

SUPPLEMENTARY INFORMATION

Toxicity of Ag, CuO and ZnO nanoparticles to selected environmentally relevant test organisms and mammalian cells in vitro: a critical review

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Supplementary Information contains the following data:

Table S1 Total number of publications in Thomson Reuters ISI Web of Science for different applications of Ag, ZnO and CuO nanoparticles. Search was performed combining the keywords "silver" or "CuO" or "ZnO" and "nano*" and "application category". The search was done on 22.03.2013

Table S2 The test species that were used the most often for determining the L(E)C50 and MIC values (Tables S3-S7) in the analysed literature

Table S3 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for silver nanoparticles for various organisms.

Table S4 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for silver salt for various organisms

Table S5 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for CuO nanoparticles for various organisms

Table S6 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for copper salt for various organisms

Table S7 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for ZnO nanoparticles for various organisms

Table S8 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for zinc salt for various organisms

Table S1 Total number of publications in Thomson Reuters ISI Web of Science for different applications of Ag, ZnO and CuO nanoparticles. Search was performed combining the keywords "silver" or "CuO" or "ZnO" and "nano*" and "application category". The search was done on 22.03.2013

Application area	Ag NPs	CuO NPs	ZnO NPs
Hydrogen storage	63	24	65
Environmental remediation	38	9	18
Catalysis	1012	138	406
Drug delivery	379	10	92
Medical imaging	51	1	8
Photovoltaic	103	1	176
Textiles	297	9	129
Therapeutics	55	0	4
Reinforced composites	50	23	48
Electronics	514	13	323
Optics	631	5	171
Coatings and pigments	10	2	23
Cosmetics	65	5	57
Ceramics applications	85	12	111
Anti-oxidants	5	0	0
Lubrication	36	5	12
Sensors	1915	341	2395
Absorbents	20	2	16
Energetics	48	3	37
Magnetics	53	3	30
Water purification	125	10	57
Air emissions reduction	10	4	12
Natural and green products	10	1	0
Quantum computing	17	2	8
Masonry and building materials	0	0	0
Photonics	183	2	105
Surfactants	447	45	217
Antimicrobials	1477	25	120

Table S2 The test species that were used the most often for determining the L(E)C50 and MIC values (Tables S3-S7) in the analysed literature

