



# *Varnost nanomaterialov: povezava med njihovimi lastnostmi in učinki na okolje*

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Biotehniška fakulteta, Oddelek za biologijo.



Biotechnical Faculty — University of *Ljubljana*

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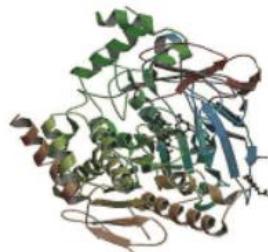
*We are looking for insights into bio-nano interactions at various levels*

The Bionanoteam strives to **holistically assess specific interactions of biological systems with nanoparticles on different levels of biological organization**. Our main topics of research are:

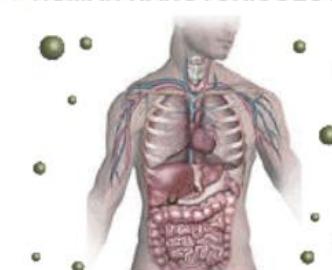
#### ENVIRONMENTAL NANOTOXICOLOGY



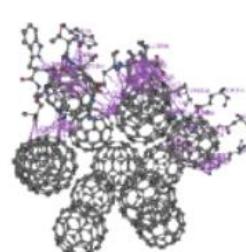
#### ACETYLCHOLINESTERASE RESEARCH



#### HUMAN NANOTOXICOLOGY



#### BIOSENSORS & NANODEVICES



*Principal Investigator*



**Prof. Dr. Damjana Drobne**

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[email](#) | [bibliography](#)

Bionanoteam is a research group at the Biotechnical Faculty, University of Ljubljana.

Founded in 2008 by Prof. Damjana Drobne, Bionanoteam currently consists of 4 experienced researchers, 4 PhD candidates, and a number of graduate and undergraduate students. Damjana is a Professor of Zoology and Professor of Toxicology from University of Ljubljana. With over twenty years of research experience, she is the founder and principal investigator of Bionanoteam.

# Kje smo?



UNILJ



Biotehniška fakulteta



Oddelek za biologijo

# SKUPINA ZA NANOBILOGIJO IN NANOTOKSIKOLOGIJO

## študije *in vivo* in *in vitro* delovanja nanomaterialov na biološke sisteme



### KOPENSKI IZOPODNI RAKI

- spremljanje dinamike prehranjevanja, preživetje
- študije bioakumualcije in biodistribucije nanomaterialov
- študije na izoliranem prebavnem organu



### VODNE BOLHE

- akutni testi toksičnosti
- kopičenje NM na površini živali



### MIGETALKARJI

- spremljanje viabilnosti modelnega organizma



### ČEBELE

- spremljanje prehranskih parametrov in preživetja
- študije nevrotoksičnosti NM



### MORSKI JEŽKI

- vpliv NM na jajčne celice, spermije in spremembe zarodka
- študije okoljskega plašča NM



### SOLINSKI RAKI

- akutni testi toksičnosti



### CELIČNE KULTURE

- spremembe integritete celične membrane, aktivnosti, proliferacijo celic, celično smrt, genotoksičnost...
- fosfolipidoza



- ERITROCITI

- spremenjena morfologija
- adsorbcija NM na površino eritrocitov
- mikrovezikulacija



### ENCIMI

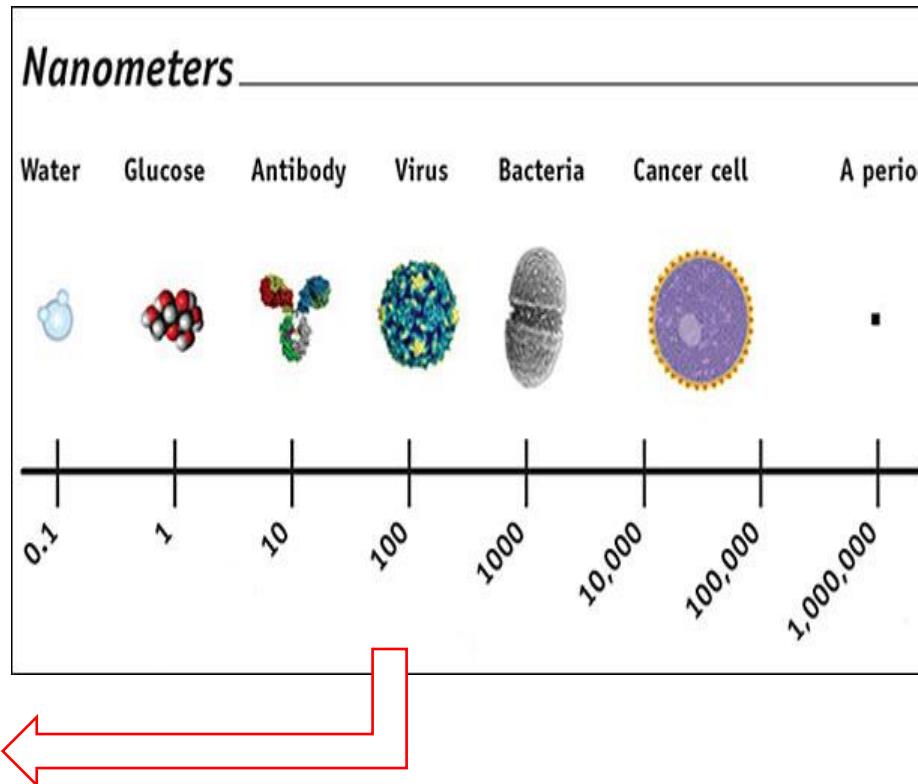
- inhibicija delovanja encima zaradi prisotnosti NM
- adsorpcija encima na NM

# Bionanoteam v EU FP7 in H2020 projektih:



- **EU FP7 NanoValid** (2011-2015) RIA
  - **EU FP7 NanoMILE** (2013-2017) RIA
  - **H2020 NanoFASE** (2015-2019) RIA
  - **H2020 Pandora** (2015-2018) MSCA-ITN-2015; Marie-Sklodowska-Curie Actions
- and a partner in
- **NanoSafety cluster**; hazard group
- a partner in two national Centers of Excellence(CO):
- **CO NAMASTE**
  - **CO Nanocenter**

# Nanomateriali



Nanomateriali

**NANO**  
Latinsko: nanus  
Grško: nanos



# Definicija nanomaterialov



## Definition of a nanomaterial

The EU adopted a definition of a nanomaterial in 2011 ([Recommendation on the definition of a nanomaterial \(2011/696/EU\)](#)). Its provisions include a requirement for review "*in the light of experience and of scientific and technological developments. The review should particularly focus on whether the number size distribution threshold of 50 %*

*A natural, incidental or manufactured material containing particles, in an unbound state or as an aggregate or as an agglomerate and where, for **50 % or more of the particles** in the number size distribution, one or more external dimensions is in the **size range 1 nm - 100 nm**.*

*In specific cases and where warranted by concerns for the environment, health, safety or competitiveness the number size distribution threshold of 50 % may be replaced by a threshold between 1 and 50 %.*

*By derogation from the above, fullerenes, graphene flakes and single wall carbon nanotubes with one or more external dimensions below 1 nm should be considered as nanomaterials.*

[http://ec.europa.eu/environment/chemicals/nanotech/faq/definition\\_en.htm](http://ec.europa.eu/environment/chemicals/nanotech/faq/definition_en.htm)

# *Naravni izvor*



# Inženirsko proizvedeni



©ADAM.

