

PROF. WOJCIECH CHRZANOWSKI

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"Professor Wojciech Chrzanowski is a nanobiomedical engineer who translates nanoscale science to human applications and transforms medical treatments and disease diagnosis."

CURRENT APPC	INTMENTS	
2021 – present	Professor of Nanomedicine, Faculty of Medicine and Health, The University of Sydney (0.8 full-time equivalent)	
2019 – present	President of High-Intensity Focused Ultrasound Biomedical Association Inc.	
2015 – present	Vice President of the Asian Federation for Pharmaceutical Sciences	
2010 – present	Founder and Director of the Nano-Bio-Engineering Group and the Nano-Bio- Characterisation Facility, Sydney Pharmacy School, The University of Sydney	
PREVIOUS APPO	DINTMENTS	
2016 – 2021	Deputy Director at The University of Sydney Nano Institute (0.2 full-time equivalent)	
2015	Research Fellow in Medicine, Harvard Medical School Harvard University	
2008 – 2009	Research Assistant, School of Mechanical Engineering, University of Glasgow, EU Framework Program 6 Project Grant	
2006 – 2008	Marie Curie Research Fellow, Eastman Dental Institute, Division of Biomaterials and Tissue Engineering, University College London	
2003 – 2006	Research Assistant/Lecturer,.	
ACADEMIC QUA	ALIFICATIONS	
2006 – 2014	Doctorate of Science in Biomedical Engineering, Polish Academy of Science, Nałęcz Institute of Biocybernetics and Biomedical Engineering, Poland	
2013	Graduate Certificate in Educational Studies, The University of Sydney	
2000 – 2003	Doctor of Philosophy in Biomedical Engineering, The Silesian University of Technology, Poland	
1995 – 2000	Master of Science in Biomedical Engineering, The Silesian University of Technology, Poland.	
GRANTS		

GRANTS

Since 2006 I have obtained \$14. 8M in competitive grant funding including: EU Horizon2020 (2×), Australian Research Council Linkage Projects (3×), NHMRC equipment grants (6×), Sydney Catalyst Grant (2×), Medical Advances Without Animals Research Grant (3×), and Harmonia Project Grant, National Science Centre Poland (1×). I was also awarded \$850k amount in internal competitive funding including a Sydney Outstanding Academic Researcher (SOAR) Prize (\$150k), Sydney Nano Grand Challenge Projects (2×), Sydney Nano Kickstarter Grants (2×) and \$160k in partnership and travel grants.

FELLOWSHIPS

Japanese Society for Promotion of Science Invitational Fellowships – Chubu University (2014), Tokyo University (2012)

Marie Curie Intra-European Fellowship (2006), University College London – European Union Framework Program 7

AWARDS	
2021	Highly Commended Finalist, Australia Research Awards, Category Frontier
2020	Winner of the Biomedical Shark Tank Competition – Institute for Biomedical Materials and Devices, University of Technology, Sydney
2019	Barry Inglis Medal for Innovation in Nanometrology – National Measurement Institute
2019	Sydney Nano Institute's Publication Award in the field Nanotoxicology

2019	Award for being Outstanding Reviewer for Nanoscale Horizons – Royal Society of Chemistry
2018	Winner of the Big Idea Award Competition – Sydney Local Health District
2018	Vice Chancellor's Award for Excellence (Outstanding Teaching and Research) – The University of Sydney
2018	2 nd Most Influential Scientific Discovery – The University of Sydney
2018	Outstanding Research Paper Award – Royal Society of Chemistry
2018, 2016	Supervisor of the Year Award – Sydney University Postgraduate Research Association
2017	Deputy Vice Chancellor Research Award for Reduction in the Use of Animals in Research (Magnetically bioprinted three-dimensional scaffold-free liver model for nanotoxicity studies) – The University of Sydney
2015	Endeavor Executive Award in the field of Nanotoxicology – The Department of Industry, Innovation, Science, Research and Tertiary Education of the Australian Government
2014, 2014	Recipient of Australian Institute Nuclear Science and Engineering Research Awards in the field of Biomedical Engineering

