

# **N2CR INSIGHTS** THE QUARTERLY NEWSLETTER OF N2CR

## LATEST NEWS



Find us at MD6 Foyer 2A

Click here for research opportunities

**MBBS undergraduates** 

Are you passionate about medical research and eager to

make a difference in the field of cancer research?

Join us at the N2CR booth to explore exciting research

opportunities that can shape your future career!

Page 1

Research News Pages 1-3

Highlights of 2024 Pages 3-4

Upcoming Events & Announcements Page 4

Follow us on social media!



## **RESEARCH NEWS (OCT - DEC 2024)**

## **Revolutionising Nanomedicine: DNA Barcodes for Gold Nanoparticles**

(Advanced Functional Materials; Xingyue Huang, Zhicheng Le ... Xuehao Tian, Andy Tay)



Cancer Researchers'

**Stories** 

interview with our

talented member,

A/Prof Deng Lih

Wen from the Dept

of Biochemistry.

with

is

an

and

N2CR's series

back

esteemed

**Read More** 

Gold nanoparticles (NPs) are promising for cancer therapy due to their versatile properties. However, their interactions with cells vary, affecting reproducibility and application. To optimise this, Dr Andy Tay Kah Ping from N2CR and his team, implemented a DNA barcoding system to label NPs of different shapes (sphere, rod, triangle) and sizes (40 nm, 80 nm). Barcodes remained attached and didn't interfere with cell interactions. Results showed that sphere NPs had poor in vitro uptake but good in vivo targeting, while the 80 nm nanotriangles worked well in both lab tests and in living organisms. These NPs were effective in siRNA delivery and photothermal therapy in a breast tumour model, highlighting their potential to enhance cancer treatment.

### ZBTB48: A New On-Switch for the Immune Response

(The EMBO Journal; Grisma Rane, Vivian Li Siew Kuan, ... Fudong Li, Dennis Kappei)

CIITA is the master regulator of MHC class-II immune genes that are important for defense mechanism against infections and cancer development. CIITA is controlled by three specific switches used in a cell-type specific manner and can be turned on by a signal called IFN- $\gamma$ . In recent work, research fellow Dr Grishma Rane and colleagues from the lab of N2CR member Dr Dennis Kappei found that a protein called ZBTB48 binds to the B-cell switch and activates CIITA in this type of white blood cells. When ZBTB48 is missing, CIITA and MHC-II gene activity are reduced and cannot be turned on even after signaling by IFN- $\gamma$ . The team will next apply this fundamental science discovery to B-cell malignancies. Watch this space!



**Read More** 

## **Magnetic Field Therapy Improves Breast Cancer Treatment with Fewer Side Effects**

(Cancers; Viresh Krishnan Sukumar, Yee Kit Tai, ... Joline Si Jing Lim, Alfredo Franco-Obregon)



Read More

Chemotherapy, although the first line of defense in cancer treatment, is indiscriminate and is damaging to the entire body, which, ironically, worsens a patient's chances of getting better from cancer. Doxorubicin is a widely used chemotherapy drug for breast cancer but has significant side effects due to its systemic delivery.

This study led by N2CR member A/Prof Alfredo Franco-Obregon with co-first authors, Viresh Krishnan Sukumar (N2CR PhD Scholarship student) and Dr Alex Tai Yee Kit, found that a 10-minute magnetic exposure increases doxorubicin uptake in breast cancer cells without harming the surrounding healthy cells. The study suggests a localised, non-invasive magnetic therapy to enhance doxorubicin efficacy with fewer side effects. This may ultimately allow for the lowering of systemic doxorubicin administration that could improve a patient's outcome against cancer. This would be a potential game changer in cancer treatment.

### **Extracellular Vesicles: Key Players in Tumour Microenvironment and Drug Resistance** (Theranostics; Jayshree Hirpara, Win Lwin Thuya, ... Shazib Pervaiz, Boon Cher Goh)

This research, led by N2CR members Prof Goh Boon Cher and Prof Shazib Pervaiz with Dr Jayshree Hirpara as the first author, investigates the role of extracellular vesicles (EVs) in the tumour microenvironment and its contribution to drug resistance. By isolating and analysing EVs from drug-resistant cells and plasma, the study revealed high levels of metabolic transporters like SLC1A5 which are associated with resistance to Tyrosine Kinase Inhibitors (TKIs), a class of targeted cancer therapy.

The results demonstrated that EVs from resistant cells can induce resistance in sensitive cells and transform fibroblasts into tumour-associated fibroblasts, thereby enhancing their invasive capabilities. These findings highlight the significance of EVs in promoting therapy resistance and suggest potential diagnostic and therapeutic targets.



**Read More** 

## **Understanding RNA Changes and Cancer Development**

(Proceedings of the National Academy of Sciences of the United States of America; Jian Han ... Sze Jing Tang, Leilei Chen)



Led by N2CR member A/Prof Polly Chen, this research highlights the crucial role of 'death associated protein 3' (DAP3) in regulating N6-methyladenosine (m6A) RNA methylation, a process often disrupted in cancers. DAP3 is frequently overexpressed in cancer cells to promote tumour growth. It maintains m6A levels by helping methyltransferase-like 3 (METTL3) bind to RNA and promoting the splicing of a protein called MAT2A, which increases S-adenosylmethionine (SAM) levels. This study enhances our understanding of how m6A dysregulation contributes to cancer, highlighting a potential new target for cancer treatment.