## Consumer Products Inventory



The Project on Emerging Nanotechnologies

— [CPI HOME](#) / [PRODUCTS](#) —

# All Products

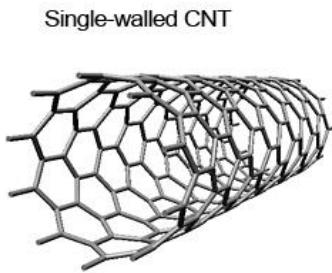
Products 1-25 of 1827

[STANJE 20-11-2016: 1827 NANO PRODUKTOV](#)

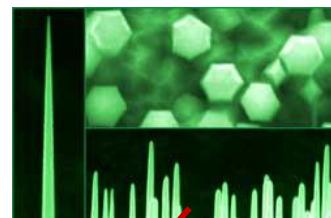
<http://www.nanotechproject.org/cpi/browse/nanomaterials/titanium-dioxide/>

# *Primeri nanomaterialov*

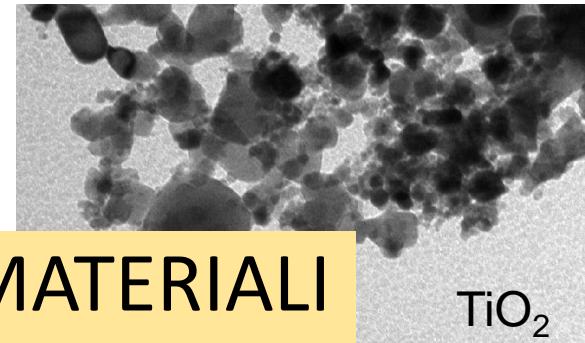
*nanoCEVKE*



*nano IGLICE*

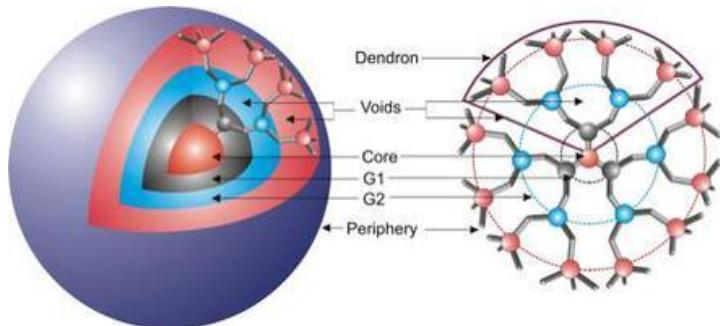


*nanoDELCI*

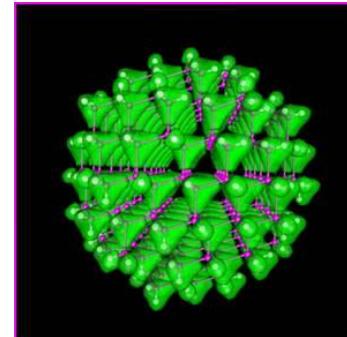


NANODELCI  = NANOMATERIALI

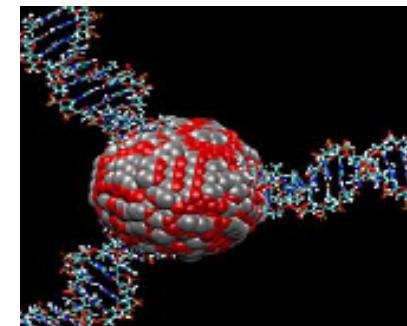
SFERIČNI DENDRIMERI



KVANTNE PIKE  
Quantum dots



*nanoBIOKOMPOZITI*  
TiO<sub>2</sub>+ DNA



# LASTNOSTI

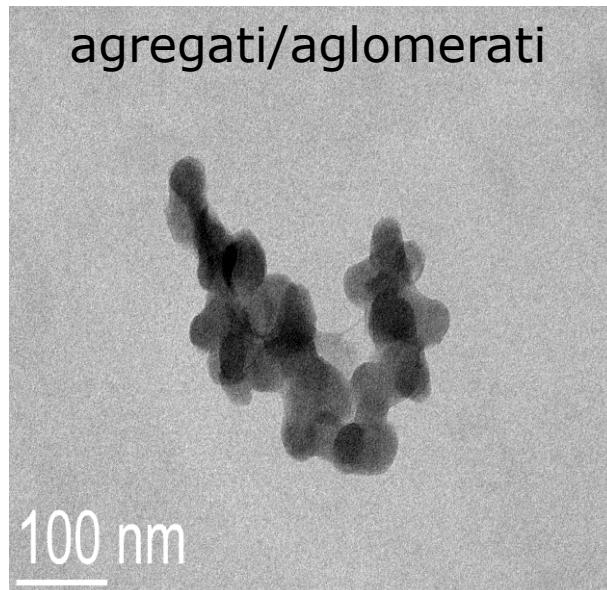
as they are

- ✓ Kemijska sestava
- ✓ Površinska kemija
- ✓ Kristalna struktura
- ✓ oblika
- ✓ velikost



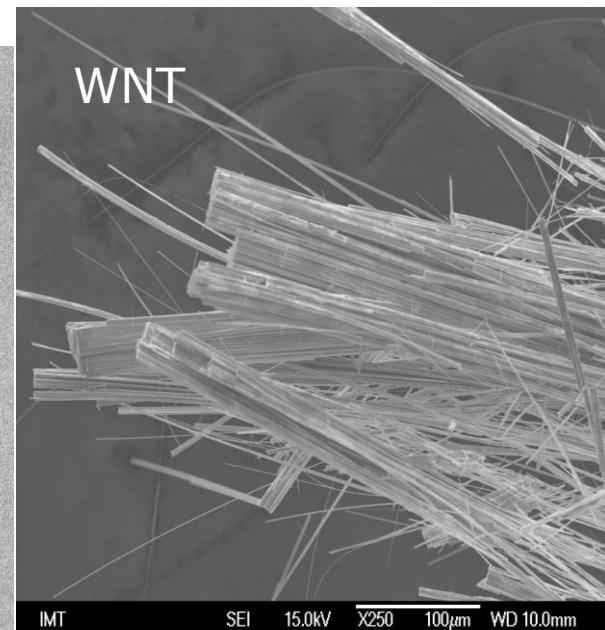
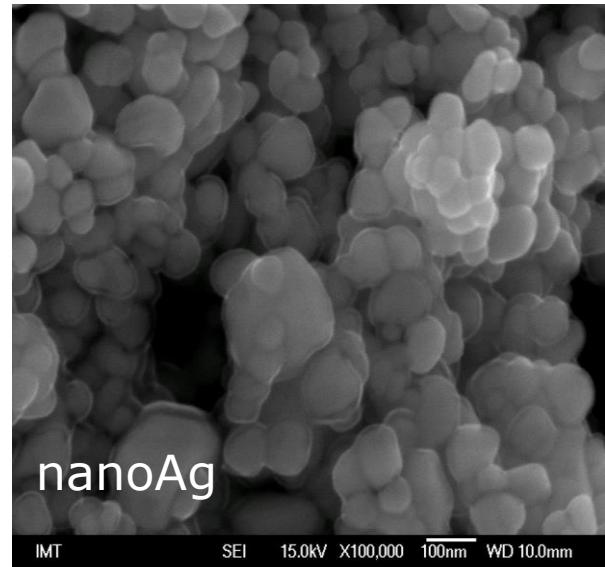
as they behave