PUBLICATIONS

- 218 peer-reviewed publications, 3 books, 6 book chapters and inventor of 6 patents; ORCID: https://orcid.org/0000-0001-8050-8821
- H-index: **41**; i10-index: 115. My h-index doubled in the last 5 years.
- Field Weighted Citation from the last 3 years is 5.29 4.1x the Australian average and 5.2x the global average for the field of Medical and Health Sciences.
- 48% of my papers have been published in journals classified in the top 10% (Scientific Journal Rankings SJR) in the field: Nature Biotechnology 2021 2nd in Biotechnology, ACS Nano 2021 1st in Nanotechnology, Analytical Chemistry 2019, 2016 1st in Analytical Chemistry, 7th Most Influential Journal Across the Board according to Innovation Index; Nature Protocols 2019 in the top 1% of journals in biochemistry; Advanced Functional Materials 2014 2nd in Biomaterials; Biomaterials 2017, 2016, 2014 4th in Biomaterials.
- I am the first or senior author on over 69% of my publications.

SEMINARS

- 65 invited seminar/conference/workshop presentations in total, 46 of which were international.
- 23 invited conference presentations in the last 5 years.

OUTREACH

- International TV, Radio, and newspaper interviews viewed over 700M times, *e.g.*, The Guardian, Project, Ch9 News (Exclusive), Ch7 News, ABC News, ABC Radio National, Sydney Morning Herald, The Age, Engineers Australia).
- VIVID Sydney 2014 I exhibited my work at the Australian Museum of Contemporary Arts.

SERVICE TO THE DISCIPLINE

- Editor in Chief Artificial Cells, Nanomedicine and Biotechnology.
- Associate Editor Nanomaterials, Journal of Tissue Engineering.

CURRENT INDUSTRY COLLABORATIONS/ENGAGEMENT

BiomeCentric (AUS), MedLab Clinical (AUS), Aspect Biosystems (Canada), Probiotics Australia (AUS), Nano Medical Technologies (AUS), Kinaltec (AUS), TheraKii (AUS), IZON (NZ), BCAL (AUS), SynThera (India).

Selected Grants

Title				
Funding agency / number	CI(s)	Amount funded	Years	
Title: Integrated Assessment and Advanced Characterization of Neuro-Nanotoxicity				
Title: Integrated Assessment ar	d Advanced Characterization of	Neuro-Nanotoxicity		

Gianni Ciofani, James Baker, Sean Kelly, Alberto Katsumiti, Isabel Rodríguez, Adrián García, pus five industry partners.

This program establishes the world's first integrated approach for the neuro-nanotoxicity assessment that understands the interconnected in vivo–in vitro relationship holistically.

Title: EV-Phage Biobots			
CDIP Fund	Wojciech Chrzanowski, Hien Duong	A\$ 69,000	2022-2023

This program pioneers the development of multifunctional biologicals that simultaneously eradicate bacteria and heal wounds.

Title: Shaping Innovative Products for Sustainable Tissue Engineering Strategies – SHIFT

Horizon 2020 Framework Programme, Call: H2020- MSCA-RISE-2020	Antonella Motta, Rui Reis, Alicia El Haj, Anthony Weiss, Wojciech Chrzanowski, Gilson Khang, Natalia Nalves, Rui L. Reis, Sorada Kanokpanont, Turmunkh	A\$ 1 540,000	2021-2024
	Gerelchuluun		

This program focuses on the development of new strategies for hard and soft tissue regeneration and wound healing.

Title: Breathe Easy - Development of the next generation treatment for chronic obstructive pulmonary
disease COPD using a COPD-on-chip model that replaces the use of animal models in lung disease and lung
injury research including COVID-19,Medical Advances WithoutWojciech Chrzanowski, Thanh
PhanA\$ 25,0002020-2021GrantKatter State State

This program aims to develops new models mimicking lung pathophysiology for testing new formulation to treat COPD.

Title: nanoJECT - light-thrusted needleless injections for pain-free cannabinoids delivery				
Office of Global	Chrzanowski W, Oh K,	A\$ 35,000	2019-2020	
Engagement/Partnership	Divakarla S, Hyeonwoo L			
Collaboration Awards	(International partner:			
	Yonsei University)			

This program focuses on the development of optic fibre based devices for pain free and needles injections of drugs.

Title: HIFU n - Nanotechnology-Enhanced High Intensity Focused Ultrasound as the New Generation ofCancer Therapy: advancing HIFU towards mainstream, non-invasive cancer therapySydney CatalystWojciech ChrzanowskiA\$ 45,0002019-2020

This program develops nano enhanced ultrasound system for cancer therapy.