Organisms	Ag NPs	CuO NPs	ZnO NPs
Crustaceans	17 LC50 values <i>Daphnia magna</i> (13); <i>D. pulex</i> (2); <i>Ceriodaphnia dubia</i> (2)	8 LC50 values <i>Daphnia magna</i> (6); <i>D. pulex</i> (1); <i>Thamnocephalus platyurus</i> (1)	10 LC50 values <i>Daphnia magna</i> (10)
Algae	17 EC50 values <i>Pseudokirchneriella subcapitata</i> (8); <i>Chlamydomonas reinhardtii</i> (5); <i>Chlorella vulgaris</i> (1); <i>Chattonella marina</i> (1); <i>Dunaliella tertiolecta</i> (1); <i>Ochromonas danica</i> (1)	5 EC50 values <i>Pseudokirchneriella subcapitata</i> (8); <i>Chlamydomonas reinhardtii</i> (5); <i>Chlorella vulgaris</i> (1); <i>Chattonella marina</i> (1); <i>Dunaliella tertiolecta</i> (1); <i>Ochromonas danica</i> (1)	5 EC50 values <i>Pseudokirchneriella subcapitata</i> (3); <i>Thalassiosira pseudonana</i> (1); <i>Skeletonema costatum</i> (1)
Fish	17 LC50 values <i>Oryzias latipes</i> (8); <i>Pimephales promelas</i> (5); <i>Danio rerio</i> embryos (3); <i>Oreochromis mossambicus</i> (1)	1 LC50 values <i>Cyprinus carpio</i> (1)	4 LC50 values <i>Danio rerio</i> (adult)(2); <i>D. rerio</i> embryo (2)
Nematodes	21 (L(E)C50 values <i>Caenorhabditis elegans</i> (21)	0 (L(E)C50 values	6 (L(E)C50 values <i>Caenorhabditis elegans</i> (6)
Bacteria	46 MIC (minimal inhibitory concentration) values <i>Escherichia coli</i> (12); <i>Staphylococcus aureus</i> (12); <i>Bacillus subtilis</i> (4); <i>Pseudomonas aeruginosa</i> (4); <i>Shewanella oneidensis</i> (2); <i>Aggregatibacter actinomycetemcomitans</i> (1); <i>Enterococcus faecalis</i> (1); <i>E. faecium</i> (1); <i>Fusobacterium nucleatum</i> (1); <i>Klebsiella aerogenes</i> (1); <i>K. pneumoniae</i> (1); <i>Salmonella abony</i> (1); <i>S. typhimurium</i> (1); <i>Porphyromonas gingivalis</i> (1); <i>Prevotella intermedia</i> (1); <i>Proteus vulgaris</i> (1); <i>Streptococcus mutans</i> (1)	13 MIC values <i>Escherichia coli</i> (5); <i>Staphylococcus aureus</i> (3); <i>Aggregatibacter actinomycetemcomitans</i> (1); <i>Bacillus subtilis</i> (1); <i>Fusobacterium nucleatum</i> (1); <i>Porphyromonas gingivalis</i> (1); <i>Prevotella intermedia</i> (1)	15 MIC values <i>Escherichia coli</i> (5); <i>Campylobacter jejuni</i> (2); <i>Staphylococcus aureus</i> (2); <i>Aggregatibacter actinomycetemcomitans</i> (1); <i>Fusobacterium nucleatum</i> (1); <i>Klebsiella pneumoniae</i> (1); <i>Streptococcus mutans</i> (1); <i>Porphyromonas gingivalis</i> (1); <i>Prevotella intermedia</i> (1)
Yeast/ fungi	14 L(E)C50 values <i>Saccharomyces cerevisiae</i> (4); <i>Candida albicans</i> (3); <i>C. krusei</i> (1); <i>C. parapsilosis</i> (1); <i>C. tropicalis</i> (1); <i>Malassezia sympodialis</i> (1); <i>T. mentagrophytes</i> (1)	4 L(E)C50 values <i>Saccharomyces cerevisiae</i> (4)	7 L(E)C50 values <i>Aspergillus niger</i> (1); <i>Fusarium oxysporum</i> (1); <i>Penicillium expansum</i> (1); <i>Botrytis cinerea</i> (1); <i>Candida albicans</i> (1); <i>Saccharomyces cerevisiae</i> (2)
Mammalian cells in vitro	29 EC50 values <i>THP-1</i> (9); <i>HepG2</i> (7); <i>A549</i> (4); <i>MCF-7</i> (3); <i>SGC-7901</i> (3); murine osteoblasts (2)	21 EC50 values <i>A549</i> (11); <i>Caco-2</i> (4); <i>Hep G2</i> (2); human keratinocytes (1); human sebocytes (1); <i>RAW 264.7</i> (1)	25 EC50 values <i>A549</i> cells (6); <i>Ana-1</i> (4); <i>MH-S</i> (4); <i>BEAS-2B</i> (2); <i>Neuro.2A</i> (2); rat alveolar epithelial cells (2); <i>RAW 264.7</i> (2); <i>HeLa</i> (1); <i>HEK296</i> (1); human skin fibroblasts (1); rat liver mitochondria (1)
Vibrio fischeri	2 EC50 values <i>Vibrio fischeri</i> (2)	4 EC50 values <i>Vibrio fischeri</i> (4)	4 EC50 values <i>Vibrio fischeri</i> (4)
Protozoa	7 EC50 values <i>Paramecium caudatum</i> (4); <i>Tetrahymena thermophila</i> (3)	6 EC50 values <i>Paramecium caudatum</i> (4); <i>Tetrahymena thermophila</i> (3)	9 EC50 values <i>Tetrahymena thermophila</i> (8); <i>Paramecium multimicronucleatum</i> (1)