- ✓ corona
- ✓ raztpljanje
- ✓ agregacija
- ✓ staranje



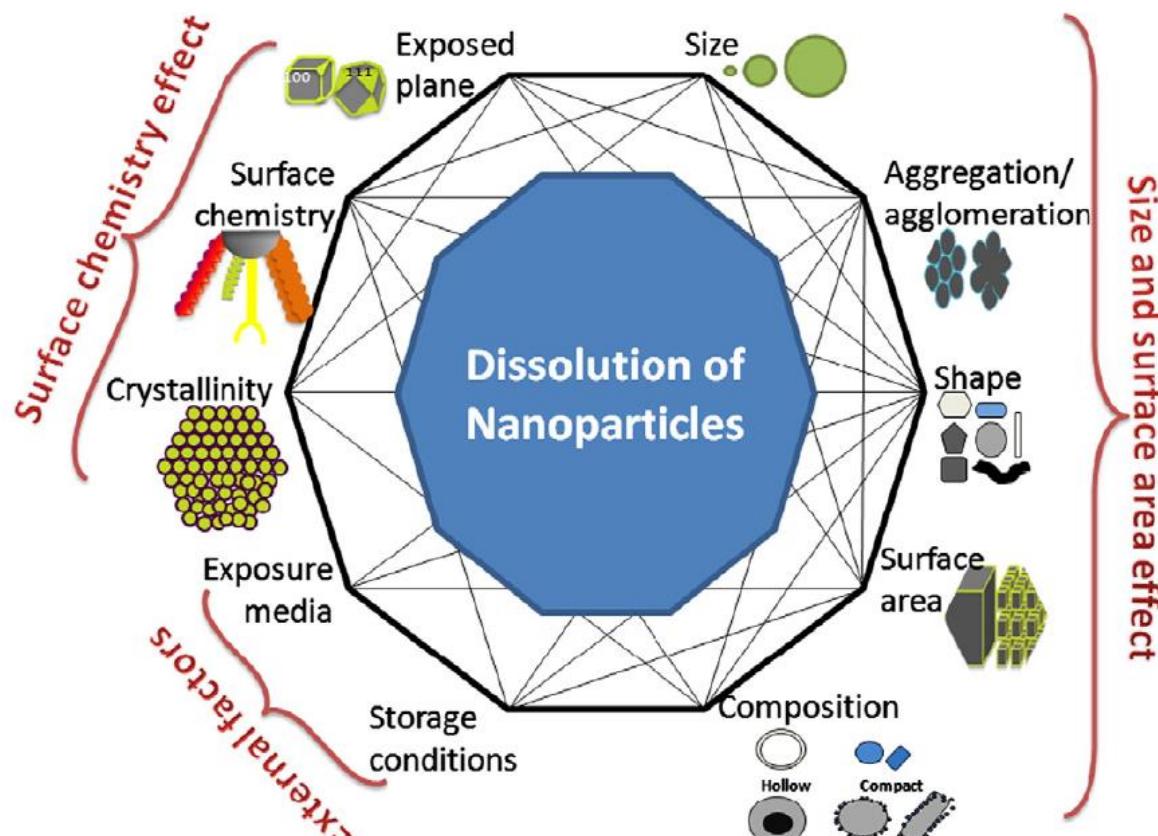
as they interact  
interract

- ✓ hidrofobnost
- ✓ adsorbcija



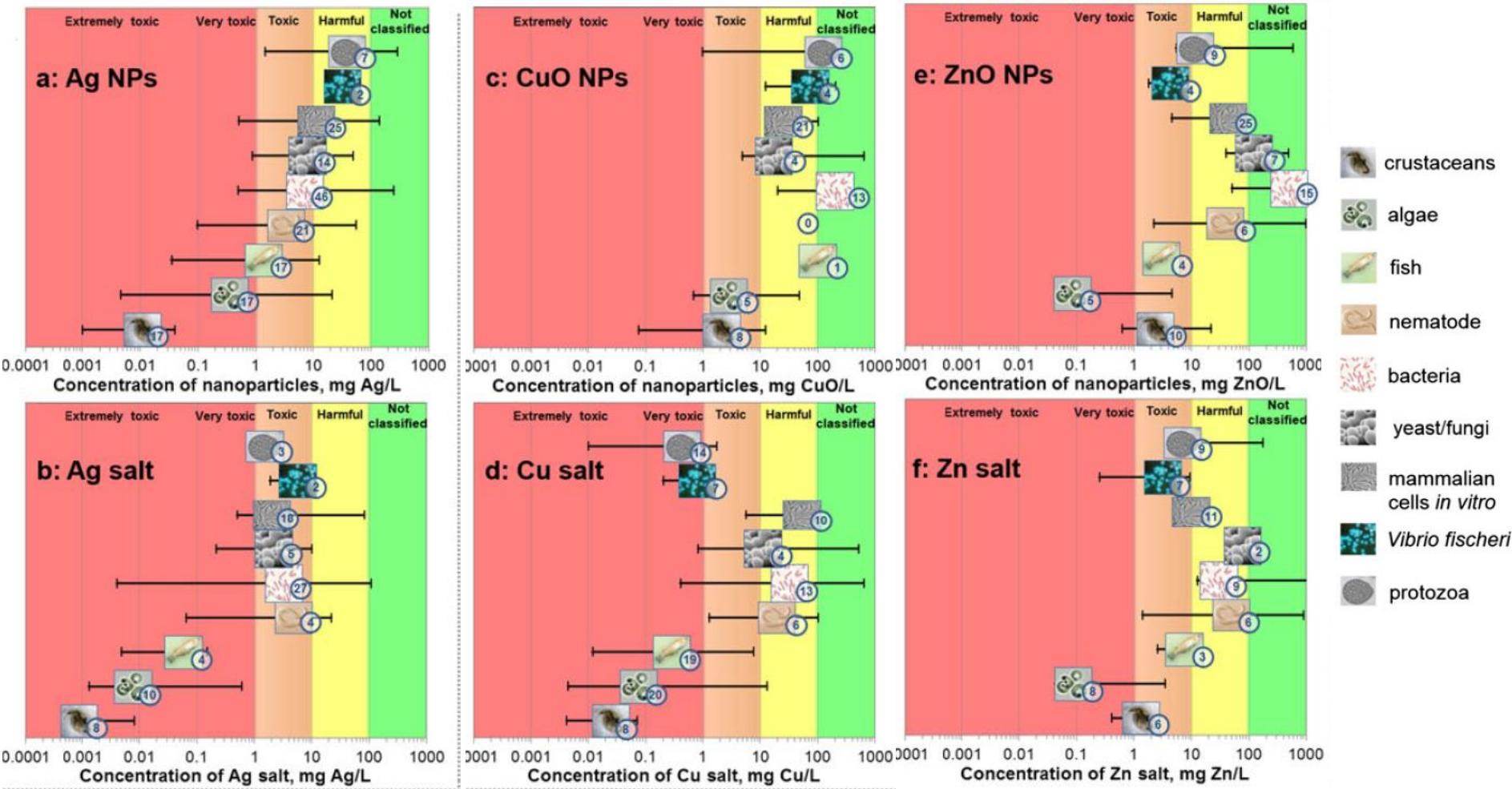
# Od LASTNOSTI do UČINKOV – znanje v 2017