Title: Skin- and skin injury-on-chip-microfluidic platform for Nano toxicity, drug discovery, and precisionbiology to replace the traditional use of animals in biomedical research and testingMedical Advances WithoutChrzanowski W, Divakarla S, A\$ 25,000Animals (MAWA)/ResearchGrant

This program develops new models of skin and skin injury for testing toxicity of nanoparticles and drugs.

eaking barriers in cancer pain trea	tment using iontophore	etic tattoo-like
	14 05 000	0040.0055
	A\$ 35,000	2018-2020
Chen S, Hall S		
(International partner:		
Chinese University of Hong		
ound-based system for transde		peutics which
Wojciech Chrzanowski, Sally	A\$ 45,000	2018-2020
Yunsun Kim		
erosol device for nulmonary de	livery of extracellular	vesicles
	-	
Wojciech Chrzanowski	A\$ 75,000	2018-2020
		2010 2020
os microneedle patch for transc	dermal devilry of cann	abinoids.
articles for controlled-release dru	gs via inhalation	
	-	
-	/\\$+00,000	2018-2021
ative nanoparticles which enca g.	psulate single drug cr	ustal and allow for
noparticles and their impact on h	ealth and environment	
Elizabeth New, Wojciech	A\$50,000	0040 0040
Chrzanowski	1 ,	2018-2019
gle nanoparticles characterisat	ion capability for test	ing nanotoxicity.
Wojcjech Chrzanowski	A\$30,000	
	Αψου,ουυ	2018-2019
	_	ar vesicles and wa
	n at master level	
eering - EU Australian cooperatio		
- · ·		+ 2014 2017
Claudio Migliaresi	197,500 Euro	+ 2014-2017
Claudio Migliaresi Rui L. Reis	197,500 Euro \$325,000 =	+ 2014-2017
Claudio Migliaresi	197,500 Euro	+ 2014-2017
Claudio Migliaresi Rui L. Reis	197,500 Euro \$325,000 =	+ 2014-2017
Claudio Migliaresi Rui L. Reis Thomas Groth	197,500 Euro \$325,000 =	+ 2014-2017
	Chrzanowski W, Zhao N, Divakarla S, Vitetta L, Liu J, Chen S, Hall S (International partner: Chinese University of Hong Kong) Dund-based system for transder g concentration in the blood stru- Wojciech Chrzanowski, Sally Yunsun Kim erosol device for pulmonary de le delivery platform for cannabino Wojciech Chrzanowski os microneedle patch for transder articles for controlled-release dru Chan H, Cipolla D, Chrzanowski W ative nanoparticles which encap g. noparticles and their impact on h Elizabeth New, Wojciech Chrzanowski gle nanoparticles characterisati tory extracellular vesicles via aero Wojciech Chrzanowski, Marca Wauben	Divakarla S, Vitetta L, Liu J, Chen S, Hall S (International partner: Chinese University of Hong Kong) Dund-based system for transdermal delivery of therage (concentration in the blood stream. Wojciech Chrzanowski, Sally A\$ 45,000 Yunsun Kim erosol device for pulmonary delivery of extracellular te delivery platform for cannabinoids for cancer patients Wojciech Chrzanowski A\$ 75,000 Dis microneedle patch for transdermal devilry of canner articles for controlled-release drugs via inhalation Chan H, Cipolla D, A\$456,000 Chrzanowski W ative nanoparticles which encapsulate single drug cr (chrzanowski gle nanoparticles characterisation capability for test tory extracellular vesicles via aerosolisation for treatment Wojciech Chrzanowski, A\$30,000

