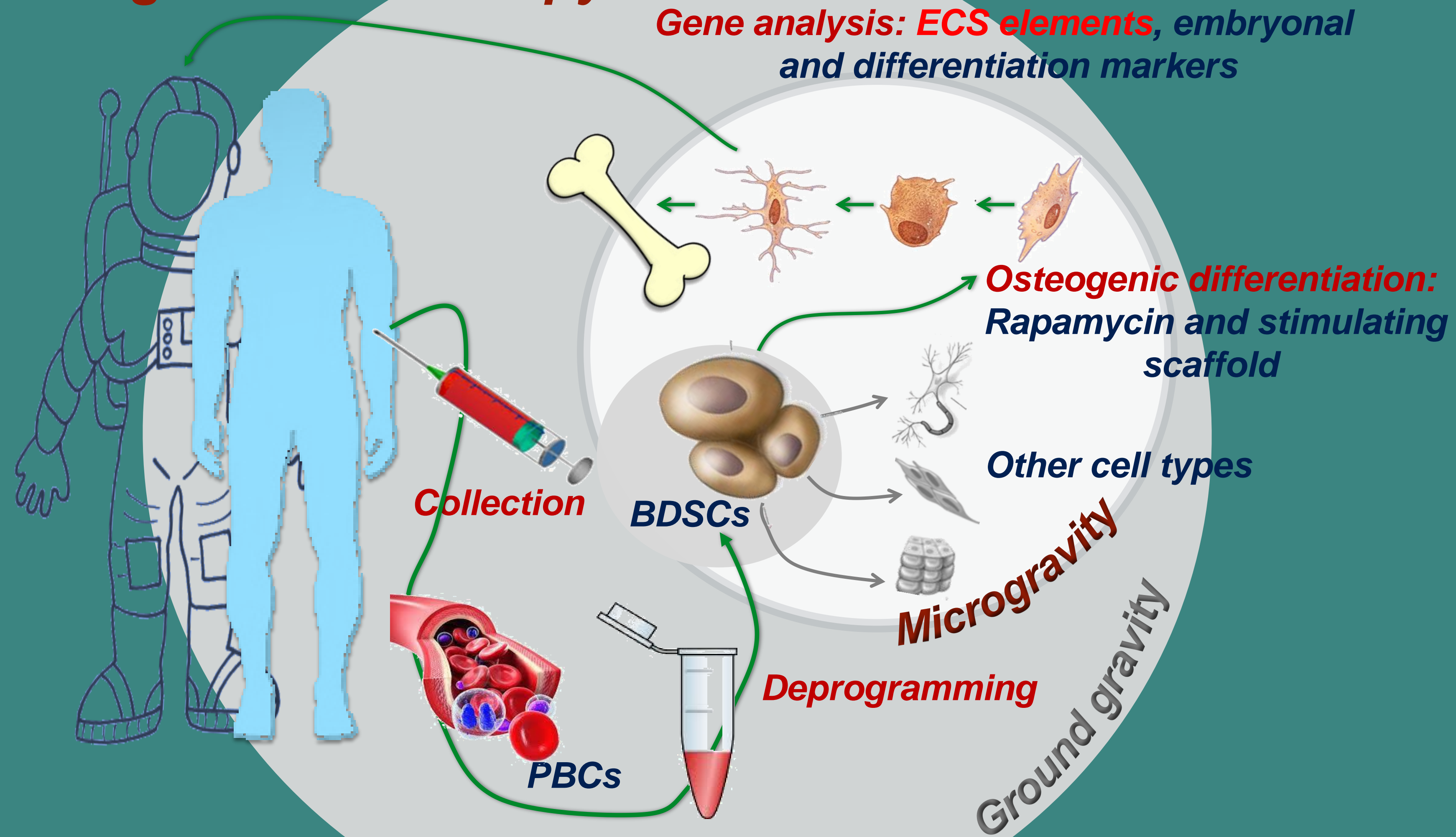
Control of folliculogenesis?
Control of oviductal transport
Inhibition of blastocyst implantation
Control of trophoblast differentiation and growth
*[Control of spermatogenesis, sperm motility
and acrosome reaction in males]*

Arrest of keratinocyte differentiation



SERiSM: Role of the Endocannabinoid System in Reprogramming Human Pluripotent Stem Cells under Microgravity

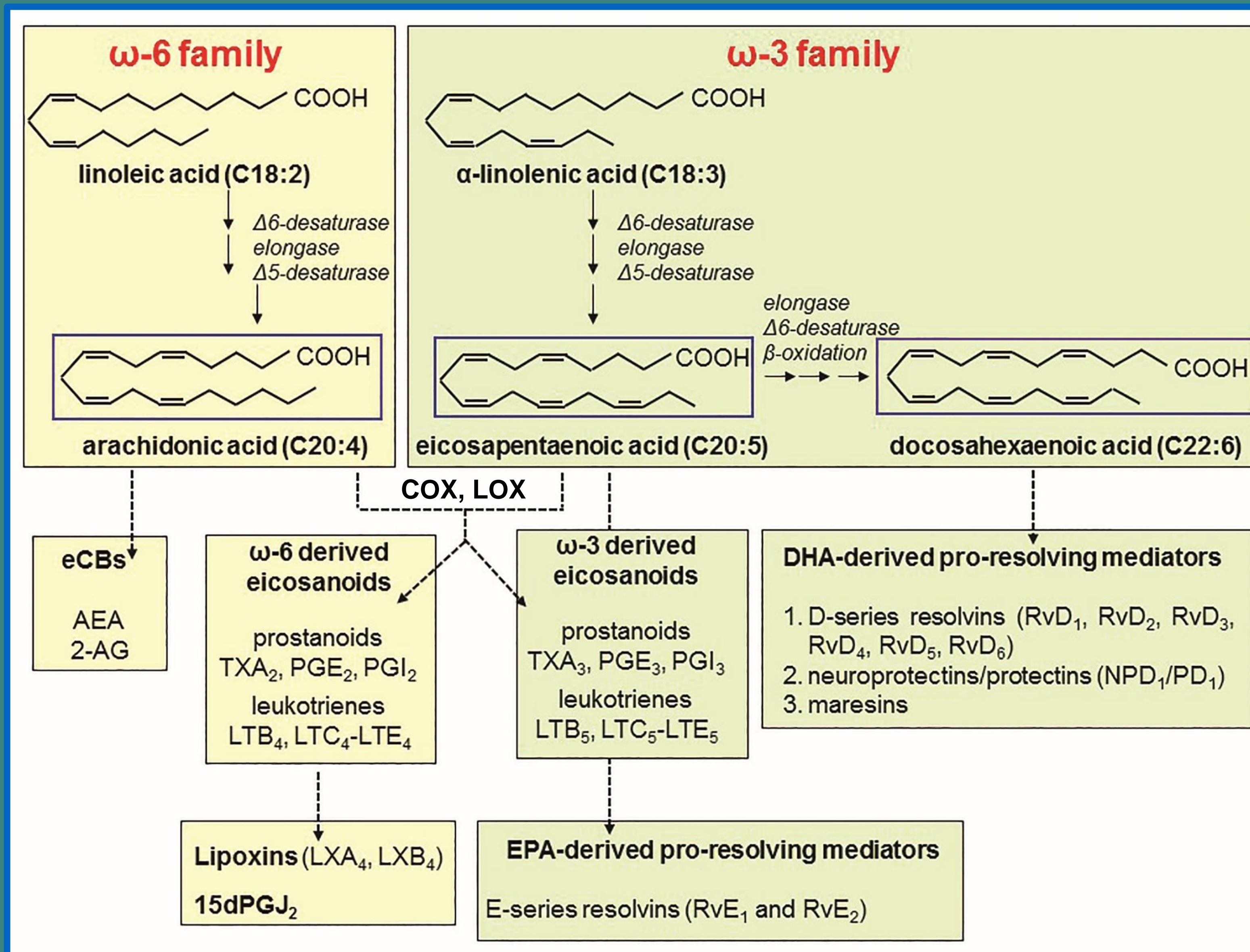
Regenerative therapy?