Table S3 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **silver nanoparticles** for various organisms. Data in Tables S3-S8 are arranged according to the decreasing sensitivity (increasing median L(E)C50 values) of test organisms to silver nanoparticles

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		0.01 (0.001-0.04)	0.01 (0.001-0.04)			
<i>Daphnia magna</i>	48 h LC50	0.007	0.007	10 nm (provider), uncoated	mineral medium	Hoheisel et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.01	0.01	20 nm (provider), uncoated	mineral medium	Hoheisel et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.02	0.02	30 nm (provider), uncoated	mineral medium	Hoheisel et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.03	0.03	50 nm (provider), uncoated	mineral medium	Hoheisel et al. 2012
<i>Ceriodaphnia dubia</i>	48 h LC50	0.005	0.005	21 ± 4 nm (TEM), citrate	mineral medium	Kennedy et al. 2012
<i>Ceriodaphnia dubia</i>	48 h LC50	0.03	0.03	102 ± 6 nm (TEM), citrate	mineral medium	Kennedy et al. 2012
<i>Daphnia magna</i>	24 h EC50	0.01	0.01	<100 nm (provider), uncoated	mineral medium	Jo et al. 2012
<i>Daphnia magna</i>	24 h LC50	0.01	0.01	35-50 (TEM), PVP	mineral medium	Poynton et al. 2012
<i>Daphnia magna</i>	24 h LC50	0.004	0.004	35 (TEM), citrate	mineral medium	Poynton et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.03	0.03	< 100 nm (TEM), lactate	mineral medium	Zhao and Wang 2012
<i>Daphnia magna</i>	48 h LC50	0.002	0.002	10-20 nm (TEM), PVP	mineral medium	Zhao and Wang 2012
<i>Daphnia magna</i>	48 h LC50	0.001	0.001	10-20 nm (TEM), sodium dodecylbenzene sulfonate	mineral medium	Zhao and Wang 2012

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Daphnia magna</i>	48 h EC50	0.001	0.001	60 nm (provider), uncoated	mineral medium	Kim et al. 2011
<i>Daphnia magna</i>	48 h EC50	0.002	0.002	300 nm (provider), uncoated	mineral medium	Kim et al. 2011
<i>Daphnia pulex</i>	48 h, LC50	0.04	0.04	26.6 ± 8.8 (SEM)	mineral medium	Griffitt et al. 2008
<i>Daphnia magna</i>	48 h EC50	0.02	0.02	8.4 ± 2.8 (TEM), PVP	mineral medium	Blinova et al. 2012
<i>Daphnia magna</i>	48 h EC50	0.04	0.04	12.5 ± 4.0 (TEM), protein	mineral medium	Blinova et al. 2012
Algae: Median (min-max) L(E)C50		0.36 (0.005-21.20)	0.36 (0.005-21.20)			
<i>Raphidocelis subcapitata</i> (<i>Pseudokirchneriella subcapitata</i>)	4.5 h EC50	21.20	21.20	35 nm (provider), uncoated	mineral medium	Wang et al. 2012a
<i>Raphidocelis subcapitata</i> (<i>Pseudokirchneriella subcapitata</i>)	4.5 h EC50	4.01	4.01	80 nm (provider), PVP	mineral medium	Wang et al. 2012a
<i>Raphidocelis subcapitata</i> (<i>Pseudokirchneriella subcapitata</i>)	4.5 h EC50	0.90	0.90	15 nm (provider), with a dispersant	mineral medium	Wang et al. 2012a
<i>Chlamydomonas reinhardtii</i>	1 h EC50	0.36	0.36	25 ± 13 nm (DLS, TEM), carbonate	mineral medium	Navarro et al. 2008
<i>Chlamydomonas reinhardtii</i>	2 h EC50	0.11	0.11	25 ± 13 nm (DLS, TEM), carbonate	mineral medium	Navarro et al. 2008
<i>Chlamydomonas reinhardtii</i>	3 h EC50	0.095	0.095	25 ± 13 nm (DLS, TEM), carbonate	mineral medium	Navarro et al. 2008
<i>Chlamydomonas reinhardtii</i>	4 h EC50	0.086	0.086	25 ± 13 nm (DLS, TEM), carbonate	mineral medium	Navarro et al. 2008
<i>Chlamydomonas reinhardtii</i>	5 h EC50	0.089	0.089	25 ± 13 nm (DLS, TEM), carbonate	mineral medium	Navarro et al. 2008
<i>Pseudokirchneriella subcapitata</i>	96 h IC50	1.60	1.60	25.4 nm (provider), uncoated	natural water	McLaughlin and Bonzongo 2011