Title: Design clean technologies f	or the synthesis and purification	of biohybrid polymers	
ARC LP LP120200489	Fariba Dehghani, Wojciech CHRZANOWSKI	\$336,000	2013-2016
Title: Engineering of intelligent inh triggered release of active pharma			
National Science Centre (Poland), Project Grant Harmonia	Elzbieta Pamula Wojciech Chrzanowski Marek Langner	\$310,000	2014-2017
Title: 'Firefighters' for smoke inha	lation injury		
DVC-R Compact Grant	Wojciech Chrzanowski Kim Chan Yiwei Wang Sally Kim	\$25,000	2016
Title: 3D cell explorer – quantita	tive 3D stain-free, high-resolutio	n imaging of cells and	tissues
NHMRC Equipment Grant	Wojciech Chrzanowski Nicholas King 1Anthony Weiss Jackob George Alan Body	\$39,500	2016
Title: ElastoSensTMBio2 – non-d cultures	estructive & contactless biophys	sical tests of soft biom	aterials and cell
University of Sydney Equipment Grant	Fariba Dehghani Wojciech Chrzanowski Geraldine O'Neil Robyn McConchie	\$55,000	2016
Title: NanoSafety – new generatio drug delivery studies	n of 3D scaffold-free tissue moc	lels for high throughpu	t nanotoxicity and
DVC Research/AINST Accelerator Scheme	Wojciech Chrzanowski Nicholas King Iqbal Ramzan Brian Hawkett Dipesh Khanal	\$94,500	2015
Title: Cancer invasion and metast environments	tasis: how actin networks contro	l cell movement in 3 d	imensional
SPARC-Cancer	Geraldine O'Neill Matheo Biro Fariba Dehghani Wojciech Chrzanowski	\$124,000	2015
Title: Focus on Nano-Antibiofilm- Associated with Biomedical Devic		to Examine and Comb	oat Infections
The São Paulo Research Foundation (FAPESP). SPRINT – São Paulo Researchers in International Collaboration FAPESP grant: 2015/50311-8	Carlos Eduardo Vergani (UNESP) Wojciech Chrzanowski (USyd)	\$19,500	2015

Title: Kicking, scorching and coc	ling cells to guide their fate – u	nderstanding effects of	vibrations and
temperatures on a single cell res	• •	•	
JSPS Invitational Fellowship	Wojciech Chrzanowski	\$21,000	2014
Title: Externally activated drug e	luting implants/devices		
DVC Research Bridging Support Grant	Wojciech Chrzanowski	\$20,000	2014
Title: Nanoparticle drug Carriers	for Externally Triggered and Ta	rgeted Chemotherapy	
Sydney Catalyst Pilot Funding	David McKenzie Natalka Suchowerska Wojciech Chrzanowski Joanne Toohey Lisa Horwath	\$50,000	2014
Title: Cell-based therapy for pul	monary injury		
USyd Faculty of Pharmacy, Innovation Challenge Award Grant	Wojciech Chrzanowski Kim Chan Sally Kim	\$30,000	2014
Title: Multifunctional surface for	implantable devices		
AINSE research award Australian Institute of Nuclear Science and Engineering (AINSE)	Wojciech Chrzanowski	\$58,000	2013
Title: nanoIR - nanoscale bioima	ging with continuous mapping	of chemical and physica	al properties
NHMRC Equipment Grant	Wojciech Chrzanowski, Filip Braet Marcela Bilek Kim Chan Sebastian Perrier Geraldine O'Neil Alaina Ammit Fariba Dehghani	\$196,151	2013
Title: Naturally good - fibres and and drug delivery	particles of natural origin as a	new biomaterials for reg	generative medicine
Australian Government Department of Education and Training	Wojciech Chrzanowski	\$31,000	2013
Endeavour Executive Award			
Title: Scanning Ion Conductance	Microscopy		
The Ramaciotti Establishment and Equipment Grants Ramaciotti Foundation	lqbal Ramzan Wojciech Chrzanowski	\$75,000	2012
Title: Talking to cells - biointerfa	ce as a key parameter in comm	nunication with cells	
DVC International/IPDF Grant	Wojciech Chrzanowski David Brown Fariba Dehghani Hae-Won Kim	\$16,000	2012

Title: Molecular force probe for nanoscale bioimaging – MFP-3D-BIO					
NHMRC Equipment Grant	Wojciech Chrzanowski Filip Braet Marcela Bilek Kim Chan Sebastian Perrier Paul Young Daniela Traini Alexey Kondyurin	\$197,555	2012		
Title: Bioimprinted implant sur	faces with multifunctional prop	perties			
Australia-Korea Early Career S&T Researchers Program	Wojciech Chrzanowski Hae-Won Kim	\$4,800	2011		
The Australian Academy of Science					
Title: 'Braille' for cells – functional surfaces that enhance implant integration					
JSPS Invitational Fellowship	Wojciech Chrzanowski Tadashi Kokubo	\$18,900	2011		
Title: qNano – scanning ion oc	Title: qNano – scanning ion occlusion spectroscopy (SIOS)				
NHMRC Equipment Grant	Wojciech Chrzanowski Mary Bebawy Kim Chan Ramin Rohanizadeh Brian Hawkett	\$16,000	2011		
Title: The Instron testing material instrument					
NHMRC Equipment Grant	Fariba Dehghani Tony Weiss Marcela Bilek Hala Zreiqat Colin Dunstan Wojciech Chrzanowski Ramin Rohanizadeh	\$64,000	2011		