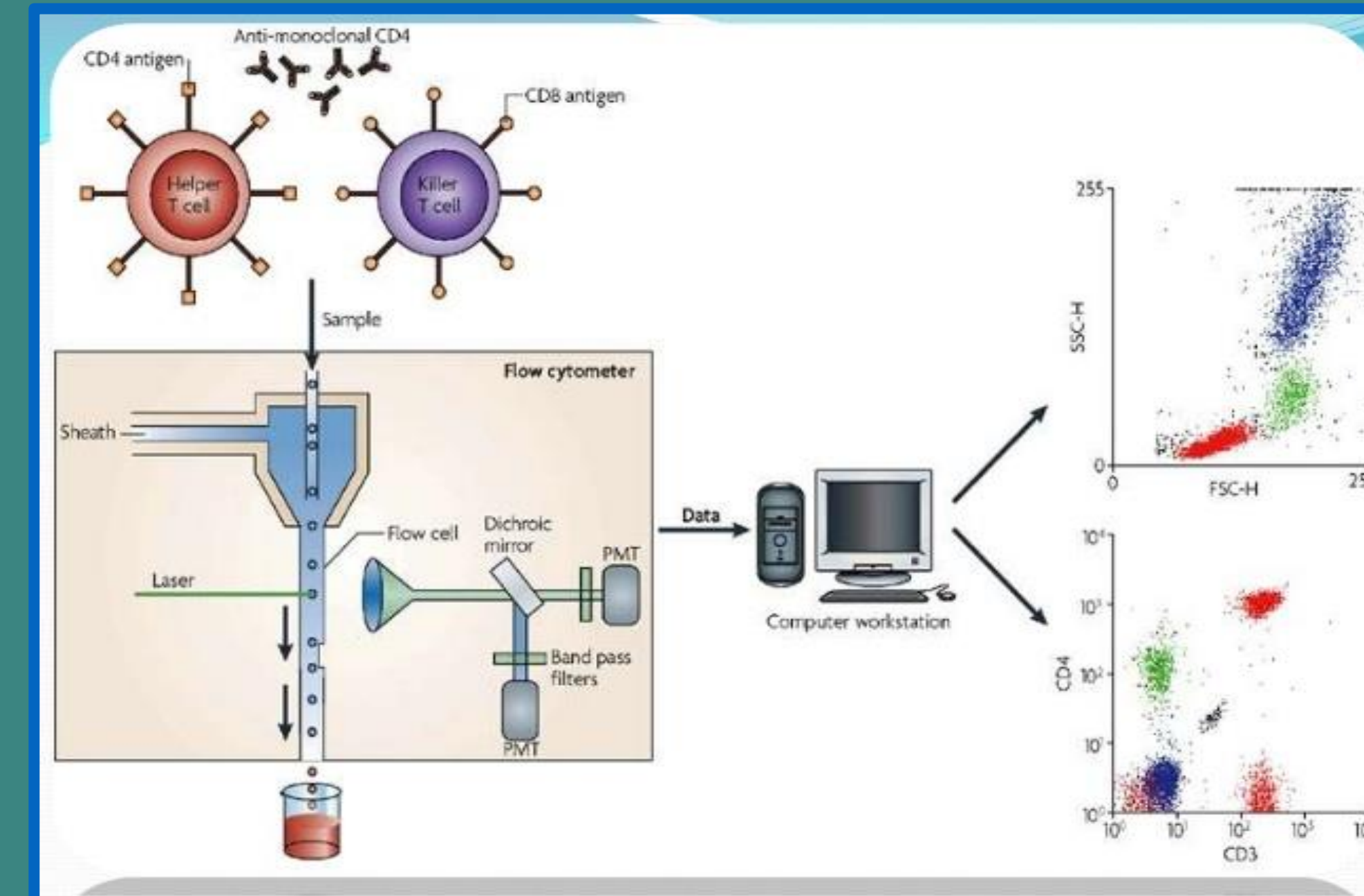
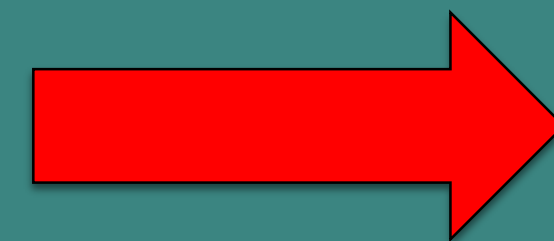
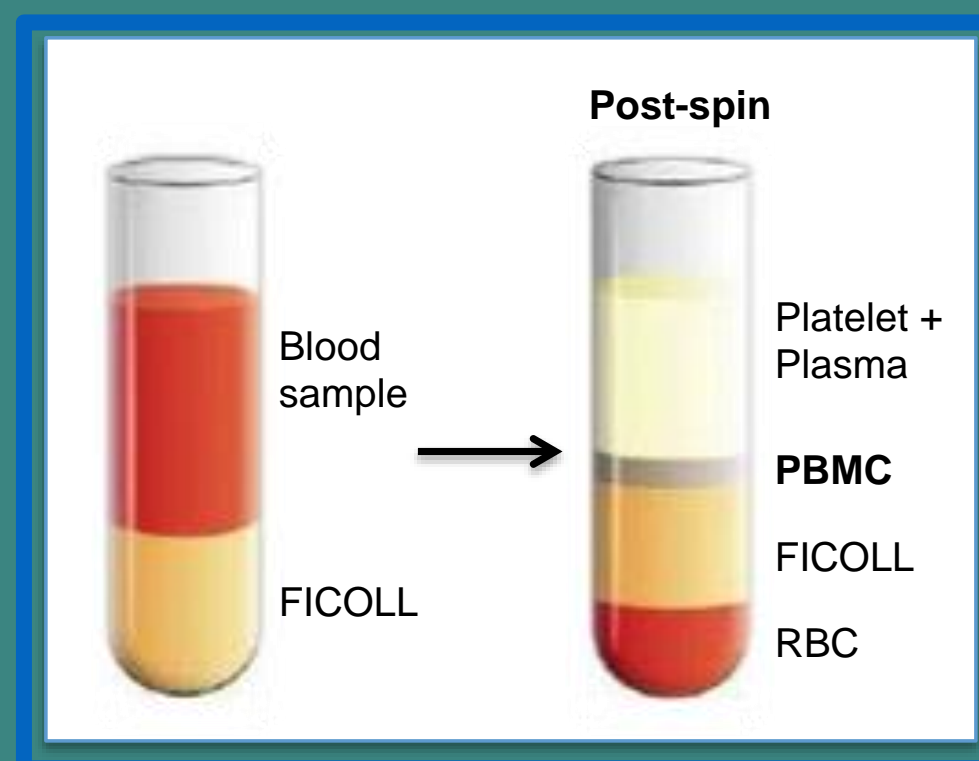
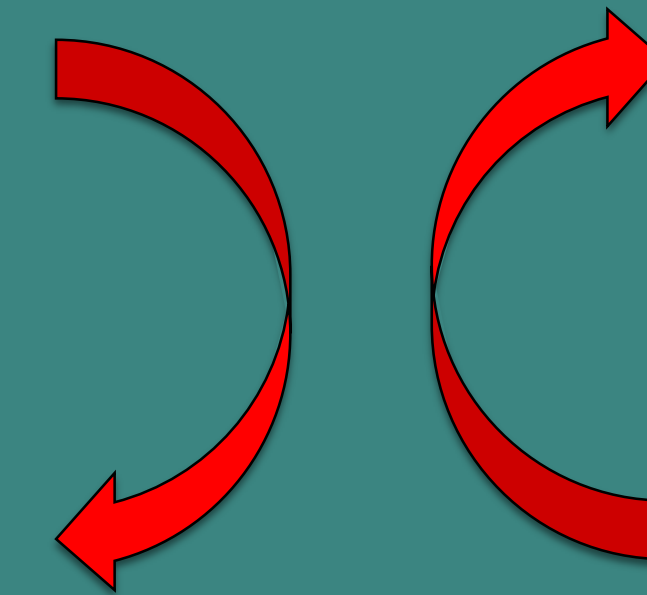
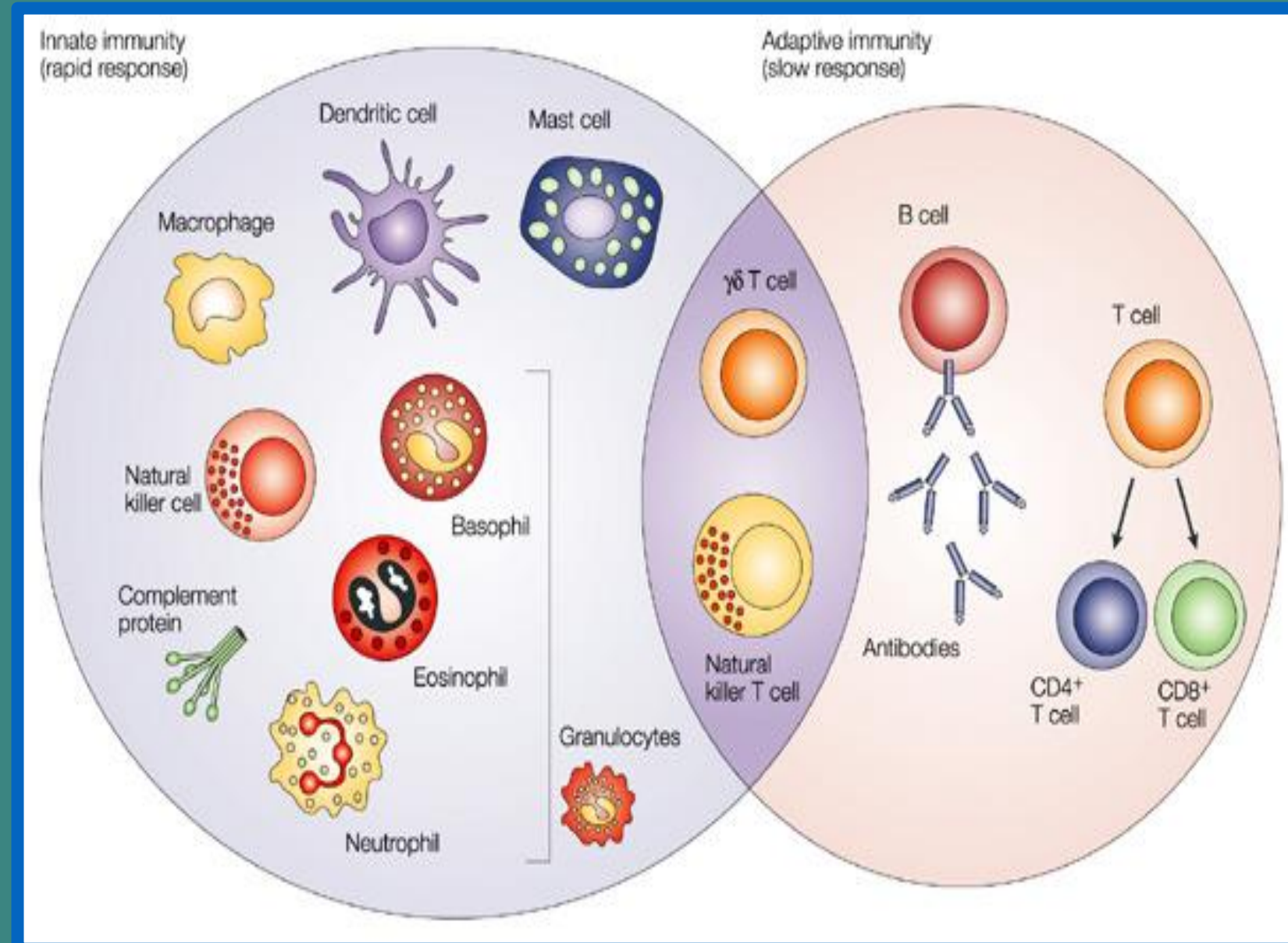