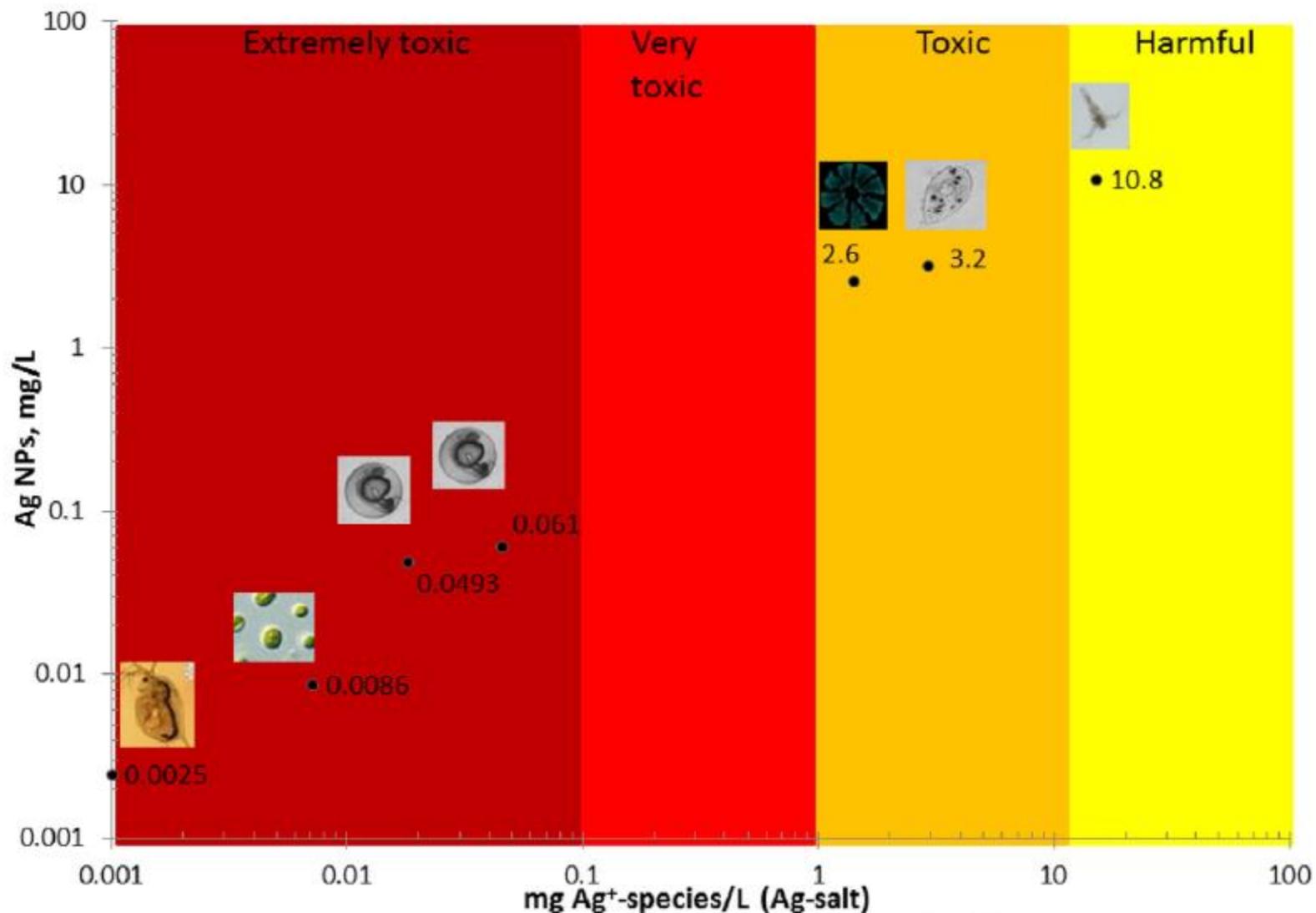
## KEMIJSKA SESTAVA: RAZTAPLJANJE



S.K. Misra et al. / Science of the Total Environment 438 (2012) 225–232

# TOKSIČNOST NEKATERIH MATERIALOV JE ODVISNA OD RAZTAPLJANJA

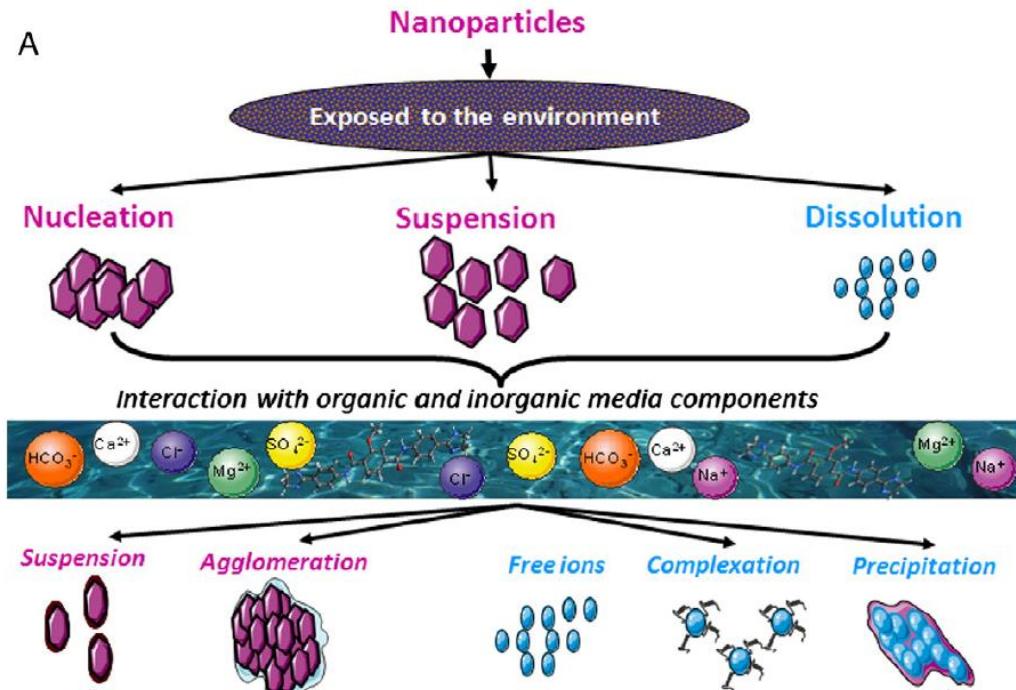




# NANOMATERIALI IMAJO INTERAKCIJO Z MEDIJEM

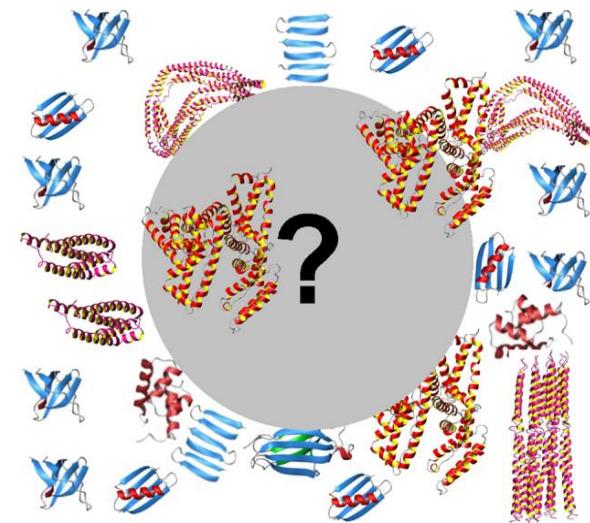
## DISSOLUTION/AGGREGATION/INTERACTION

A



S.K. Misra et al. / Science of the Total Environment 438 (2012) 225–232

## PROTEIN-NANOPARTICLE COMPLEX „PROTEIN CORONA“



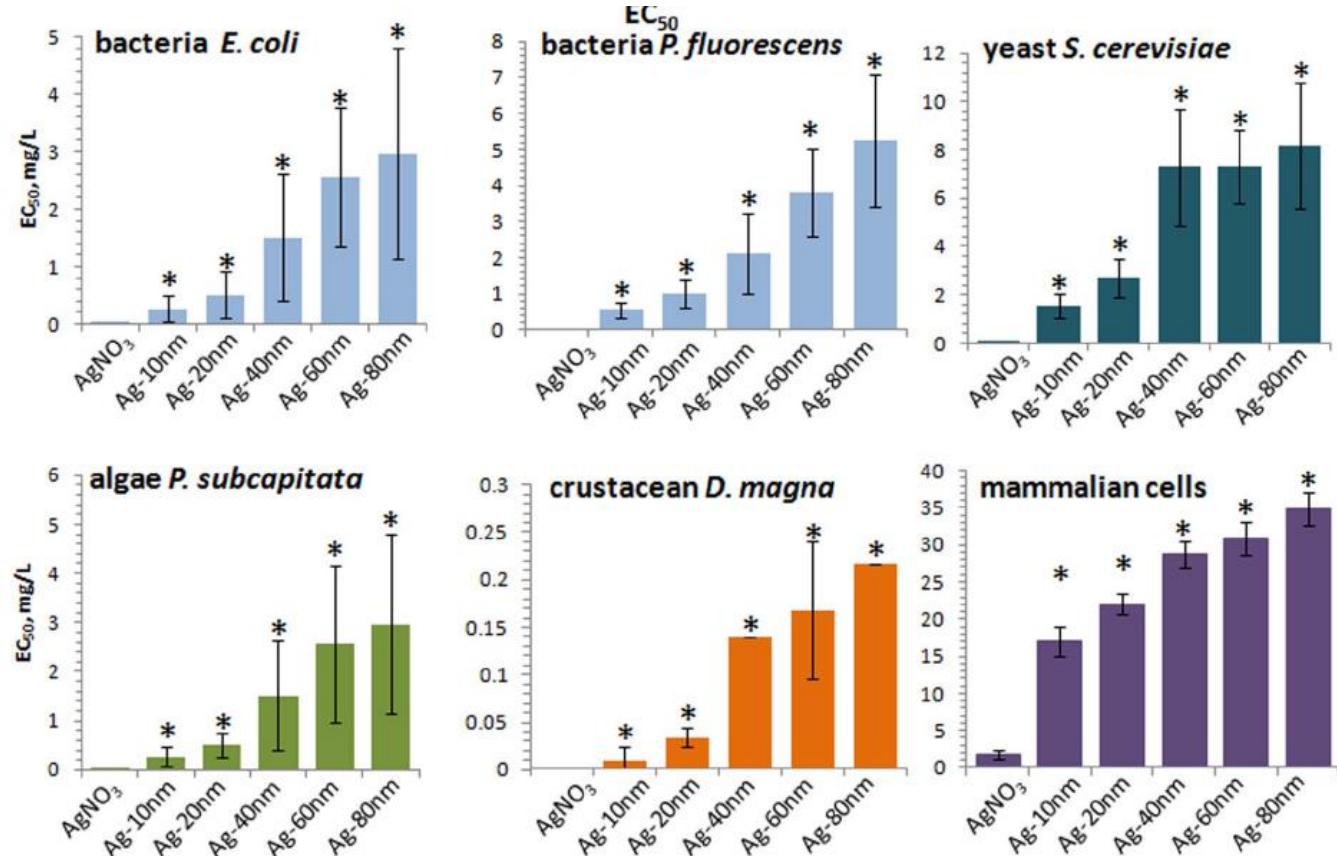
I. Lynch et al. / Advances in Colloid and Interface Science 134–135 (2007) 167–174

# Je UČINEK ODVISEN OD VELIKOSTI?

## Size-Dependent Toxicity of Silver Nanoparticles to Bacteria, Yeast, Algae, Crustaceans and Mammalian Cells *In Vitro*

Angela Ivask<sup>1\*</sup>, Imbi Kurvet<sup>1</sup>, Kaja Kasemets<sup>1</sup>, Irina Blinova<sup>1</sup>, Villem Aruoja<sup>1</sup>, Sandra Suppi<sup>1</sup>, Heiki Viia<sup>1</sup>,

PLoS ONE 9(7): e102108.



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PLoS ONE 9(7): e102108.