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Pseudokirchneriella subcapitata</i>	96 h IC50	0.023	0.023	25.4 nm (provider), uncoated	natural water	McLaughlin and Bonzongo 2011
<i>Pseudokirchneriella subcapitata</i>	96 h IC50	0.0046	0.0046	25.4 nm (provider), uncoated	mineral medium	McLaughlin and Bonzongo 2011
<i>Pseudokirchneriella subcapitata</i>	96 h EC50	0.020	0.020	41 nm (TEM), PVP	mineral medium	Kennedy et al. 2010
<i>Chlorella vulgaris</i>	24 h EC50	8.9**	8.9**	50 nm (provider), uncoated	mineral medium	Oukarroum et al. 2012
<i>Dunaliella tertiolecta</i>	24 h EC50	2.0**	2.0**	50 nm (provider), uncoated	complex medium	Oukarroum et al. 2012
<i>Ochromonas danica</i>	48 h EC50	20.8**	20.8**	90% of the particles were in the 1-10 nm range (provider), carboxy-functionalized	complex medium	Miao et al. 2010
<i>Chattonella marina</i>	1 h EC50	1.7**	1.7**	56 ± 5 nm (DLS), citrate stabilized	complex medium	He et al. 2012
<i>Pseudokirchneriella subcapitata</i>	96 h EC50	0.19	0.19	20-30 nm (provider), sodium citrate stabilized	mineral medium	Griffitt et al. 2008
Fish: Median (min-max) L(E)C50		1.36 (0.03-12.6)	1.36 (0.03-12.6)			
<i>Oryzias latipes</i>	48 h LC50	1.38	1.38	29.9 nm (TEM), PVP	water	Wu and Zhou 2013
<i>Oryzias latipes</i>	72 h LC50	1.12	1.12	29.9 nm (TEM), PVP	water	Wu and Zhou 2013
<i>Oryzias latipes</i>	96 h LC50	0.87	0.87	29.9 nm (TEM), PVP	water	Wu and Zhou 2013
<i>Pimephales promelas</i>	96 h LC50	0.09	0.09	10 nm (provider)	water	Hoheisel et al. 2012
<i>Danio rerio</i>	96 h EC50	1.61	1.61	35 nm (provider)	water	Wang et al. 2012a
<i>Danio rerio</i>	96 h EC50	1.36	1.36	80 nm (provider), PVP	water	Wang et al. 2012a

