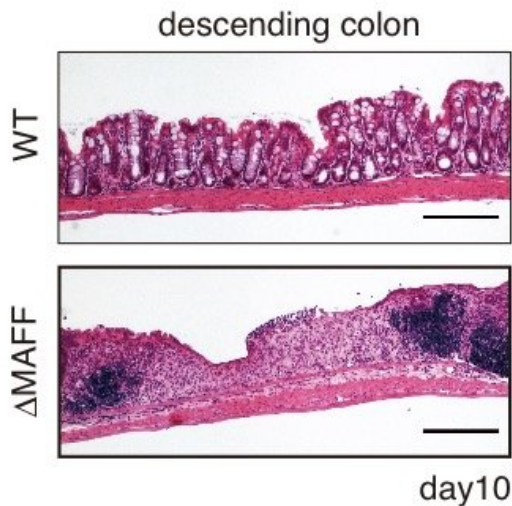


Host antibodies shape gut microbiome by changing bacteria gene expression

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Ten days after being treated with an intestine-damaging drug, the colons of mice inoculated with wild-type *B. theta* have largely recovered (top row). But the colons of mice inoculated with a *B. theta* strain lacking MAFF proteins still show signs of damage and intestinal inflammation (bottom row). Credit: Nakajima et al., 2018

Researchers at the RIKEN Center for Integrative Medical Science in Japan have discovered how antibodies secreted in the gut promote the growth of beneficial bacteria. Their study, which will be published July 24 in the *Journal of Experimental Medicine*, shows that immunoglobulin A (IgA) antibodies can alter the expression of bacterial genes, allowing different bacterial species to cooperate with each other and form a community that can protect the body from disease.

It is estimated that between two and five grams of IgA are produced in adult intestines each day. IgA is the first line of defense against harmful, [pathogenic bacteria](#) in the gastrointestinal tract, but it also promotes the growth of a healthy

microbiome—the community of "good" [bacteria](#) that reside in the gut. An average adult has over a thousand species of bacteria in their intestines that improve the function of the intestines and immune system, significantly impacting human health and disease.

Led by Keiichiro Suzuki, first author Akira Nakajima and colleagues at Kyoto University and RIKEN Yokohama Institute set out to investigate how IgA promotes the growth of healthy gut bacteria. Antibodies usually fight off [harmful bacteria](#) by recognizing specific proteins on the bacteria's surface. But Suzuki and colleagues found that even IgAs that recognize a non-bacterial protein can coat the surface of healthy gut bacteria because sugar molecules attached to the tails of IgAs can bind to a molecule called lipopolysaccharide that is found on many different species of bacteria.

The prominent human gut bacterium *Bacteroides thetaiotaomicron* (*B. theta*) was particularly susceptible to coating by IgA. Suzuki and colleagues discovered that exposure to IgA causes *B. theta* to up-regulate two proteins that were structurally shared with many other bacterial strains, and the researchers named these molecules Mucus-Associated Functional Factors (MAFFs). These proteins appear to help *B. theta* grow in the mucous membrane of the intestine and produce metabolites that allow other [beneficial bacteria](#), such as Clostridiales, to thrive as well. Mice inoculated with a strain of *B. theta* unable to up-regulate MAFFs had an altered gut microbiome and were more susceptible to intestinal inflammation, or colitis.

"We knew that immunoglobulin A contributed in some way to gut health, but it was exciting to discover this new mechanism—the MAFF system—that actually promotes symbiosis among the bacteria that inhabit the mucus membrane of the gut," Dr. Suzuki says. "The MAFF system is also present in humans," he continues, "so it is an

interesting target of research, but there is still much to be investigated. We need to identify its molecular target and find out how its expression is regulated along with neighboring genetic elements, with the hope that answering these questions could lead to the development of new methods for therapies or prophylaxis of [inflammatory bowel disease](#)."

More information: Nakajima et al., 2018. *J. Exp. Med.* [DOI: 10.1084/jem.20180427](https://doi.org/10.1084/jem.20180427)

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