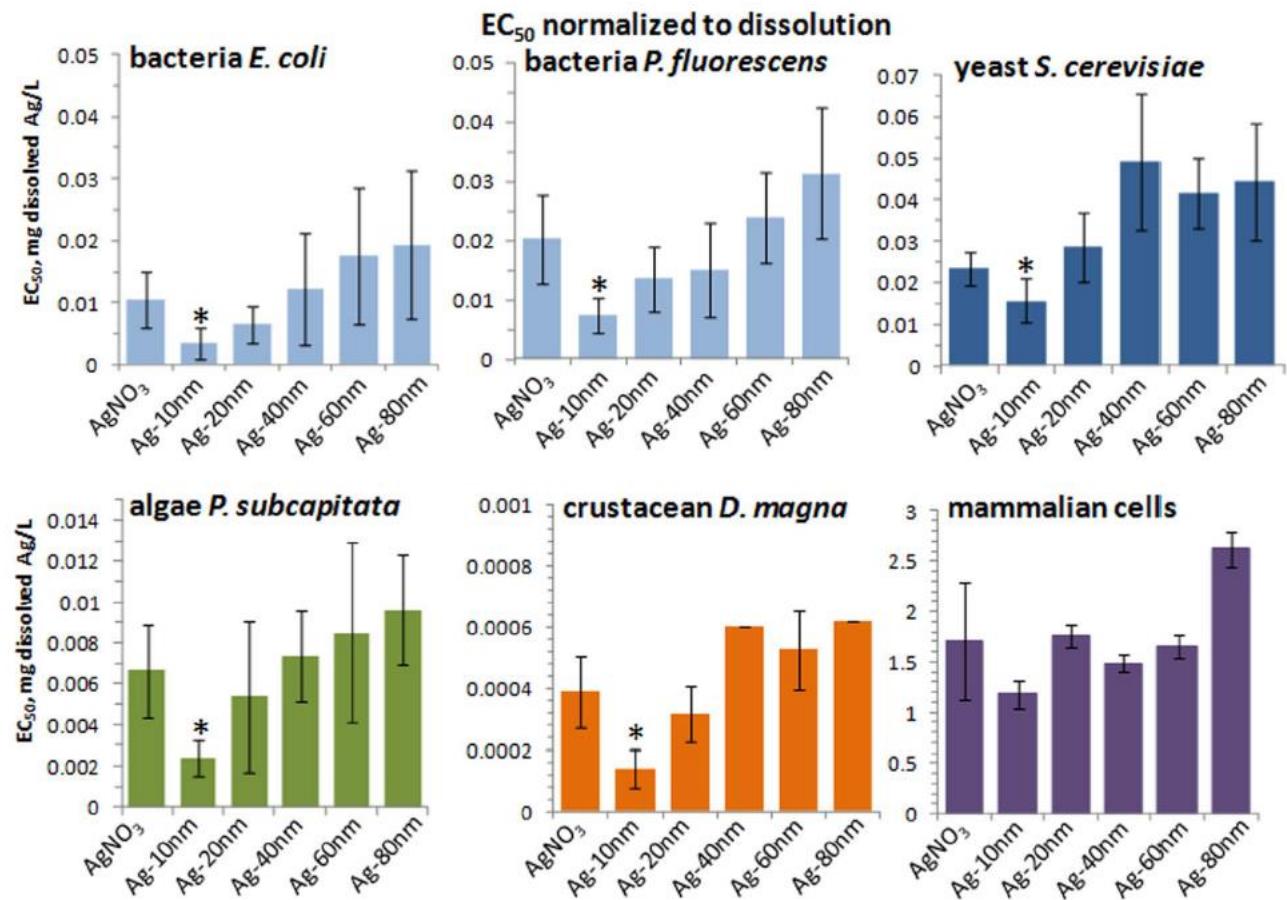
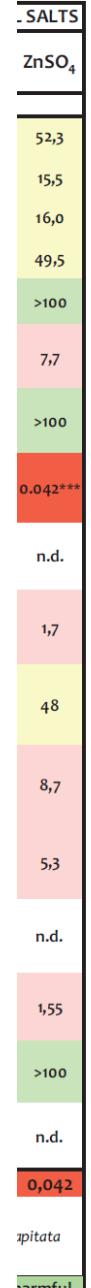


Table 4. Tox

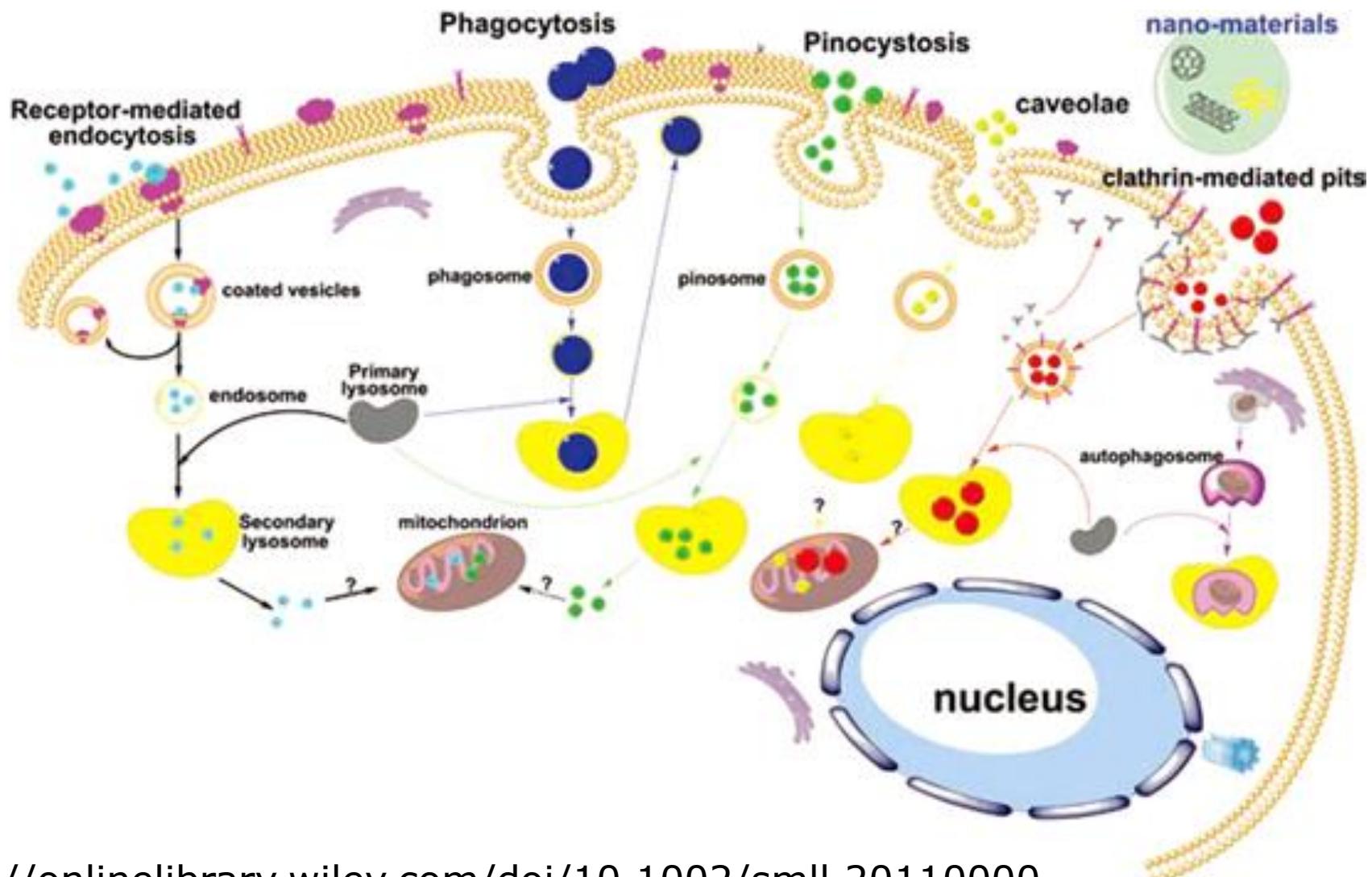
## GRUPIRANJE NANOMATERIALOV GLEDE NA LASTNOSTI