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Danio rerio</i>	96 h EC50	0.78	0.78	15 nm (provider)	water	Wang et al. 2012a
<i>Oryzias latipes</i>	72 h LC50	10.0**	10.0**	10 nm (provider), PVP	water	Kwok et al. 2012
<i>Oryzias latipes</i>	72 h LC50	2.50**	2.50**	50 nm (provider), PVP	water	Kwok et al. 2012
<i>Oryzias latipes</i>	72 h LC50	10.0**	10.0**	7 nm (provider), citrate	water	Kwok et al. 2012
<i>Oryzias latipes</i>	96 h LC50	1.39	1.39	3.6 nm (provider)	water	Kashiwada et al. 2012
<i>Oreochromis mossambicus</i>	8 d LC50	12.6	12.6	<100 nm (TEM), citrate	water	Govindasamy and Rahuman 2012
<i>Pimephales promelas</i>	96 h LC50	9.40	9.40	35 nm (provider)	water	Laban et al. 2010
<i>Pimephales promelas</i>	96 h LC50	1.25	1.25	35 nm (provider)	water	Laban et al. 2010
<i>Pimephales promelas</i>	96 h LC50	10.6	10.6	>100 nm (provider)	water	Laban et al. 2010
<i>Pimephales promelas</i>	96 h LC50	1.36	1.36	>100 nm (provider)	water	Laban et al. 2010
<i>Oryzias latipes</i>	96 h LC50	0.03	0.03	49.6 (TEM)	water	Chae et al. 2009
Nematode: Median (min-max) L(E)C50		3.34 (0.1-55)	3.34 (0.1-55)			
<i>Caenorhabditis elegans</i>	72 h EC50	13.4	13.4	1 nm	mineral medium	Ellegaard-Jensen et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	2.80	2.80	28 nm, PVP	mineral medium	Ellegaard-Jensen et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.10**	0.10**	<100 nm	mineral medium	Lim et al. 2012

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Caenorhabditis elegans</i>	72 h EC50	55.0	55.0	50.6 nm (TEM), citrate	complex medium	Kim et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	1.40	1.40	8 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.86	0.86	8 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	1.83	1.83	38 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.38	0.38	38 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	49.9	49.9	21 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	4.31	4.31	21 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	45.0	45.0	75 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	11.6	11.6	75 nm, PVP	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	39.8	39.8	7 nm, citrate	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	3.34	3.34	7 nm, citrate	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.86	0.86	5 nm, GA	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.10	0.10	5 nm, GA	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	1.51	1.51	5 nm, GA	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.29	0.29	5 nm, GA	mineral medium	Yang et al. 2012

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Caenorhabditis elegans</i>	72 h EC50	~25.0	25.0**	25 nm, PVP	mineral medium	Meyer et al. 2010
<i>Caenorhabditis elegans</i>	72 h EC50	~25.0	25.0**	50 nm, PVP	mineral medium	Meyer et al. 2010
<i>Caenorhabditis elegans</i>	72 h EC50	~25.0	25.0**	5 nm, citrate	mineral medium	Meyer et al. 2010
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		7.1 (0.5-250)	7.1 (0.5-250)			
<i>Prevotella intermedia</i>	48 h MIC	100	100	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Porphyromonas gingivalis</i>	48 h MIC	250	250	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Fusobacterium nucleatum</i>	48 h MIC	100	100	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Aggregatibacter actinomycetemcomitans</i>	48 h MIC	100	100	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Streptococcus mutants</i>	24 h MIC	4.9	4.9	25 nm (TEM), PVP	complex medium	Hernández-Sierra et al. 2008
<i>Staphylococcus aureus</i>	24 h MIC	10	10	10 nm (TEM), PVP	complex medium	Cho et al. 2005
<i>Escherichia coli</i>	24 h MIC	10	10	10 nm (TEM), PVP	complex medium	Cho et al. 2005
<i>Escherichia coli</i>	24 h MIC	6.3	6.3	7 nm (TEM)	complex medium	Martínez-Castanón et al. 2008
<i>Escherichia coli</i>	24 h MIC	13	13	29 nm (TEM)	complex medium	Martínez-Castanón et al. 2008
<i>Escherichia coli</i>	24 h MIC	11.8	11.8	89 nm (TEM)	complex medium	Martínez-Castanón et al. 2008
<i>Staphylococcus aureus</i>	24 h MIC	7.5	7.5	7 nm (TEM)	complex medium	Martínez-Castanón et al. 2008

