

Deep-sea osmolyte makes biomolecular machines heat-tolerant

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Conceptual illustration showing deep-sea osmolyte TMAO stabilizing the interaction between microtubules and kinesins. Credit: Munmun, T. et. al., Chemical Communications, December 26, 2019

Researchers have discovered a method to control biomolecular machines over a wide temperature range using deep-sea osmolyte trimethylamine N-oxide (TMAO). This finding could open a new dimension in the application of artificial machines fabricated from biomolecular motors and other proteins.

Biomolecular motors are the smallest natural machines that keep living



organisms dynamic. They can generate force and perform work on their own by consuming chemical energy. In recent years, reconstructed biomolecular motors have appeared as promising substitutes of synthetic motors and expected to be key components in biomimetic artificial micro- or nano-devices. However, reconstructed biomolecular motors lose their ability to function due to thermal instability in artificial environments.

Tasrina Munmun, Arif Md. Rashedul Kabir, Kazuki Sada and Akira Kakugo of Hokkaido University and Yukiteru Katsumoto of Fukuoka University were inspired by seeing how proteins remain stable in living organisms such as sharks, teleosts, skates, and crabs that survive in harsh environments like deep sea hydrothermal vents or under thermal perturbations. Although proteins are generally denatured by heat, the proteins in deep-sea animals remain stable and active with heat thanks to TMAO.

"Based on this fascinating defense mechanism in <u>deep-sea animals</u>, we attempted to control the activity of kinesin, a biomolecular <u>motor</u> associated with microtubule proteins, over a wide <u>temperature</u> range," said Arif Md. Rashedul Kabir. To investigate the activity of kinesins, the team conducted in vitro motility assays in which <u>kinesin</u> motors propelled the microtubules on a two-dimensional substrate.





In the motility assay, microtubules are propelled by kinesin motors attached to the substrate surface. Credit: Munmun, T. et. al., Chemical Communications, December 26, 2019





TMAO is effective in prolonging the lifetime of microtubule-kinesin interaction over a wide range of temperatures. Credit: Munmun, T. et. al., Chemical Communications, December 26, 2019

According to the study published in *Chemical Communications*, they discovered that TMAO suppresses thermal denaturation of kinesins in a concentration dependent manner. Within a temperature range of 22-46 degrees C, kinesins propelled microtubules for a prolonged time (almost 2.5 times longer) when TMAO was present. This shows the team successfully controlled the dynamics between kinesins and microtubules over a broad temperature range. "This study is the first example showing successful utilization of a deep-sea osmolyte in maintaining



biomolecular motors for a prolonged time over a wide temperature range in engineered environments," Arif Md. Rashedul Kabir commented.

Arif Md. Rashedul Kabir continued, "The idea of utilizing natural defense mechanisms against heat-induced inactivation of proteins and enzymes will now be encouraged further."

"Our work will open a new dimension in sustainable applications of reconstructed biomolecules which will benefit various fields including biomimetic engineering, biochemical and <u>biomedical engineering</u> as well as materials science," Akira Kakugo added.

More information: Tasrina Munmun et al. Controlling the kinetics of interaction between microtubules and kinesins over a wide temperature range using the deep-sea osmolyte trimethylamine N-oxide, *Chemical Communications* (2019). DOI: 10.1039/C9CC09324A

Provided by Hokkaido University

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