Back to the International Space Station in August 2017! (*Space Mission Expedition 53*)

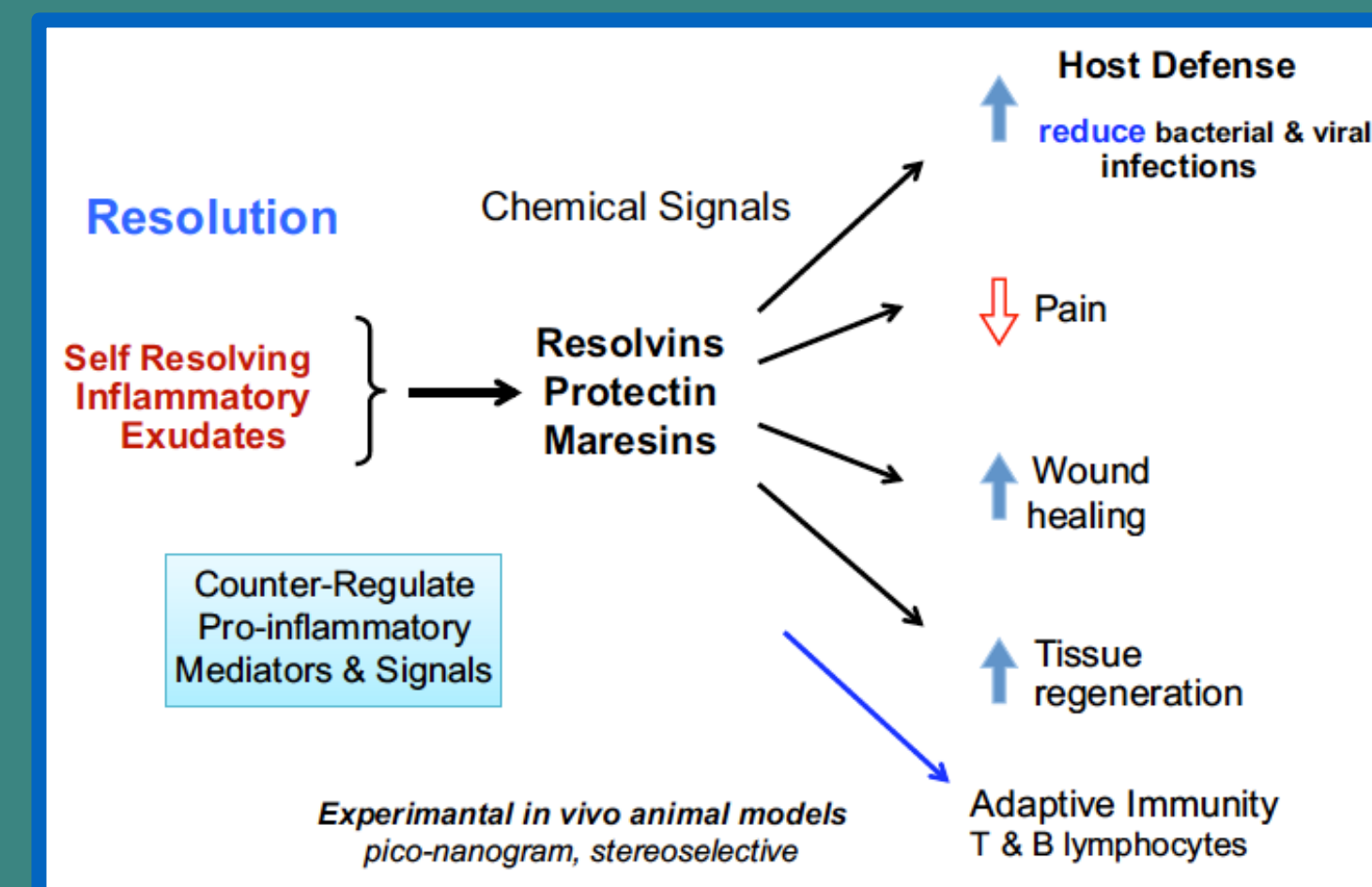
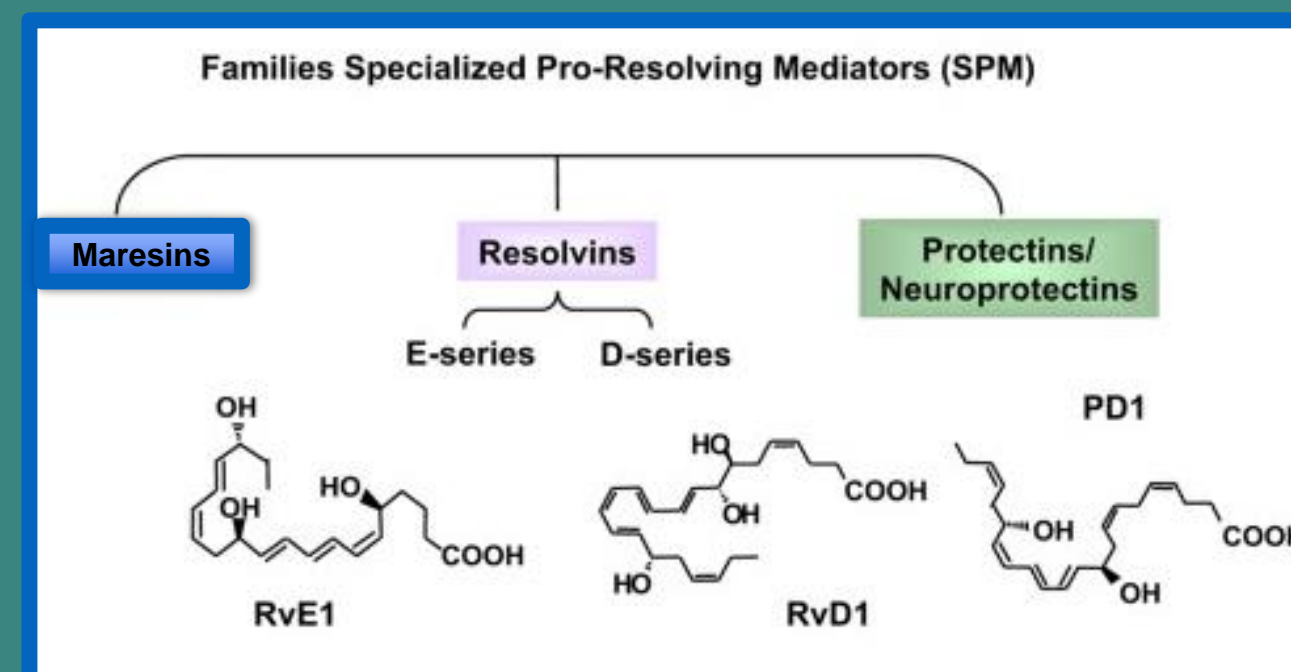