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Staphylococcus aureus</i>	24 h MIC	16.7	16.7	29 nm (TEM)	complex medium	Martínez-Castanón et al. 2008
<i>Staphylococcus aureus</i>	24 h MIC	33.7	33.7	89 nm (TEM)	complex medium	Martínez-Castanón et al. 2008
<i>Pseudomonas aeruginosa</i> , <i>Bacillus subtilis</i> , <i>Salmonella abony</i> , <i>Salmonella typhimurium</i> , <i>Klebsiella aerogenes</i> , <i>Proteus vulgaris</i> , <i>Bacillus subtilis</i> , <i>Staphylococcus epidermidis</i> , <i>Escherichia coli</i>	24 h MIC	6.3	6.3	7-20 nm, biostabilized	complex medium	Jain et al. 2009
<i>Staphylococcus aureus</i> ,	24 h MIC	12.5	12.5	7-20 nm, biostabilized	complex medium	Jain et al. 2009
<i>Escherichia coli</i> (4 different strains)	MIC	40-180	40-180	3.3 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Bacillus subtilis</i>	MIC	40	40	3.3 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Staphylococcus aureus</i> (3 different strains)	MIC	120	120	3.3 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Escherichia coli</i>	8 h MIC	2	2	4 nm (TEM), biogenic	complex medium	Suresh et al. 2010
<i>Bacillus subtilis</i>	8 h MIC	0.5	0.5	4 nm (TEM), biogenic	complex medium	Suresh et al. 2010
<i>Shewanella oneidensis</i>	8 h MIC	3	3	4 nm (TEM), biogenic	complex medium	Suresh et al. 2010
<i>Escherichia coli</i>	8 h MIC	6	6	<20 nm	complex medium	Suresh et al. 2010
<i>Bacillus subtilis</i>	8 h MIC	2	2	<20 nm	complex medium	Suresh et al. 2010
<i>Shewanella oneidensis</i>	8 h MIC	6.5	6.5	<20 nm	complex medium	Suresh et al. 2010
<i>Enterococcus faecalis</i>	24 h MIC	13.5	13.5	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Staphylococcus aureus</i>	24 h MIC	6.8	6.8	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Escherichia coli</i>	24 h MIC	3.4	3.4	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Pseudomonas aeruginosa</i> (2 different strains)	24 h MIC	3.4-6.8	3.4-6.8	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Staphylococcus epidermidis</i>	24 h MIC	1.6	1.6	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Staphylococcus aureus</i>	24 h MIC	6.8	6.8	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Enterococcus faecium</i>	24 h MIC	13.5	13.5	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
<i>Klebsiella pneumoniae</i>	24 h MIC	6.8	6.8	25 nm (TEM), mono and disaccharides	complex medium	Panáček et al. 2006
Yeast/fungi: Median (min-max) L(E)C50		7.90 (0.88-48.5)	7.90 (0.88-48.5)			
<i>Saccharomyces cerevisiae</i>	MIC	48.5	48.5	8.6 nm (TEM)	complex medium	Debabrata and Giasuddin 2013
<i>Candida albicans</i> , <i>Candida glabrata</i>	48 h MIC	0.88**	0.88**	50 nm (provider)	complex medium	Brandt et al. 2012
<i>Malassezia sympodialis</i>	48 h MIC	8.80**	8.80**	50 nm (provider)	complex medium	Brandt et al. 2012
<i>Saccharomyces cerevisiae</i>	IC30	15.1	15.1	5-10 nm	complex medium	Niazi et al. 2011
<i>Saccharomyces cerevisiae</i>	MIC	20.0**	20.0**	5-10 nm	complex medium	Niazi et al. 2011
<i>Candida glabrata</i>	24 h IC50	13.5**	13.5**	5-60 nm	complex medium	Monteiro et al. 2012
<i>Saccharomyces cerevisiae</i>	48 h MIC	2.0	2.0	3 nm (TEM)	complex medium	Kim et al. 2009a

