

Tumour blood supply stopped in its tracks by modified natural compound

August 10 2017, by Carrie Bengston



Dr Orazio Vittorio of Children's Cancer Institute. Credit: University of New South Wales

Researchers have discovered how the modified natural compound dextran-catechin disrupts formation of blood vessels that fuel growth in the childhood cancer neuroblastoma.



Researchers have discovered how a modified <u>natural compound</u> disrupts angiogenesis, the formation of <u>blood vessel</u> networks, in neuroblastoma tumours, stopping them laying down the vital supply lines that fuel cancer growth and spread.

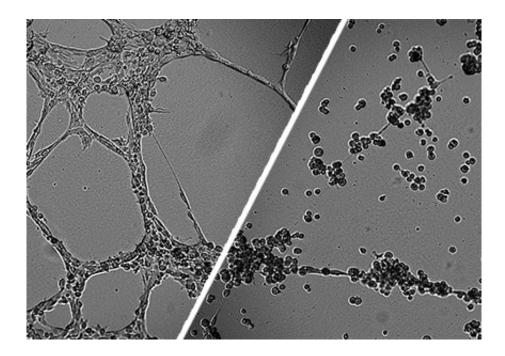
The international study, led by scientists at Children's Cancer Institute and UNSW, is published in the journal *Scientific Reports* and paves the way for less toxic treatments for neuroblastoma, a childhood cancer with an average age of diagnosis of just one to two years old.

Lead author Dr Orazio Vittorio of Children's Cancer Institute found in 2012 that the natural polyphenol catechin slows tumour growth in the laboratory but breaks down too quickly in the body to be effective.

"We joined catechin with a sugar called dextran. We found this dextrancatechin complex is much more stable in the body and that it slows <u>tumour growth</u> by affecting copper levels – but we didn't know precisely how," he says.

The new research showed that copper is needed by <u>endothelial cells</u>, the cells that line <u>blood vessels</u>, and that dextran-catechin disrupts the cells' copper levels in several ways.





Images of untreated cells (left) and dextran-catechin treated cells (right). Credit: University of New South Wales

"Dextran-catechin inhibits specific copper transport proteins in the cells that line the blood vessels, altering intracellular copper levels. This prevents the cells joining together to form the network of vessels that supplies tumours with blood to keep growing" he says.

The result is clearly visible down a microscope. When <u>human endothelial</u> <u>cells</u> were cultured and treated with dextran-catechin, the normally branching networks of blood vessels failed to form properly. Likewise, in mice, neuroblastoma tumours treated with dextran-catechin had significantly fewer blood vessels than tumours treated with saline control.

"Instead of forming a neat, branching network of blood vessels, you see a mess of cells all over the place, which means cancer cells can't get the blood supply they need," says Dr Vittorio.



"This is exciting because it's a new target for the childhood cancer neuroblastoma that appears safe and has minimal side effects."

Copper is needed to switch on genes that help endothelial cells grow and form blood vessels during angiogenesis. The study found that dextrancatechin targets specific proteins that control copper uptake and removal in endothelial cells, reducing the copper available to switch on the genes.

Further laboratory research into dextran-catechin's potential as a future treatment is underway.

More information: Eugene M. H. Yee et al. Dextran-Catechin inhibits angiogenesis by disrupting copper homeostasis in endothelial cells, *Scientific Reports* (2017). DOI: 10.1038/s41598-017-07452-w

Provided by University of New South Wales

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