PUBLICATIONS

- 1. Calder, D.; Fathi, A.; Oveissi, F.; Maleknia, S.; Abrams, T.; Wang, Y.; Maitz, J.; Tsai, K. H.-Y.; Maitz, P.; Chrzanowski, W., **Thermoresponsive and Injectable Hydrogel for Tissue Agnostic Regeneration.** *Advamced Healthcare Materials* **2022**, *11* (23).
- Divakarla, S. K.; Das, T.; Chatterjee, C.; Ionescu, M.; Pastuovic, Z.; Jang, J.-H.; Al-Khoury, H.; Loppnow, H.; Yamaguchi, S.; Groth, T., Antimicrobial and Anti-inflammatory Gallium– Defensin Surface Coatings for Implantable Devices. ACS Applied Materials & Interfaces 2022, 14 (7), 9685-9696.
- 3. Limantoro, C.; Das, T.; He, M.; Dirin, D.; Manos, J.; Kovalenko, M. V.; Chrzanowski, W., Synthesis of Antimicrobial Gallium Nanoparticles Using the Hot Injection Method. ACS Materials Au 2023.
- 4. Phan, T. H.; Shi, H.; Denes, C. E.; Cole, A. J.; Wang, Y.; Cheng, Y. Y.; Hesselson, D.; Roelofs, S. H.; Neely, G. G.; Jang, J.-H., Advanced pathophysiology mimicking lung models for accelerated drug discovery. *Biomaterials Research* **2023**, *27* (1), 35.
- 5. Tsai, K. H.; Shi, H.; Parungao, R. J.; Naficy, S.; Ding, X.; Ding, X.; Hew, J. J.; Wang, X.; Chrzanowski, W.; Lavery, G. G., Skin 11β-hydroxysteroid dehydrogenase type 1 enzyme expression regulates burn wound healing and can be targeted to modify scar characteristics.

Burns & Trauma **2023**, *11*.Pokrajac L, Abbas A, Chrzanowski W, Dias G, Eggleton B, Maguire S, Maine E, Malloy TF, Nathwani J, Nazar L: **Nanotechnology for a Sustainable Future: Addressing Global Challenges with the International Network4Sustainable Nanotechnology**. *UCLA School of Law, Public Law Research Paper* 2022(22-03):15.