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Candida albicans</i>	48 h MIC	2.0	2.0	3 nm (TEM)	complex medium	Kim et al. 2009a
<i>Candida albicans</i>	24 h IC80	2-4	2-4	3 nm (TEM)	complex medium	Kim et al. 2008
<i>Candida tropicalis</i>	24 h IC80	7.0	7.0	3 nm (TEM)	complex medium	Kim et al. 2008
<i>Candida glabrata</i>	24 h IC80	1-7	1-7	3 nm (TEM)	complex medium	Kim et al. 2008
<i>Candida parapsilosis</i>	24 h IC80	4-25	4-25	3 nm (TEM)	complex medium	Kim et al. 2008
<i>Candida krusei</i>	24 h IC50	13.0	13.0	3 nm (TEM)	complex medium	Kim et al. 2008
<i>Trichophyton mentagrophytes</i>	24 h IC50	1-4	1-4	3 nm (TEM)	complex medium	Kim et al. 2008
Mammalian cells in vitro: Median (min-max) L(E)C50		11.3 (0.5-140)	11.3 (0.5-140)			
Murine osteoclasts	IC50 (time not specified)	14	14	50 nm	not specified	Albers et al. 2012
Murine osteoblasts	IC50 (time not specified)	1	1	50 nm	not specified	Albers et al. 2012
A549 cells	24 h IC50	11.25	11.25	30-50 nm (provider), PVP	complex medium	Foldbjerg et al. 2011
A549 cells	24 h IC50	7.50	7.50	30-50 nm (provider), PVP	complex medium	Foldbjerg et al. 2011
THP-1 cells	24 h EC50	2.44	2.44	69 ± 3 nm (TEM), PVP	complex medium	Foldbjerg et al. 2009
HepG2 cells	28 h IC50	3.38	3.38	<10 nm (TEM), uncoated	complex medium	Kim et al. 2009b
HepG2 cells	28 h IC50	1.95	1.95	<10 nm (TEM), uncoated	complex medium	Kim et al. 2009b

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
HepG2 cells	28 h IC50	0.53	0.53	<10 nm (TEM), uncoated	complex medium	Kim et al. 2009b
THP-1 cells- derived macrophages	24 h IC50	110	110	20 nm (TEM), peptides	complex medium	Haase et al. 2012
THP-1 cells- derived macrophages	48 h IC50	18	18	20 nm (TEM), peptides	complex medium	Haase et al. 2012
THP-1 cells- derived macrophages	24 h IC50	140	140	40 nm (TEM), peptides	complex medium	Haase et al. 2012
THP-1 cells- derived macrophages	48 h IC50	30	30	40 nm (TEM), peptides	complex medium	Haase et al. 2012
A549 cells	24 h EC50	1.0	1.0	5 nm (TEM), PVP	complex medium	Liu et al. 2010
A549 cells	24 h EC50	10.0	10.0	20 nm (TEM), PVP	complex medium	Liu et al. 2010
A549 cells	24 h EC50	13.3	13.3	50 nm (TEM), PVP	complex medium	Liu et al. 2010
HepG2 cells	24 h EC50	0.59	0.59	5 nm (TEM), PVP	complex medium	Liu et al. 2010
HepG2 cells	24 h EC50	25.5	25.5	20 nm (TEM), PVP	complex medium	Liu et al. 2010
HepG2 cells	24 h EC50	33.6	33.6	50 nm (TEM), PVP	complex medium	Liu et al. 2010
MCF-7 cells	24 h EC50	0.5	0.5	5 nm (TEM), PVP	complex medium	Liu et al. 2010
MCF-7 cells	24 h EC50	14.3	14.3	20 nm (TEM), PVP	complex medium	Liu et al. 2010
MCF-7 cells	24 h EC50	47.6	47.6	50 nm (TEM), PVP	complex medium	Liu et al. 2010
SGC-7901 cells	24 h EC50	9.9	9.9	5 nm (TEM), PVP	complex medium	Liu et al. 2010