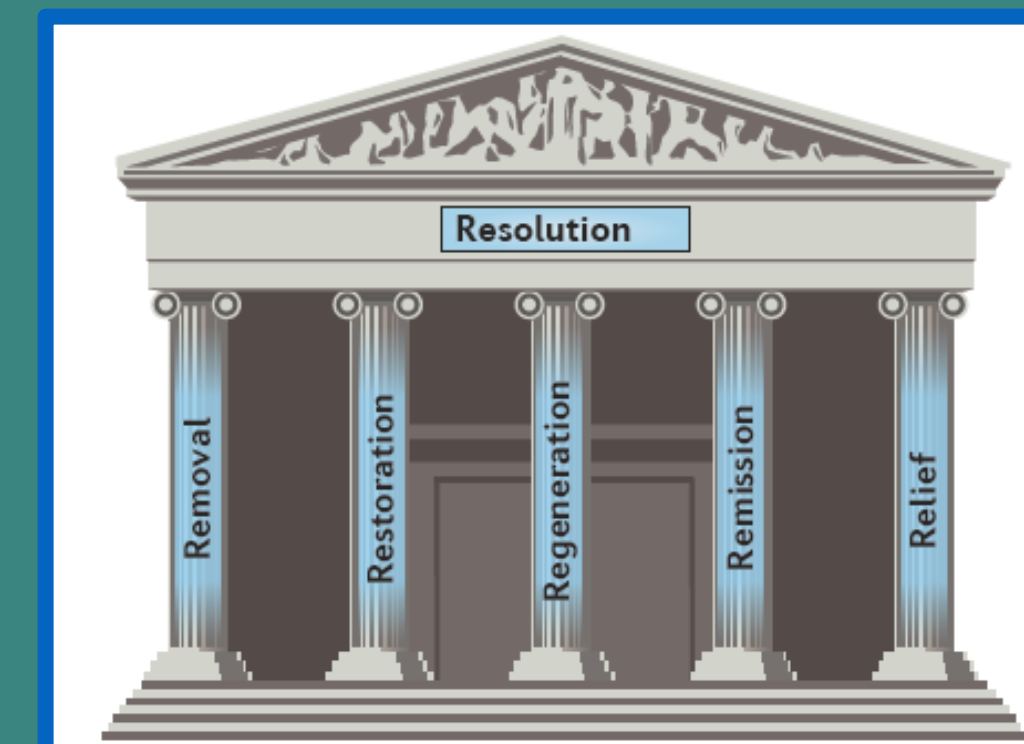
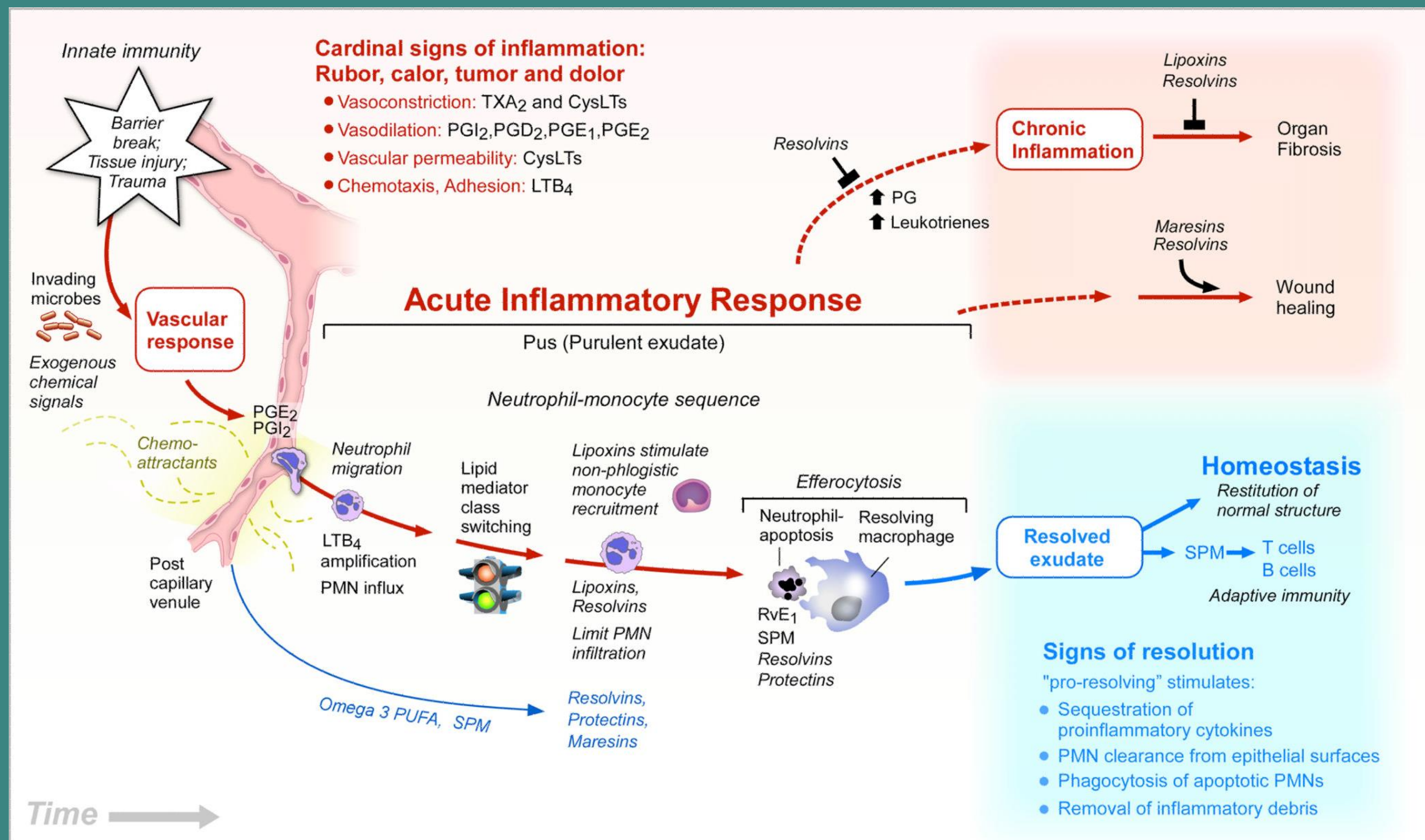
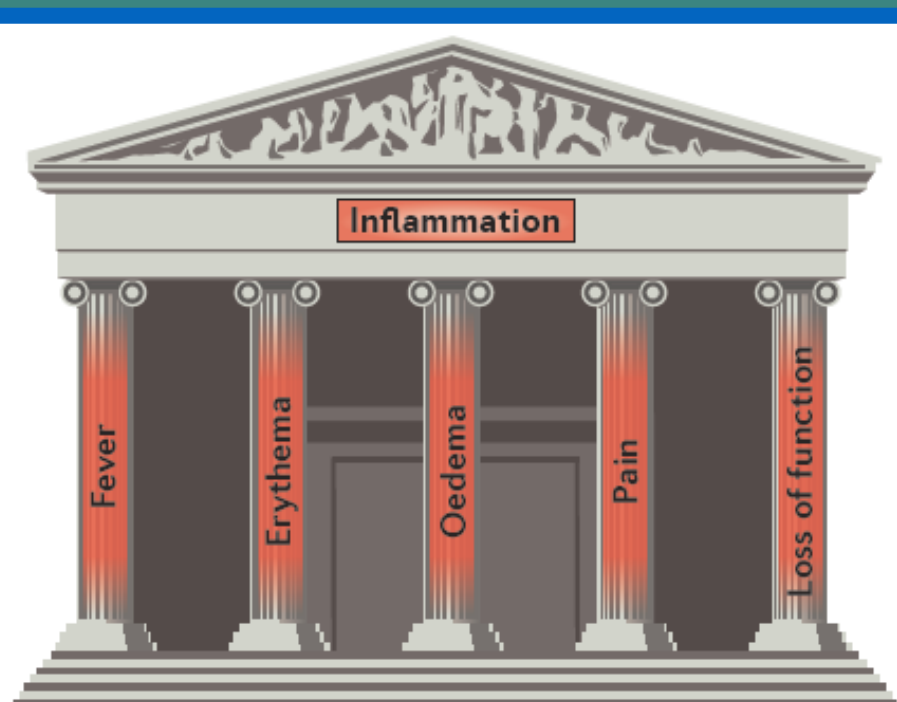


Beyond endocannabinoids: Pro-resolving lipid mediators



