

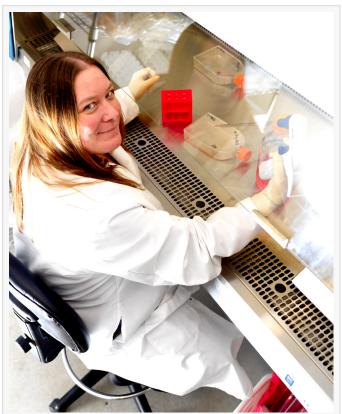
Circular Peptides in Violets May Aid in the Fight Against Glioblastoma

Circular peptides found in miniscule quantities in violets provide in vitro an over ten-fold increase in the efficacy of TMZ chemotherapy for glioblastoma.

JACKSON, WY, UNITED STATES, September 30, 2024 /EINPresswire.com/ -- Glioblastoma is one of the most serious brain diseases known. More than 45% of brain cancers are gliomas. Only half of glioblastoma patients respond to the FDAapproved chemotherapy Temozolomide (TMZ). Even for those patients, the cancer cells quickly evolve resistance. Most patients pass away within 12 to 16 months after diagnosis, and few make it beyond five years.

Now a glimmer of hope for patients comes from an unlikely place: Jackson Hole, Wyoming, where scientists at the non-profit <u>Brain Chemistry Labs</u> have been studying molecules found in violets.

Violets produce a dazzling suite of small circular peptides called cyclotides. They roughly appear in



Violet researcher Dr. Samantha Gerlach at Brain Chemistry Labs

shape "like floppy frisbees," says Dr. Samantha L. Gerlach. "They have been found active in the test tube against certain types of human cancer cells."

Disulfide crosslinks which maintain the shape of cyclotides may help them create pores in the membranes of cancer cells. Within the plant, cyclotides provide protection against insect herbivores, fungal infections, and viruses. Cyclotides were originally discovered from an herbal tea used by indigenous people in Africa to ease the course of childbirth. The tea was made from a plant they call kalata-kalata and which scientists call Oldenlandia affinis.

An international team led by scientists in Jackson Hole announced last week in the Swiss journal <u>biomedicines</u> that the cyclotide kalata B1 turbocharges the activity of the chemotherapy TMZ, decreasing the amount necessary to kill glioblastoma cells by over ten-fold. Senior author Dr.

Gerlach and her colleagues demonstrated that a synthetic version of kalata B1 has equal efficacy to the natural molecule.

"While kalata B1 commonly occurs in violet species, extraction from plant material yields only miniscule amounts," Gerlach states. "Working day and night for months, the minimal quantities we obtain are insufficient for clinical research."

Through a collaboration with CSBio in California, the scientists were able to obtain much larger quantities of the synthetic version sufficient for testing in mouse models of glioblastoma.



Wyoming violets

The structure and efficacy of synthetic kalata B1 was found to be equivalent in all respects to the naturally occurring molecule. Dr. Krish Krishnan at California State University, Fresno used Nuclear Magnetic Resonance (NMR) spectroscopy to confirm the shape and folding of the synthetic molecule. "Our cell data suggest that we can now move forward with the synthetic version in mice models," Dr. Rachael Dunlop at the Brain Chemistry Labs stated. This next step of testing in mice will occur in Vienna, Austria.

While Brain Chemistry Labs Director Dr. Paul Alan Cox believes that the advent of synthetic kalata B1 could be a major step forward, he is cautious about overstating the significance for patients. "We are still a long ways from clinical trials, but now the way is clear to determine if it might be safe for further testing."

Marya King Brain Chemistry Labs +1 224-358-6578 marya@ethnomedicine.org

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