- 6. Phan TH, Kim SY, Rudge C, Chrzanowski W: **Made by cells for cells-extracellular vesicles as next-generation mainstream medicines**. *Journal of Cell Science* 2022, **135**(1):jcs259166.
- 7. Divakarla SK, Das T, Chatterjee C, Ionescu M, Pastuovic Z, Jang J-H, Al-Khoury H, Loppnow H, Yamaguchi S, Groth T: Antimicrobial and Anti-inflammatory Gallium–Defensin Surface Coatings for Implantable Devices. ACS Applied Materials & Interfaces 2022, 14(7):9685-9696.
- 8. Cheng Z, Li Y, Wang K, Zhu X, Tharkar P, Shu W, Zhang T, Zeng S, Zhu L, Murray M: **Compritol solid lipid nanoparticle formulations enhance the protective effect of betulinic acid derivatives in human Müller cells against oxidative injury**. *Experimental Eye Research* 2022, **215**:108906.
- 9. Wang H, Zhang C, Yu J, Song Y, Liu S, Chrzanowski W, Cai W: **Voxel-wise cross-volume representation learning for 3d neuron reconstruction**. In: *International Workshop on Machine Learning in Medical Imaging: 2021*: Springer; 2021: 248-257.
- Tong A, Sorrell TC, Black AJ, Caillaud C, Chrzanowski W, Li E, Martinez-Martin D, McEwan A, Wang R, Motion A: Research priorities for COVID-19 sensor technology. *Nature Biotechnology* 2021, 39(2):144-147.
- 11. Pokrajac L, Abbas A, Chrzanowski W, Dias GM, Eggleton BJ, Maguire S, Maine E, Malloy T, Nathwani J, Nazar L: Nanotechnology for a sustainable future: Addressing global challenges with the international network4sustainable nanotechnology. In.: ACS Publications; 2021.
- Phan TH, Divakarla SK, Yeo JH, Lei Q, Tharkar P, Pansani TN, Leslie KG, Tong M, Coleman VA, Jämting Å: New Multiscale Characterization Methodology for Effective Determination of Isolation–Structure–Function Relationship of Extracellular Vesicles. Frontiers in bioengineering and biotechnology 2021, 9:669537.
- 13. Park SH, Phan TH, Kim JE, Chrzanowski W: **Physicochemical Characterisation of Extracellular Vesicles**. In: *Extracellular Vesicles*. edn.; 2021: 45-75.
- 14. Pansani TN, Phan TH, Lei Q, Kondyurin A, Kalionis B, Chrzanowski W: **Extracellular Vesicle-Based Coatings Enhance Bioactivity of Titanium Implants—SurfEV**. *Nanomaterials* 2021, **11**(6):1445.
- 15. Hunt NJ, Lockwood GP, Kang SW, Westwood LJ, Limantoro C, Chrzanowski W, McCourt PA, Kuncic Z, Le Couteur DG, Cogger VC: Quantum dot nanomedicine formulations dramatically improve pharmacological properties and alter uptake pathways of metformin and nicotinamide mononucleotide in aging mice. ACS nano 2021, **15**(3):4710-4727.
- 16. Chrzanowski W, Lim CT, Kim SY: Extracellular Vesicles: Applications to Regenerative Medicine, Therapeutics and Diagnostics, vol. 9: Royal Society of Chemistry; 2021.
- Chen Y, Song Y, Zhang C, Zhang F, O'Donnell L, Chrzanowski W, Cai W: CellTrack R-CNN: A novel end-to-end deep neural network for cell segmentation and tracking in microscopy images. In: 2021 IEEE 18th International Symposium on Biomedical Imaging (ISBI): 2021: IEEE; 2021: 779-782.
- 18. Zhang B, Zhu M, Li Z, Lung PS, Chrzanowski W, Kwok CT, Lu J, Li Q: **Cellular fate of deformable needle-shaped PLGA-PEG fibers**. *Acta Biomaterialia* 2020, **112**:182-189.
- 19. Reczyńska K, Marszałek M, Zarzycki A, Reczyński W, Kornaus K, Pamuła E, Chrzanowski W: Superparamagnetic iron oxide nanoparticles modified with silica layers as potential agents for lung cancer treatment. *Nanomaterials* 2020, **10**(6):1076.
- 20. Reczyńska K, Marchwica P, Khanal D, Borowik T, Langner M, Pamuła E, Chrzanowski W: **Stimuli-sensitive fatty acid-based microparticles for the treatment of lung cancer**. *Materials Science and Engineering:* C 2020, **111**:110801.
- 21. Reczyńska K, Khanal D, Pielichowska K, Pamuła E, Chrzanowski W: Distinct Influence of Saturated Fatty Acids on Malignant and Nonmalignant Human Lung Epithelial Cells. *Lipids* 2020, **55**(2):117-126.
- 22. Liu Y, Zhang W, Phan TH, Chrzanowski W, Rodger A, Wang Y: **Positively charged gold-silver nanostar enabled molecular characterization of cancer associated extracellular vesicles**. *Analytical Methods* 2020, **12**(48):5908-5915.