## **RESEARCH NEWS (OCT - DEC 2024)**

## **Unveiling p53's Role in DNA Protection: New Cancer Insights**

### (Nucleic Acids Research; Gamal Ahmed Elfar, Obed Aning ... Ying Swan Ho, Chit Fang Cheok)

Research led by N2CR member Dr. Cheok Chit Fang found that the tumour suppressor protein p53 protects DNA during replication by preventing the hyperactivation of PARP1 caused by oxidative stress. In the absence of p53, cells experience increased replication stress and DNA damage. The study revealed that p53 collaborates with the RRM2B gene to prevent this damage. Without both p53 and RRM2B, cells activate an antioxidant stress response involving NRF2, leading to PARP1 activation and degradation of newly replicated DNA. The enzyme G6PD was also identified as a key player in this process. This deeper understanding of the interactions between p53, PARP1, and redox metabolism could provide valuable insights for cancer treatments.



**Read More** 



## **ECI Symposium**

ECI held its first symposium from 13 to 14 May 2024, featuring Guest of Honour Prof Tan Chorh Chuan, who delivered an inspiring address, keynote speakers, and numerous local & international experts.

## Speakers

Pierce CHOW National Cancer Centre Singapore Jeffrey HALTES University of Michigan, USA George HANNA Imperial College London, UK Milaed ILARTMAN National University Hospital, Singapore LJ Jingmei GIS, A\*STAR, Singapore LIM Chower Teck NUS, Singapore Robert SCHARPF Johan Hopking Bloomberg School of Public Health, USA

NUS second

SILAO Hai Lin NUS, Singapore TAM Wai Leong GIS, A'STAR, Singapore TAN Bee Huat Iain National Cancer Centre Singapore TAN Ker Kan National University Hospital, Singapore Cristic for Cancer Prevention and Early Detection, USA TOO Henge Phon NUS, Singapore Abole VENETIABAMAN NUS, Singapore from Laboratory Discoveries to the Community

13 - 14 May 2024 NUS MD6 Lecture Theatre 35



lest of Honour

YEOH Khay Guan Nulis, singapata Kaynate Speaker Nitzen ROSENFE University of Camerica Keynote Speake



Eleynote Speake

Our retreat at Mount Faber on 8 March 2024 was a blast!

We kicked things off with a scenic cable car ride, enjoyed stunning views, delicious food, and had some very lively discussions.

Can't wait for the next one later this year with more camaraderie and scientific synergies!



#### **N2CR INSIGHTS**

### **RESEARCH HIGHLIGHTS 2024**

'New CAR T-Cell Therapy Offers Hope for Relapsed or Refractory T-cell Leukaemia'

Nature Medicine, 3 Sep 2024

A new CAR T-cell therapy, led by N2CR members, shows promise in treating treatment resistant Tcell acute lymphoblastic leukaemia (T-ALL), with most patients achieving remission and acceptable side effects.

#### **Prof Allen Yeoh and Prof Dario Campana**

**Read More** 



#### 'Clinical Trial Shows Promise in Treating Advanced Gastric Cancer' ESMO Open, 17 Sept 2024

A recent clinical trial found that combining aerosol chemotherapy (PIPAC-OX) with an immunotherapy drug (nivolumab) was safe and effective for treating advanced gastric cancer, reducing cancer and boosting the immune response.

#### **Prof Jimmy So**

**Read More** 



'Missing link between poor diet and higher cancer risk uncovered' Cell, 25 April 2024

Research has unearthed new findings which may help explain the connection between cancer risk and poor diet. The insights gained from this study hold promise for advancing cancer prevention strategies aimed at promoting healthy ageing.

Prof Ashok Venkitaraman and Dr Kong Li Ren

**Read More** 



### 'A New Pathway in Colorectal Cancer Cells Triggered by Drug-Activated Mutant KRAS' Autophagy, 25 Feb 2024

Researchers exploited cancer cells' addiction to mutant KRAS oncoprotein by demonstrating hyperactivation-induced mitochondrial damage and cell death, which provides a novel therapeutic strategy against KRAS-driven cancers.

**Prof Shazib Pervaiz** 

动

**Read More** 

## **UPCOMING EVENTS & ANNOUNCEMENTS**



"Cell Fate and Its Regulation In Health and Disease"

MAY 28 - 30, 2025 MD11, CRC Auditorium, YLLSOM, NUS, Singapore

AWARD LECTURES

TY CHEPUK navi af thei IN PERVAIL eel of Heckins, Holio by of Engagers, Sings

INVITED SPEAKERS Yunnus ARCOC

Ladislay ANDERA TU ADAMA ne Sophie AlthAND

ARSENIAN HENRIGSON NO DIATT

the set lines

---Out Farg CHECK sha of Lines hen CHONG d of Mark CHUA International C DIEDENION **Hares** -Cher GON HUNNE RATZ KENNEDY INRYSKO

ring Yo LIANG COLUMN DESIGNATIONS NUT PLATTER ick Sek Tong OND Frank OURY of Street, or other in Francis IN LABARATHY AU SHAZE **LUNA** BY TERGADORAR ATCH AR, Disapports IND. YIS TEAL

ige YV

### Let's Welcome our Primary Members (from 1 Jan 2025)



Dr Tan Kar Tong Dept of Pharmacy & Pharmaceutica Sciences, NUS



Dr Tan Yong Zi Dept of Biological Sciences, NUS

visit our website

# Visit our Site-Specific Resources for the following services

Please click on services **Tissue Requisition and** for further information. **Tissue Microarray (TMA) For Consultation Services on Experiment Histology Services** Set-Up and Protocol **Discussion**, please email **Facility Booking** radhika@nus.edu.sg or

N2CR INSIGHTS