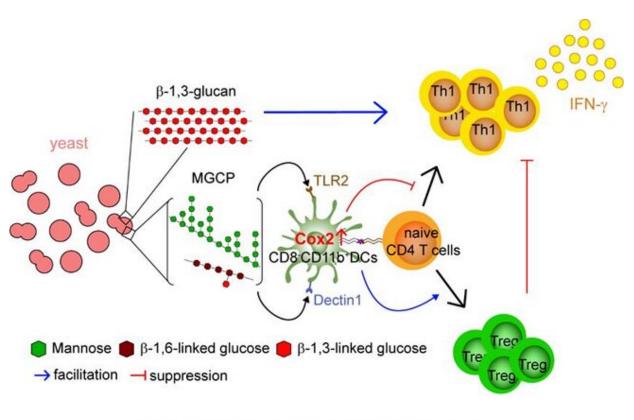


Stopping the onset and progression of intractable immune diseases

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Schematic diagram depicting anti-inflammatory response mechanisms mediated by MGCP

Credit: Pohang University of Science & Technology (POSTECH)

The development of therapeutic drugs for inflammatory bowel disease, an intractable immune disease, and multiple sclerosis—an autoimmune



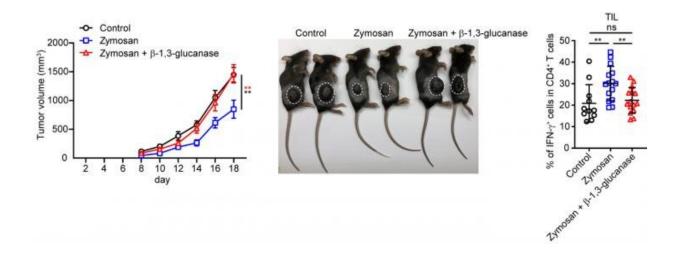
disorder—is gaining traction. A research team from the Department of Life Sciences at POSTECH and a joint research team at ImmunoBiome Inc. have uncovered that a yeast-derived polysaccharide mixture inhibits the onset and progression of immune disorders.

The number of cases of Crohn's <u>disease</u> and ulcerative colitis—both inflammatory bowel diseases—in Korea was about 18,000 and 37,000 respectively as of 2019, increasing about 2.3 times over the past decade. Multiple sclerosis is a rare and incurable disease that affects about 2,500 patients in Korea. Both diseases are intractable inflammatory diseases caused by abnormalities in the human immune system. The exact cause of the <u>inflammatory bowel disease</u> and multiple sclerosis is still unknown but it is presumed that environmental and genetic factors play a role.

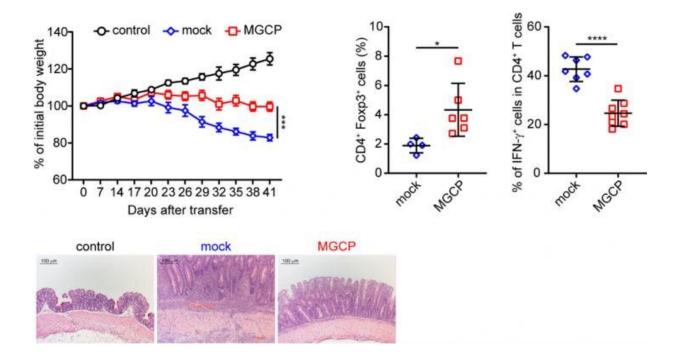
Various immune cells such as monocytes, macrophages, dendritic cells, and T cells are involved in the onset and development of these diseases, but T cells in particular play a pivotal role. Currently, agents that suppress the overall inflammatory response are used in clinical practice, but this method has major side effects including vulnerability to infections and there are no clear effective treatments as of now.

To this, the POSTECH researchers and the joint research team from ImmunoBiome Inc. focused on the microbiome and the active substances derived from them, which have a big impact on the development and regulation of the immune system. The joint research team isolated specific polysaccharides from yeast—one of the symbiotic microorganisms in the human body—and first observed their anti-inflammatory effects. Then, using high-performance liquid chromatography and nuclear magnetic resonance (NMR) techniques, the researchers identified the constituents and chemical structures of the polysaccharides and named it MGCP (Mannan/ β -1,6-Glucan-containing polysaccharides).





Anti-tumor immunity enhancing effects of cell wall β -1,3-glucan. Credit: POSTECH



Inhibitory effects of yeast-derived MGCP in inflammatory bowel disease. Credit: POSTECH



Using a <u>mouse model</u> for inflammatory bowel disease and multiple sclerosis, MGCP-administered mice suppressed the generation of inflammatory cells, which are T helper type 1 cells (Th1 cells). On the other hand, it selectively inhibited the progression of inflammatory diseases by inducing the generation of regulatory T cells (Regulatory T cells, hereinafter Treg cells) with anti-inflammatory function. The researchers additionally confirmed that the mechanism of action of the immunosuppressive response by MGCP is mediated by TLR2 and Dectin1, which are the two different innate immune receptors expressed in <u>dendritic cells</u>.

This study also presents a clear solution to an unsolved immunological question about beta-glucan (β -glucan), which is known to suppress hypersensitivity while inducing immune enhancement at the same time. The research team has uncovered that the immune response enhancing effect of β -glucan identified so far is due to β -1,3-glucan, and found that β -1,6-glucan that makes up the MGCP has a hypersensitivity immunosuppressive effect. These findings show that β -glucan of a specific structure is applicable as an immune enhancing or anti-inflammatory response inducer. These active substances show promise to be developed into the next-generation microbiome treatment.

This study revealed that the immunological efficacy is determined by the chemical structure of the obscure polysaccharide, which was largely unknown. It is significant that a novel <u>polysaccharide</u> MGCP that effectively inhibits inflammatory diseases has been discovered. "We have successfully demonstrated that MGCP can selectively suppress inflammatory T-<u>cells</u>," explained Professor Sin-Hyeog Im. "This will help to provide a great turning point for a new anti-inflammatory treatment with no side effects that can selectively suppress the inflammatory response."



More information: Changhon Lee et al, Structural specificities of cell surface β -glucan polysaccharides determine commensal yeast mediated immuno-modulatory activities, *Nature Communications* (2021). DOI: 10.1038/s41467-021-23929-9

Provided by Pohang University of Science & Technology (POSTECH)

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