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- 24. Khanal D, Lei Q, Pinget G, Cheong DA, Gautam A, Yusoff R, Su B, Yamaguchi S, Kondyurin A, Knowles JC: The protein corona determines the cytotoxicity of nanodiamonds: implications of corona formation and its remodelling on nanodiamond applications in biomedical imaging and drug delivery. *Nanoscale Advances* 2020, **2**(10):4798-4812.
- 25. Khanal D, Khatib I, Ruan J, Cipolla D, Dayton F, Blanchard JD, Chan H-K, Chrzanowski W: Nanoscale probing of liposome encapsulating drug nanocrystal using atomic force microscopy-infrared spectroscopy. *Analytical chemistry* 2020, **92**(14):9922-9931.
- 26. Dodballapur V, Song Y, Huang H, Chen M, Chrzanowski W, Cai W: **Dual-Stage Domain Adaptive Mitosis Detection for Histopathology Images**. In: *2020 Digital Image Computing: Techniques and Applications (DICTA): 2020*: IEEE; 2020: 1-7.
- 27. Dałek P, Borowik T, Reczyńska K, Pamuła Eb, Chrzanowski W, Langner M: **Evaluation of the in** vitro stability of stimuli-sensitive fatty acid-based microparticles for the treatment of lung cancer. *Langmuir* 2020, **36**(37):11138-11146.
- 28. Chrzanowski W, Kim SY, McClements L: Can stem cells beat COVID-19: advancing stem cells and extracellular vesicles toward mainstream medicine for lung injuries associated with SARS-CoV-2 infections. Frontiers in Bioengineering and Biotechnology 2020:554.
- 29. Chiari MD, Rodrigues MC, Pinto MF, Vieira DN, Vichi FM, Vega O, Chrzanowski W, Nagaoka N, Braga RR: Development of brushite particles synthesized in the presence of acidic monomers for dental applications. *Materials Science and Engineering: C* 2020, **116**:111178.
- 30. Tharkar P, Varanasi R, Wong WSF, Jin CT, Chrzanowski W: **Nano-enhanced drug delivery and therapeutic ultrasound for cancer treatment and beyond**. *Frontiers in Bioengineering and Biotechnology* 2019, **7**:324.
- 31. Pinget G, Tan J, Janac B, Kaakoush NO, Angelatos AS, O'Sullivan J, Koay YC, Sierro F, Davis J, Divakarla SK: Corrigendum: Impact of the Food Additive Titanium Dioxide (E171) on Gut Microbiota-Host Interaction. *Frontiers in Nutrition* 2019, **6**:100.
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- 38. Kim SY, Chrzanowski W: **Stem Cell Delivery Systems and Devices-Spraying**. In: *Stem Cell-Based Therapy for Lung Disease*. edn.: Springer, Cham; 2019: 241-253.
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REACH AND SIGNIFICANCE OF THE RESEARCH

My research program on nanomedicine has led to knowledge and health impact on the safety and efficacy of nanoparticles for medical applications. This has driven my development of (i) innovative high-content and high-throughput nanoparticle assessment methods & tools, and (ii) advanced therapeutic formulations, resulting in clinical & commercial outcomes.

My research program has established human pathophysiology-mimicking models and innovative methodologies for the multiscale determination of nanoparticles' interactions with biological systems, thus their safety and efficacy [e.g. Biomaterials Research 2023; Frontiers in Bioengineering and Biotechnology 2021; International Journal of Nanotechnology 2017].

The use of my models led to the 2016 Vice Chancellor's Award for Reduction in Use of Animals, and the 2022 Annual Bioprocessing Network Conference award for the development of non-animal strategy for the quality control of biologics. I also established an international consortium (iCare) and led the submission of a successful EU H2020 grant (\$5M; CI: iCare: Integrated Assessment and Advanced Characterization of Neuro-Nanotoxicity; HE-01-35, 2022). My work on alternative models to animals led to establishing NSW Centre for Organoid Innovation (CI). My work on human-mimicking models provided also key evidence presented during in NSW Inquiry into Alternatives to Animal Research

(2022), and was pivotal to the USyd participation in developing a strategy and roadmap for Australia: 'Non-animal models for preclinical development' (led by CSIRO) of which I am the USyd lead.

The assessment methods & tools I established were/are essential to establish innovative, safe and affordable nano-based formulations (aka nanomedicines) for different indications, *e.g.* gene therapy for macular degeneration (CI: Macular Disease Foundation Australia Research Grant G212978; AI: NHMRC Ideas 2020950); nano-Q10 formulation to treat metabolic disease (AI: NHMRC Ideas 2013621), and nano-cannabinoid formulation delivery using a microneedle patch (CI: Commercial Development and Industry Partnership Fund & NSW BOOST Program 2018).

