# Here begins our new MIRAI

# Elucidation of the mechanism of serotonin over optimism and pessimism



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#### Summary of the project

The optimism that "I'm sure things will go well" is one of the essential mental vitalities for believing in an uncertain future and for overcoming difficulties and paving the way for your life. By 2050, this research aims to visualize this mental vitality and power to live, and to create a future in which we can further improve it ourselves.

This project aims to examine what kind of difference will occur in the serotonin neural network, which has been shown to play a role in regulating patience for future rewards, when mice perform the same behavior but the purpose of the behavior is different for "attainment of reward" or "avoidance of punishment". We hypothesize that serotonin works to regulate "optimism / pessimism" toward achieving the goal and we will examine the serotonin neural network by neural recording and neural manipulation of task performing mice. By clarifying the neural mechanism of "the optimism that creates patience" or "the pessimism that leads to giving up", we aim to realize a society in which people can improve their "ability to overcome the difficulties of life" and "vitality of the mind".

### Milestone by the end of project (year 2024)

Elucidation of the serotonin subsystem in the brain that is causally related to optimism (confidence in the future) and pessimism (giving up on the future).

## R&D theme structure of the project Observation Manipulation Fiber photometry Compact fluorescence microscope camer Optogenetics Na Activation DRN future reward or avoiding future punishment Brain regions of interest PM(Katsuhiko Miyazaki) Research and development 1 1. Observation of serotonergic activity in the dorsal raphe nucleus Fiber photometry (Kayoko Miyazaki) Compact fluorescence microscope camera (Katsuhiko Miyazaki) 2. Measurement of serotonin extracellular concentration at brain region of serotonin projection Fiber photometry (Kayoko) 3. Observation of neural activity in the brain region of serotonin Fiber photometry (Kayoko) Compact fluorescence microscope camera (Katsuhiko) Research and development 2 1. Behavioral changes by serotonergic manipulation of the dorsal raphe nucleus Optogenetics (Kayoko) 2. Changes in neuronal activity by serotonin manipulation (excitement/inhibition) Fiber photometry (Kayoko)

Compact fluorescence microscope camera (Katsuhiko)