

## More targeted, less toxic: The golden future of cancer treatment

September 6 2019



Distinguished Professor Suresh Bhargava and Dr Neda Mirzadeh, co-leaders of the Molecular Engineering Group at RMIT University. Credit: RMIT University

Researchers have engineered gold-based molecules that target cancer cells and leave healthy cells unharmed, in a critical step towards



precision cancer drugs with fewer toxic side effects.

Pre-clinical studies have shown the molecules were up to 24 times more effective at killing cancer cells than the widely used anti-cancer drug cisplatin and were also better at inhibiting tumour growth.

The molecules were also more targeted and selective, making them promising candidates for development into a new class of gold-based drugs that can wipe out the cancer without destroying <u>healthy cells</u>.

Significantly, the synthetic molecules are built with resistance-fighting features to keep them effective over time, unlike current chemotherapies.

The study by researchers at RMIT University, detailing four new bioactive molecules and their effectiveness against five types of cancer cells, is published in *Chemistry—A European Journal*.

Dr. Neda Mirzadeh, co-leader of the Molecular Engineering Group at RMIT, said the limitations of commonly used metal-based chemotherapies drove a need to find better alternatives.

"We've made huge inroads in <u>cancer treatment</u> over recent decades, but the disease still kills over 9.6 million people globally each year and remains the world's second leading cause of death," Mirzadeh said.

"While metal-based drugs have successfully pushed survival rates up, their effectiveness is limited because of toxic side effects, drug resistance and poor stability.

"The gold-based molecules we've designed are far more selective and stable.



"Our results show there's incredible potential here for the development of new cancer-fighting therapeutics that can deliver lasting power and precision."

The molecules were evaluated in pre-clinical tests and found to be highly cytotoxic against prostate, breast, cervical, melanoma and colon cancer cells. Animal trials showed treatment with the molecules significantly inhibited tumour growth (up to 46.9%, compared with 29% for cisplatin).

The gold-based compounds also inhibit the action of an enzyme found in <u>cancer cells</u>, thioredoxin reductase, that is linked with <u>cancer</u> progression and the development of drug resistance.

In addition, the molecules have strong anti-inflammatory properties, giving them a dual therapeutic effect and potential application in the treatment of chronic inflammatory conditions like arthritis.

## Molecular design pioneers

Part of RMIT's Centre for Advanced Materials and Industrial Chemistry (CAMIC), the Molecular Engineering Group brings together synthetic chemists with pharmacologists and has over two decades of expertise in designing gold molecules for practical applications.

The pioneering group, one of just a few in the world focusing on the medical potential of gold-based molecules, has had two patents awarded for their work to date.

Team leader and CAMIC Director, Distinguished Professor Suresh Bhargava, said gold had an ancient history of medical use, particularly in India and China.



"We know that gold is readily accepted by the human body and we know it has been used for thousands of years in treating various conditions," Bhargava said.

"Essentially, gold has been market tested, but not scientifically validated.

"Our work is helping both provide the evidence base that's missing, as well as delivering new families of <u>molecules</u> that are tailor-made to amplify the natural healing properties of gold."

The research team has completed successful in vitro and in vivo preclinical studies and is seeking funding to support the next stage of the research—clinical studies and regulatory approval.

The team includes Dr. Srinivasareddy Telukutla, a graduate from RMIT's joint Ph.D. program with the Indian Institute of Chemical Technology (IICT), Ganga Reddy Velma, a Ph.D. candidate at IICT and Visiting Research Fellow at RMIT, and synthetic chemist Dr. Steven Priver.

The paper, Potent and selective cytotoxic and anti-inflammatory gold(III) compounds containing cyclometallated phosphine sulfide ligands, with co-authors from the IICT and University of Melbourne, will feature on the cover of a forthcoming edition of *Chemistry—A European Journal*.

**More information:** Suresh Kumar Bhargava et al, Potent and selective cytotoxic and anti-inflammatory gold(III) compounds containing cyclometallated phosphine sulfide ligands., *Chemistry - A European Journal* (2019). DOI: 10.1002/chem.201903388



## Provided by RMIT University

Citation: More targeted, less toxic: The golden future of cancer treatment (2019, September 6) retrieved 3 May 2024 from

https://medicalxpress.com/news/2019-09-toxic-golden-future-cancer-treatment.html

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