In contrast to conventional approaches to nanomedicines design that sparingly consider the social, legal and sustainability aspects of new technologies, my approach comprehends the interconnected nanoparticle-cell-organ-patient-society relationship holistically (ACS Nano 2021; Nature Biotechnology 2021) and has had the following knowledge and health impacts:

KNOWLEDGE IMPACT

- 156 (218 GS) publications, 7360 citations, FWCI of 3.34 (5.29, last 3 years) in Biomedical and Clinical Sciences
- 21 prizes/awards since 2011 (6 international, 15 national); e.g. Highly Commended Finalist Research Australia Awards, category Frontier
- \$14.8M from grants since 2010, including 3x ARC Linkage grants, 3x EU (FP7/H2020)
- Reduction in use of experimental animals 3R Vice-Chancellor Award 2017, 3x research grants (\$80k) from Medical Advances Without Animals and Ethical Australia
- Protocols for determining the pharmacokinetics of nanoparticles at the single-cell level
- Current industry collaborations: BiomeCentric, Probiotics Australia, nanoMT, MedLab Clinical, ExoPharm, BCAL Diagnostics, Therakii, IZON, SynThera, CYTIVA.
- Adviser to WHO in preparation of a 2020 handbook on nanoparticle safety testing.

HEALTH IMPACT

- Development of the human lung models to reduce animal in research and boost success in clinical trials https://www.theguardian.com/science/2023/apr/11/australian-scientists-grow-replica-human-lungs-and-call-for-end-to-animal-testing
- Demonstration that TiO2 nanoparticles in food are harmful, leading to media with >300M views and contributing to EU banning these nanoparticles in food, <u>http://www.abc.net.au/news/2017-12-12/associate-professor-wojciech-chrzanowski-nanoparticles-food/9249038</u>,

https://www.sbs.com.au/news/health-concerns-over-nanomaterials-spark-call-for-safety-body

- Demonstration that NanaBis[™] from MedLab Clinical (https://www.medlab.co/science/scientific_advisory_board), a novel medication for chronic pain, is unstable. This led directly to changes in the manufacturing, dosing, delivery device, and formulation of NanaBis[™]; Phase II NCT04808531 in 360 participants.
- Demonstration that nano360, an anti-viral agent, is ineffective against viruses and has likely harmful biological effects, contributing to the company (nanoMT) withdrawing their application to the TGA.

KNOWLEDGE IMPACT

It is estimated that globally more than 10000 groups work on nanoparticles, my program is the only one which integrated nanoparticle-cell-organ-patient-society relationship holistically and translated from the laboratory through to routine applications. My program established:

A. Methods for determining: (i) the composition of single nanoparticles (engineered and natural such as EVs), and (ii) pharmacokinetics of nanoparticles at single cell level. The former outcome – previously considered impossible – was achieved by innovative combination of nanoscale infrared spectroscopy,

atomic force microscopy and nanoindentation (resolution <10 nm). The later was achieved by correlative nanoimaging using holotomography and confocal microscopy (resolution <100 nm)

B. Multi-organ-like systems that mimic human pathophysiology. These consist of 'mini organs' cultured using patent-derived cells and established using PhysioMimics[™] system (CN Bio), which emulates dynamic conditions of human body. I was the first to integrate lung mini organs with the PhysioMimics[™] system. This system is currently used by FDA and AstraZeneca.

C. Integration of legal framework and safe-sustainable-by-design as well as responsible innovation principles in the research program [ACS Nano, Cell science]

This integrated system (A+B+C) enables my research program to guide a design of advanced drug delivery system (e.g. NanoBis), nano-diagnostics (breast cancer screening test, research collaboration with BCAL) and new biologicals for lung injury (research contract Therakii Biotherapeutics LTD).

HEALTH IMPACT

For the ban of synthesized/engineered nanoparticles from food in EU, I claim partial responsibility (https://www.abc.net.au/radio/programs/am/scientists-call-for-national-body-to-regulate-

nanoparticles/9248522). The >300M media hits and over 20 media interviews about my research program's paper showing harmful effects of a common food nanoadditive (TiO2), on gut microbiota and overall health happened 18 months prior to the ban in EU. Leading to this ban I was an invited speaker and invited panel chair at Food Toxicology Conference in Singapore (January 2021), where I debated and discussed with representatives from the FDA, EU and Singapore the health impact and sustainability of nanoparticles.

Without the outcomes from my research program, the withdrawal of nano360 from a TGA application for its widespread use as an anti-viral spray for domestic and public applications would likely not have been made. My research program conducted the multifaceted testing of efficacy and safety and showed that this formulation is unsafe for humans and environment.

My research program's approach to characterising the interaction of single nanoparticles with single biomolecule, cells, organ-like structures and multi-organ-like systems was uniquely responsible for identifying the stable and safe formulation of NanaBis[™] by MedLab Clinical being used in the above-listed Phase II RCT.