Bioactive lipids and immunity: from cell isolation to immunophenotyping and immune function



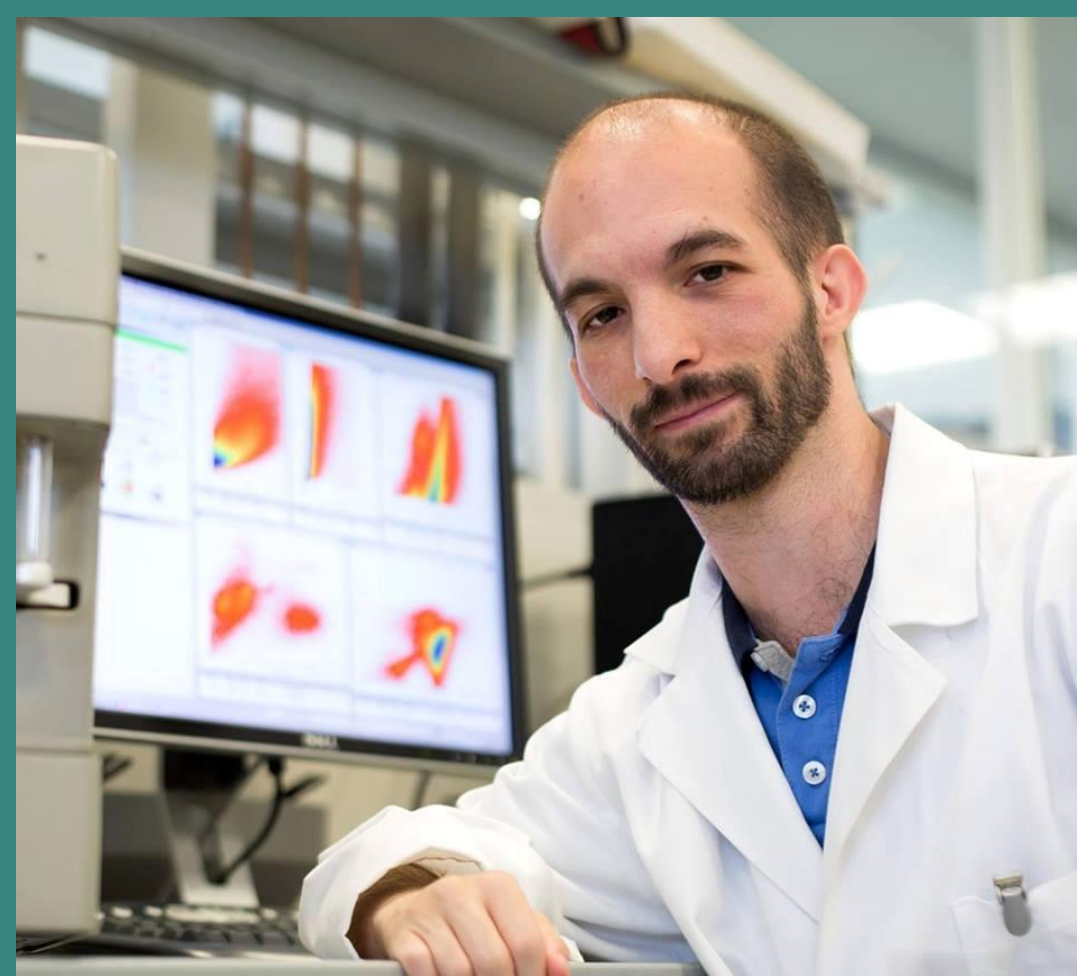


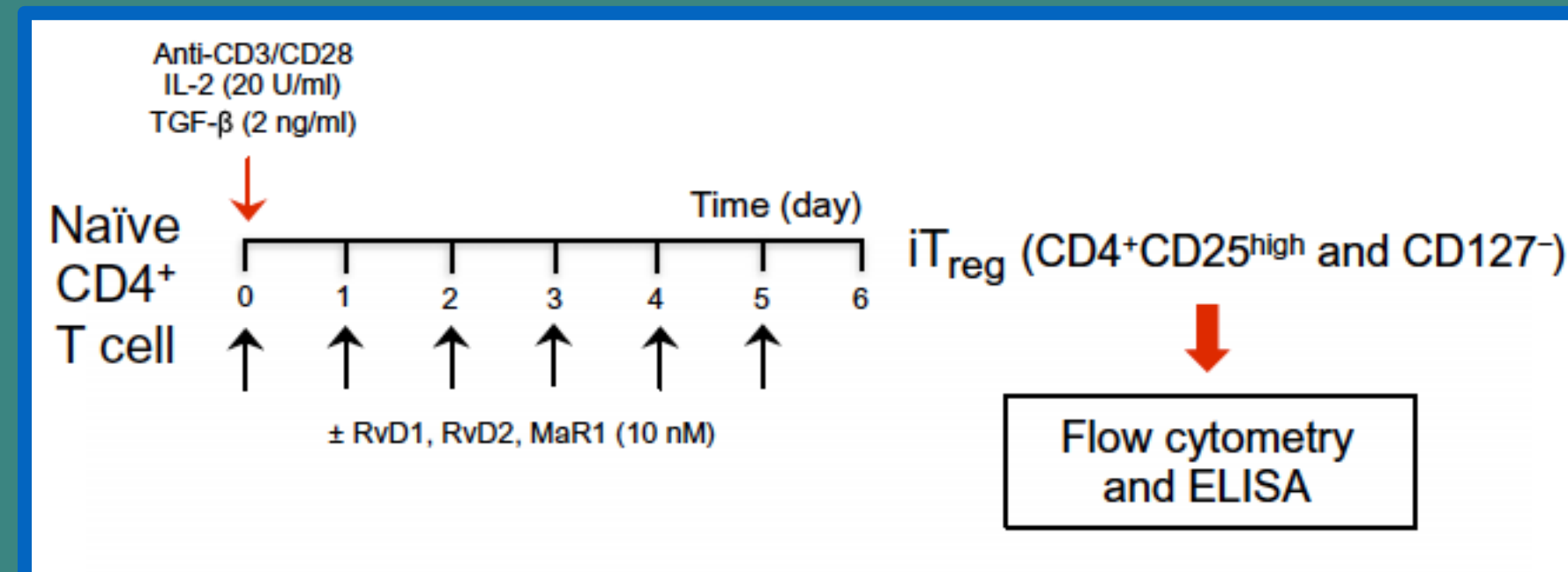
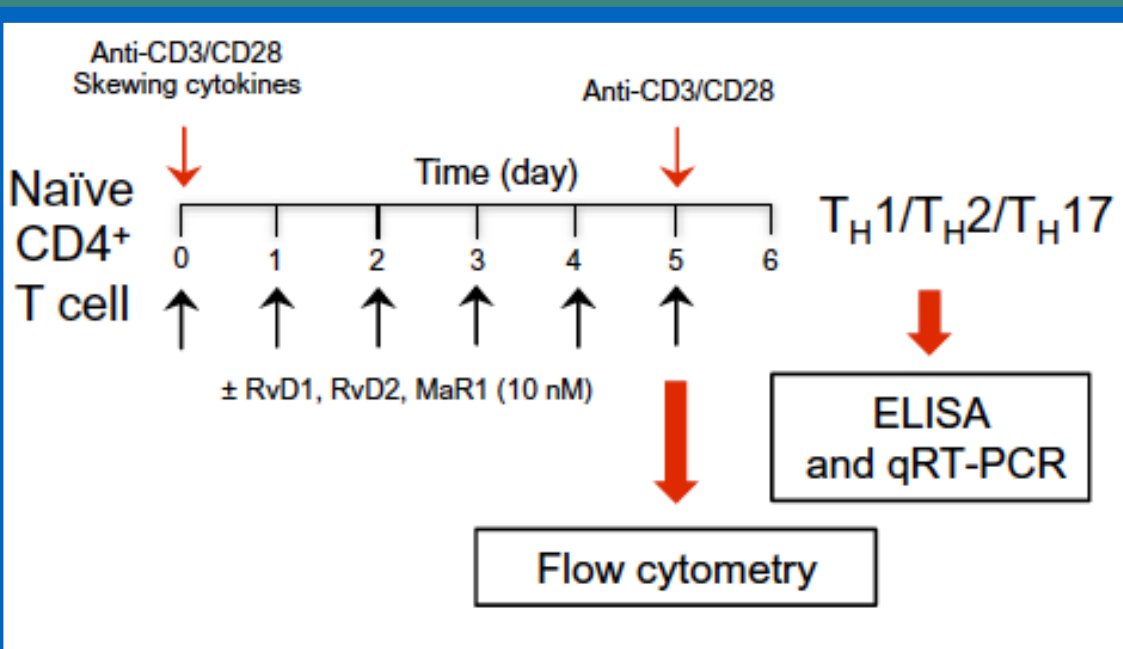
