

Novel small molecule mast cell activator vaccine adjuvants

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Value Proposition

Mast cells (MCs) are innate immune cells highly prevalent in the dermal region of the skin and in the respiratory, gastrointestinal, and urinary tract mucosa. The strategic location at the host-environment interface and the expression of a wide range of pattern recognition receptors (PRRs) allow MCs to serve as immune surveillance cells, and these cells play a critical host-protective role as a first line of defense through innate immune responses. In addition, the expression of receptors on MCs recognizing activated complement, endogenous peptides (antimicrobial peptides, neuropeptides, etc.), exogenous venom peptides, and interleukin-33 (IL33) can amplify proinflammatory responses and modulate immune responses. The immune regulatory role of MCs is particularly supported by the fact that MCs have a capacity to pre-store proteases, cytokines, biogenic amines, and other mediators that can be instantly released at the recognition of trigger stimuli. In addition to pre-stored mediators, MCs produce and secrete a wide range of cytokines and chemokines through de novo synthesis. While MCs are known to regulate innate and adaptive immunity, and MC activators have recently been described as safe and effective vaccine adjuvants, many currently known MC activators are inadequate for in vivo applications, however, and research on identifying novel MC activators is limited.

Technology

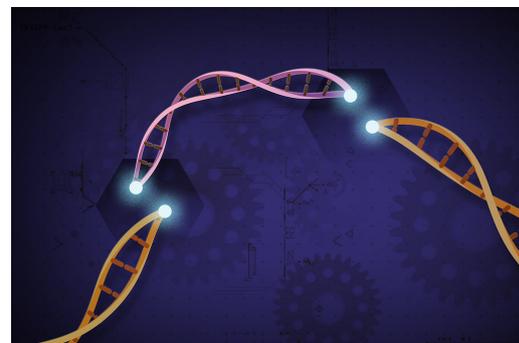
Duke inventors have reported novel small molecules that act as mast cell activators intended to be developed as vaccine adjuvants. After identification of small molecules with mast cell activating activity, the inventors performed medicinal chemistry to develop novel small molecules with mast cell activating activity similar to or superior to the parent compounds. These novel small molecules were tested for vaccine adjuvant activity in mice.

COVID-19 Application

This technology could be used as adjuvants for COVID-19 vaccines.

Advantages

- Small molecules that activate mast cells (MCs) and can be used as vaccine adjuvants
- Synthesis of compounds is easier than related polymers or synthetic peptides and is expected to cost less
- Promote de novo synthesis of cytokines and induce the



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Inventor(s)

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- Choi, Hae Woong
- Gooden, David
- Johnson-Weaver, Brandi
- Kalpathy, Santhosh

Links

- [From the lab of Dr. Herman Staats](#)
- [From the lab of Dr. Soman Abraham](#)

College

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release of eicosanoids from human to mouse MCs

Publications

- [Identification of Novel Mast Cell Activators Using Cell-Based High-Throughput Screening \(SLAS Discovery, 2019\)](#)