	Tests models	Time, medium, endpoint	NANOMATERIALS						SOLUBLE METAL SALTS			
			SiO <sub>2</sub>	TiO <sub>2</sub>	Au	MWCNT	Ag	CuO	ZnO	AgNO <sub>3</sub>	CuSO <sub>4</sub>	ZnSO <sub>4</sub>
Prokaryotes, single cell	<b>BACTERIA:</b>		EC50, MBC or LOEC (mg metal/l or mg MWCNTs/l)									
	<i>Escherichia coli</i>	4 h EC50, LB, growth inhibition	>100	>100	>10	n.d.	3,1	>100	67	1,3	>100	52,3
	<i>Staphylococcus aureus</i>	4 h EC50, LB, growth inhibition	>100	>100	>10	n.d.	5,2	>100	16	2,2	>100	15,5
	<i>Bacillus subtilis</i>	4 h EC50, LB, growth inhibition	>100	>100	>10	n.d.	4,5	>100	14	3,0	>100	16,0
	<i>Pseudomonas putida</i>	4 h EC50, LB, growth inhibition	>100	>100	>10	n.d.	3,8	>100	69	2,0	>100	49,5
	<i>Pseudomonas aeruginosa</i>	4 h EC50, LB, growth inhibition	>100	>100	>10	n.d.	3,2	>100	>100	2,1	>100	>100
	<i>Vibrio fischeri</i>	0,5 h EC50, 2% NaCl, bioluminescence inhibition	>100	>100	4,8*	n.d.	2,9	3,4	8,3	1,4	0,3	7,7
	<b>YEAST:</b>		24 h MBC, deionized water, viability	n.d.	>100	>100	1	>100	20	1	4	>100
	<b>ALGA:</b>		72 h EC50, OECD201 medium, growth inhibition	83,6	6,8	n.d.	n.d.	0,0086**	0,7	0,14	0,0071**	0,011
	<b>PROTOZOAN:</b>		24 h EC50, deionized water, viability	>100	>100	>30	>100	3,9	>100	3,9	2,9	n.d.
Eukaryotes, single cell	human mesenchymal stem cells	24 h EC50, complete medium, membrane integrity (PI)	>100	>100	>30	100	6,4	2,2	1,9	n.d.	2,73	1,7
	human mesenchymal stem cells	24 h EC50, complete medium, mitochondrial activity (MTT)	>100	>100	>30	n.d.	4,6	>100	16,3	n.d.	>100	48
	murine fibroblasts	24 h, 50% complete medium, mitochondrial	>100	>100	>6	>100	3****	0,7	9,1	2	2,9	8,7
	BALB/c3T3											
	HeLa											
Human cell cultures in vitro	<b>Harmful</b>		AO/EB, mitotically active cells									
	<b>Biopata</b>		AO/EB, mitotically active cells									
	<b>harmful</b>		AO/EB, mitotically active cells									
	<b>harmful</b>		AO/EB, mitotically active cells									
	<b>harmful</b>		AO/EB, mitotically active cells									



. AO/EB,

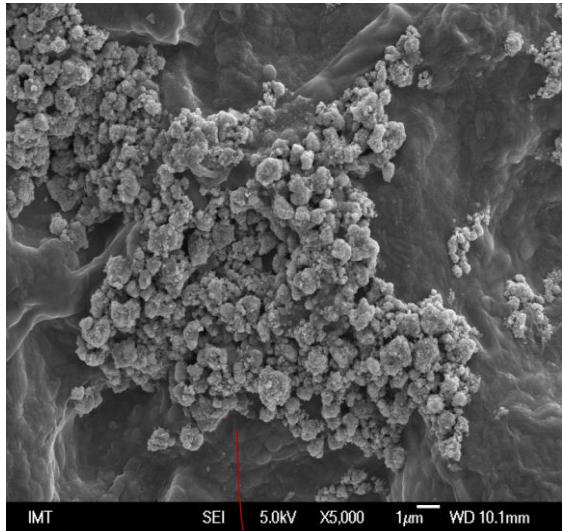
# VSTOP NANODELCEV V CELICO ?



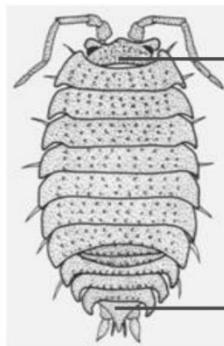
<http://onlinelibrary.wiley.com/doi/10.1002/smll.201100001/abstract>

NP na listih

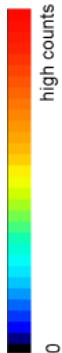
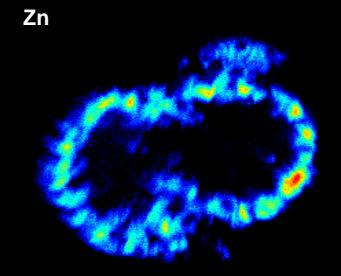
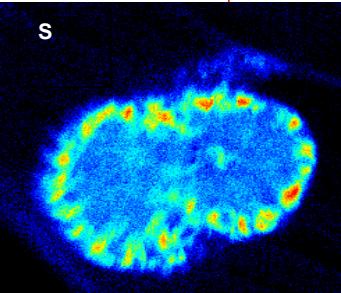
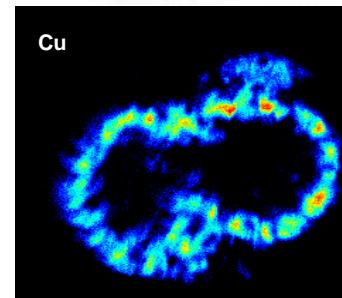
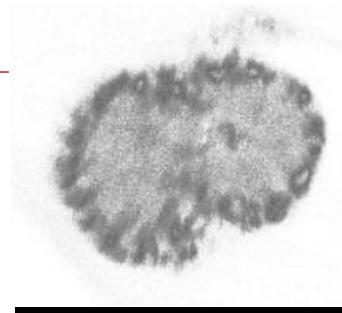
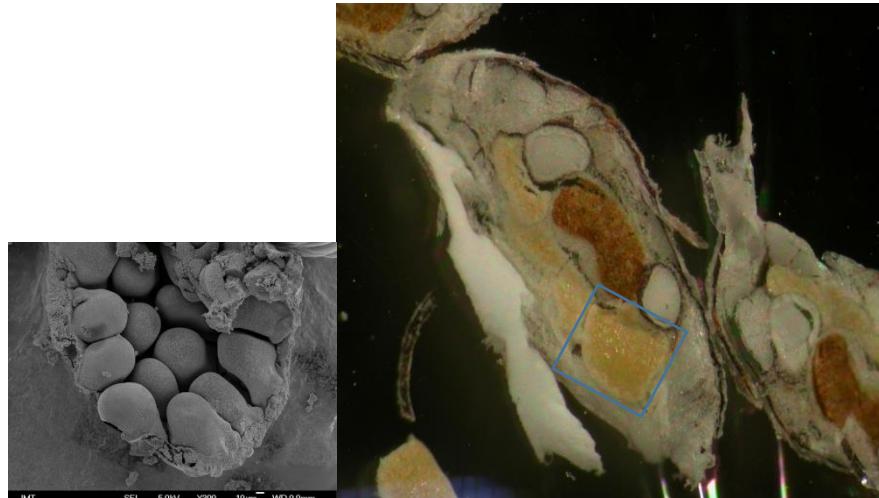
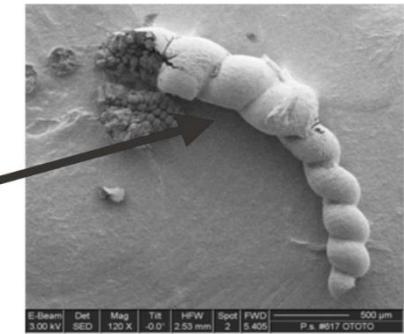
*Porcellio scaber*