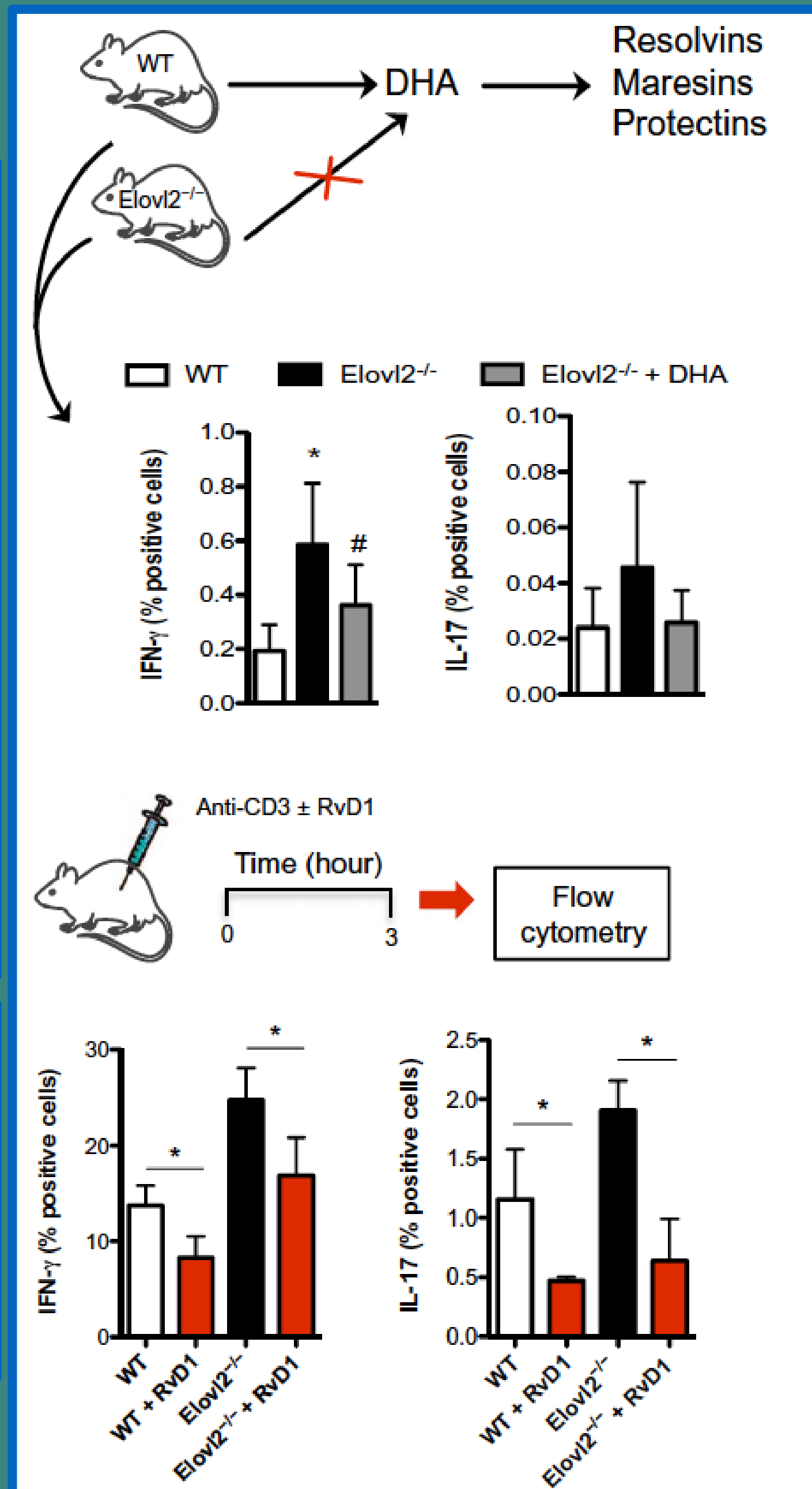
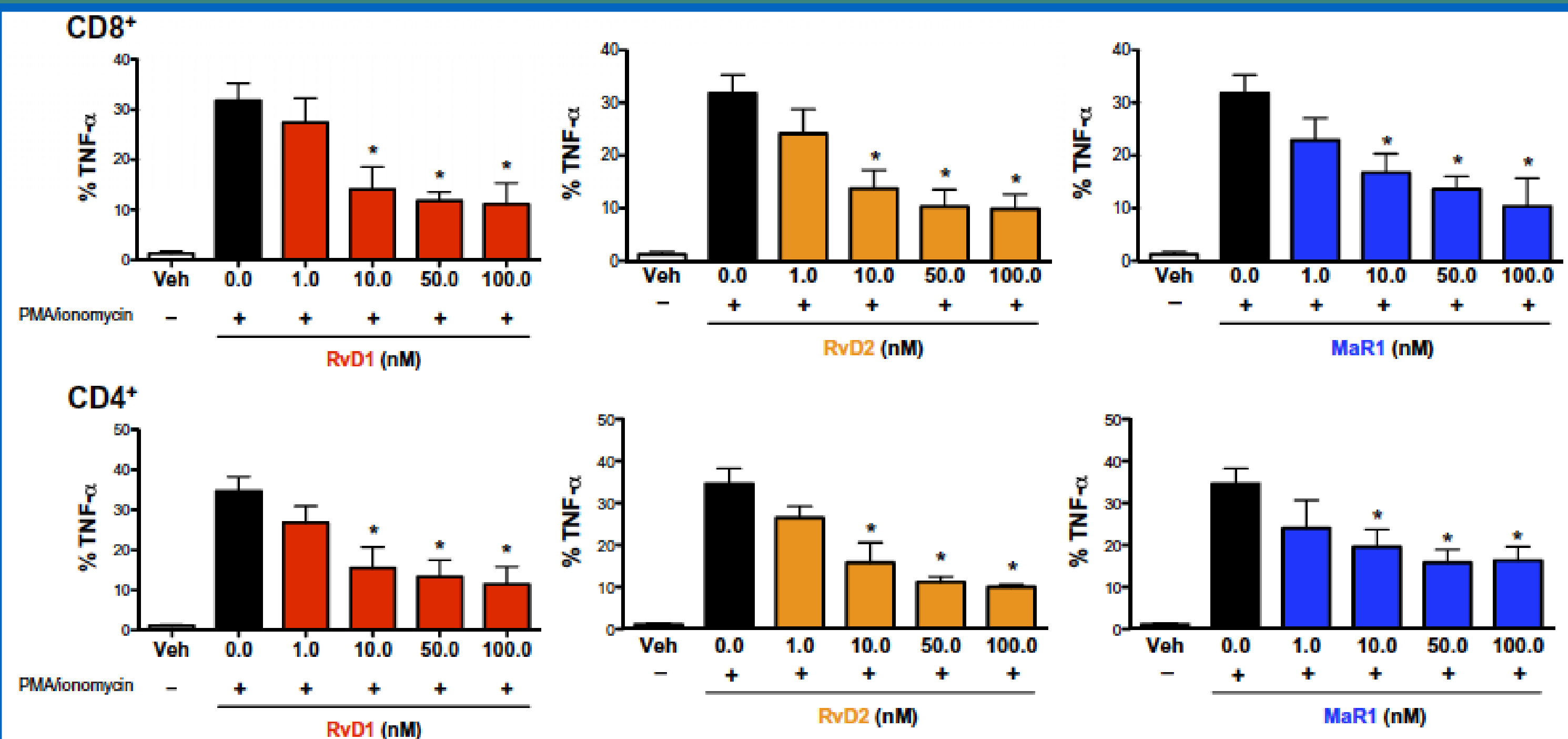
RESEARCH ARTICLE

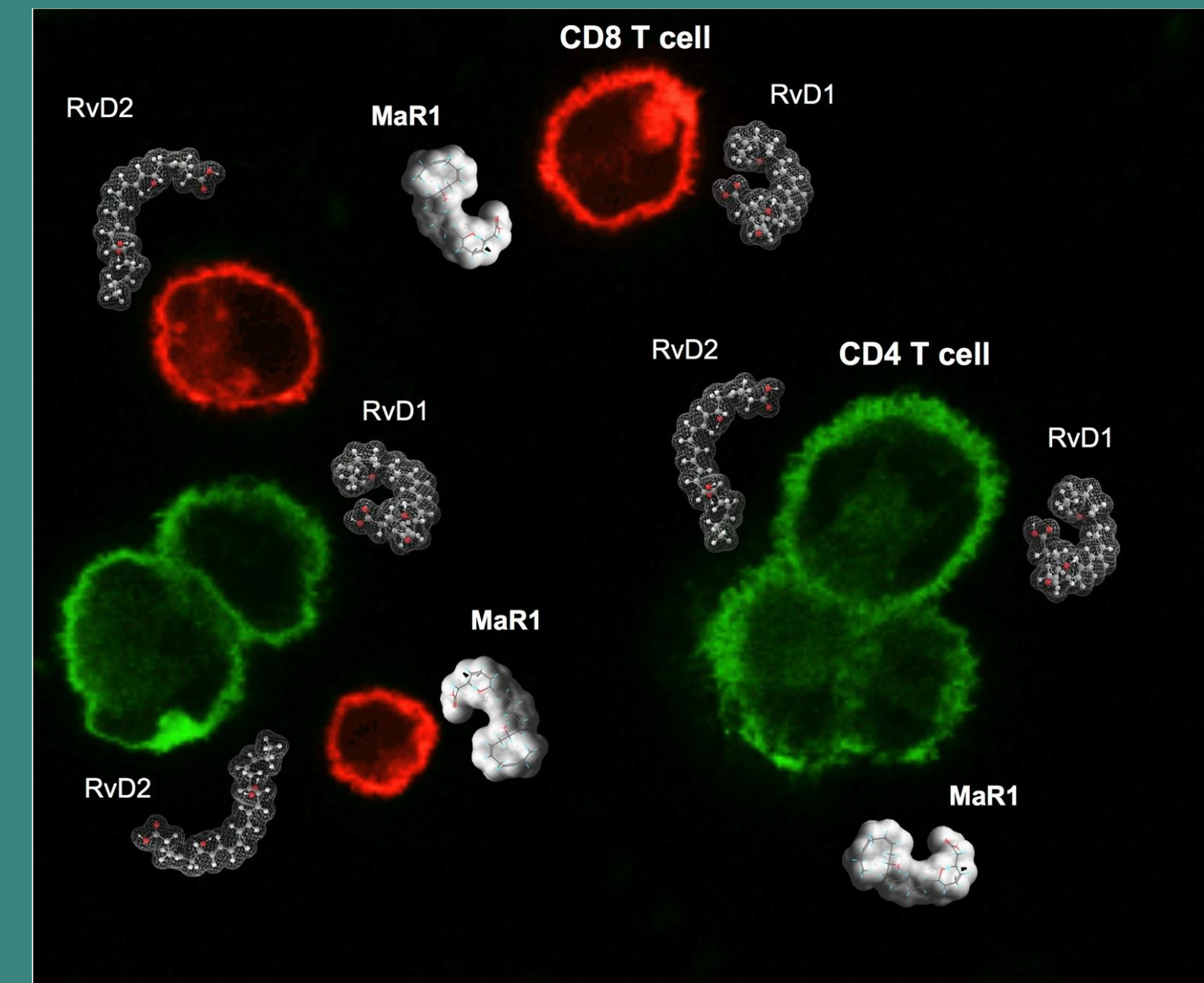
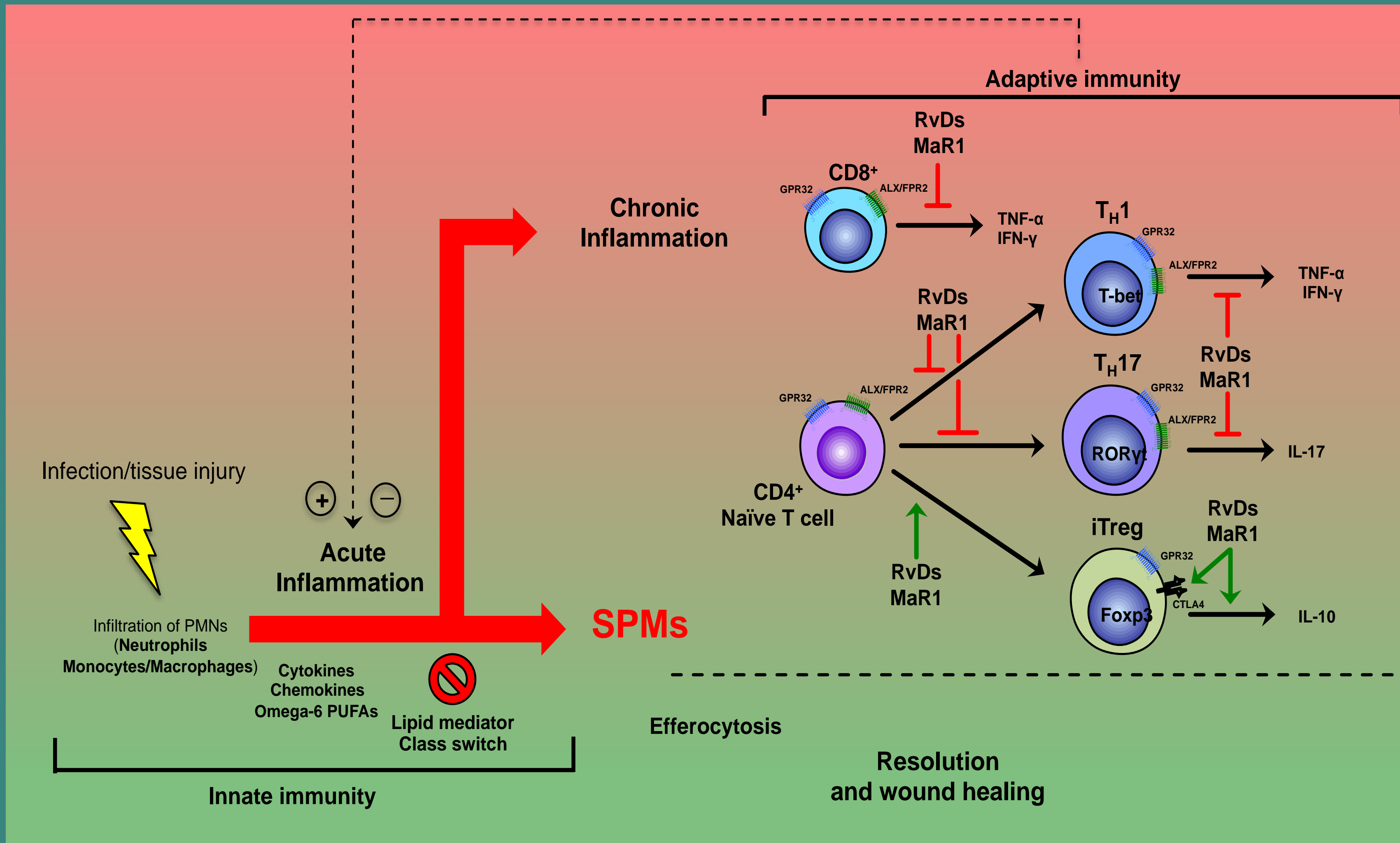
INFLAMMATION

Proresolving lipid mediators resolvin D1, resolvin D2, and maresin 1 are critical in modulating T cell responses

Valerio Chiurchiù,^{1,2*} Alessandro Leuti,^{1,2} Jesmond Dalli,³ Anders Jacobsson,⁴ Luca Battistini,⁵ Mauro Maccarrone,^{1,2†} Charles N. Serhan^{3†*} [†]Equally senior authors







Modified from Chiurchiù et al., Science Transl Med, 2016