+



*Porcellio scaber*



PIXE elementna analiza

# Dokumentirani učinki na organizme

## Stabilnost membrane celic prebavnih žlez – fluorescentna mikroskopija

Hranjenje modelnih živali z  
nanomateriali

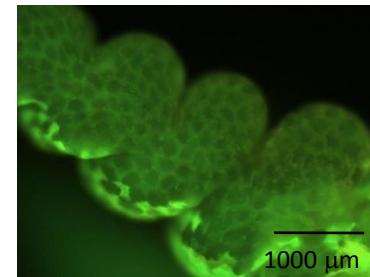


Izolacija prebavnih žlez  
Barvanje z mešanico  
barvil AO/EB

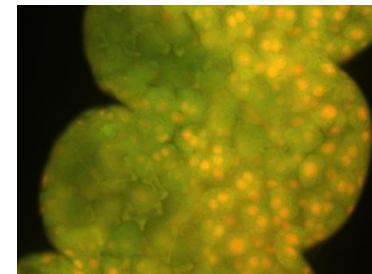
Fluorescentna mikroskopija



Mikrografije pobarvanih prebavnih žlez  
of *P. scaber* (Valant et al., 2009)

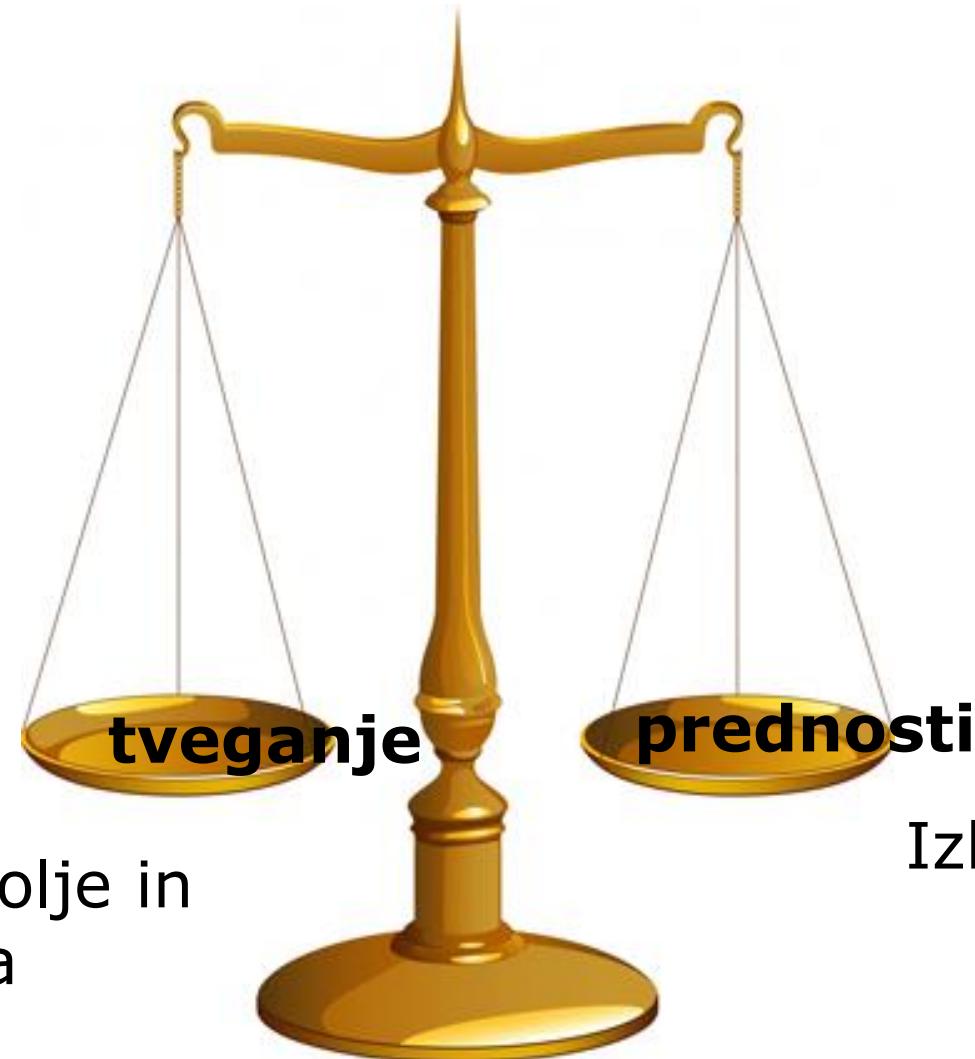


(A)  
Negativna  
kontrola,  
jedra niso  
obarvana



(B) Pozitivna  
kontrola,  
jedra svetijo  
oranžno (EB)

# *Kdaj so nanodelci varni, kdaj niso*



Učinki na okolje in  
človeka

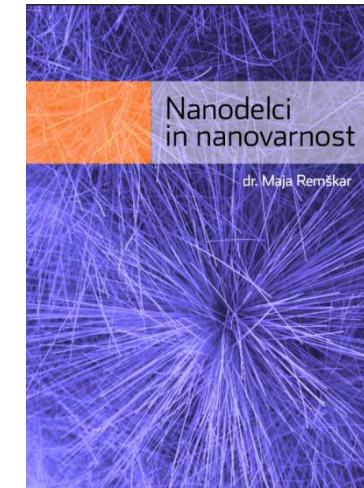
Izboljšave za  
naše  
življenje

# Priporočamo v branje

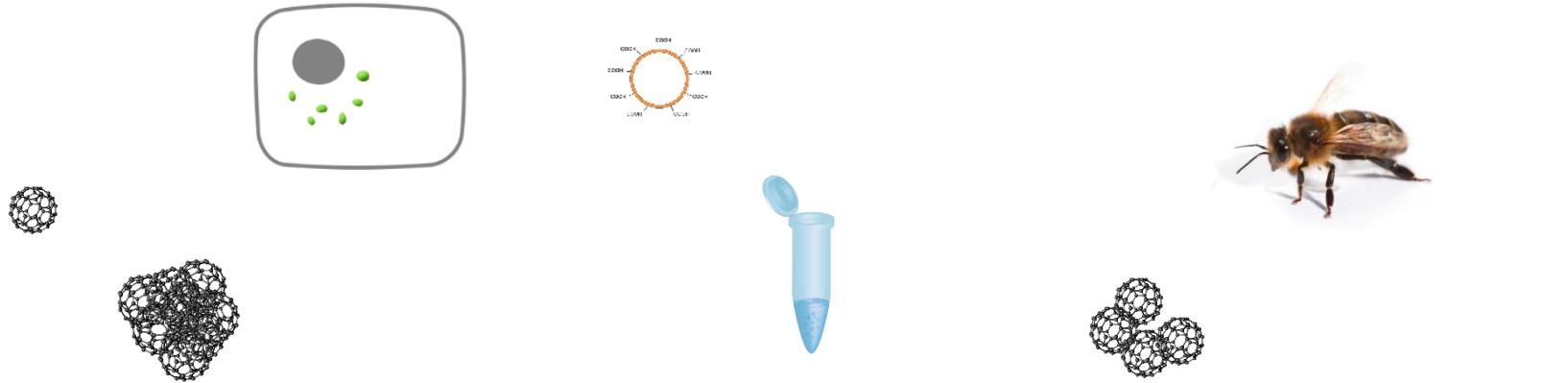


Acquisition, evaluation and public-oriented presentation of society-relevant data and findings relating to nanomaterials (DaNa)

<http://nanopartikel.info/en/projects/completed-projects/dana>



[http://www.kemijskovaren.si/files/nano\\_knjiga.pdf](http://www.kemijskovaren.si/files/nano_knjiga.pdf)



# Hvala za pozornost.