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Ag/L	Primary size (method used for determination), coating	Test medium*	Reference
SGC-7901 cells	24 h EC50	50.9	50.9	20 nm (TEM), PVP	complex medium	Liu et al. 2010
SGC-7901 cells	24 h EC50	112.0	112.0	50 nm (TEM), PVP	complex medium	Liu et al. 2010
HepG2 cells	24 h IC50	2.9	2.9	5-10 nm (provider)	complex medium	Nowrouzi et al. 2010
<i>Vibrio fischeri</i> : Median (min-max) L(E)C50		32 (29-35)	32 (29-35)			
<i>Vibrio fischeri</i>	30 min EC50	35	35	10-25 nm (TEM), uncoated	mineral medium	Binaeian et al. 2012
<i>Vibrio fischeri</i>	30 min EC50	29	29	50-100 nm (TEM), uncoated	mineral medium	Binaeian et al. 2012
Protozoa : Median (min-max) L(E)C50		38.0 (1.46-286)	38.0 (1.46-286)			
<i>Tetrahymena thermophila</i>	24 h EC50	205.0	205.0	<100 nm (provider), uncoated	water	Juganson et al. 2013
<i>Tetrahymena thermophila</i>	2 h EC50	286.0	286.0	<100 nm (provider), uncoated	water	Juganson et al. 2013
<i>Tetrahymena pyriformis</i>	24 h IC50	1.46	1.46	9 ± 1.6 nm (TEM), ATP stabilized	complex medium	Shi et al. 2012
<i>Paramecium caudatum</i>	1 h LC50	39.0	39.0	40 nm (DLS), uncoated	aqueous medium	Kvitek et al. 2009
<i>Paramecium caudatum</i>	1 h LC50	16.0	16.0	40 nm (DLS), Tween 80 (1% w/w)	aqueous medium	Kvitek et al. 2009
<i>Paramecium caudatum</i>	1 h LC50	38.0	38.0	40 nm (DLS), PEG 35000 (1% w/w)	aqueous medium	Kvitek et al. 2009
<i>Paramecium caudatum</i>	1 h LC50	36.0	36.0	40 nm (DLS), PVP 360 (1% w/w)	aqueous medium	Kvitek et al. 2009

DLS – dynamic light scattering; GA – gum arabic; PVP – polyvinylpyrrolidone; SEM – scanning electron microscopy; TEM – transmission electron microscopy

*Mineral medium - without organic components; Complex medium - with organic components; ** estimated value

Table S4 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **silver salt** for various organisms

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		0.0009 (0.0005-0.008)			
<i>Daphnia magna</i>	48 h LC50	0.001	AgNO ₃	mineral medium	Hoheisel et al. 2012
<i>Ceriodaphnia dubia</i>	48 h LC50	0.0007	AgNO ₃	mineral medium	Kennedy et al. 2012
<i>Daphnia magna</i>	24 h EC50	0.002	AgNO ₃	mineral medium	Jo et al. 2012
<i>Daphnia magna</i>	24 h LC50	0.0007	AgNO ₃	mineral medium	Poynton et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.0006	AgNO ₃	mineral medium	Zhao and Wang 2012
<i>Daphnia magna</i>	48 h EC50	0.0005	AgNO ₃	mineral medium	Kim et al. 2011
<i>Daphnia pulex</i>	48 h LC50	0.008	AgNO ₃	mineral medium	Griffitt et al. 2008
<i>Daphnia magna</i>	48 h EC50	0.002	AgNO ₃	mineral medium	Blinova et al. 2012
Algae: Median (min-max) L(E)C50		0.008 (0.0013-0.59)			
<i>Raphidocelis subcapitata</i> (<i>Pseudokirchneriella subcapitata</i>)	4.5 h EC50	0.031	AgNO ₃	mineral medium	Wang et al. 2012a
<i>Chlamydomonas reinhardtii</i>	1 h EC50	0.020	AgNO ₃	mineral medium	Navarro et al. 2008
<i>Chlamydomonas reinhardtii</i>	2 h EC50	0.020	AgNO ₃	mineral medium	Navarro et al. 2008
<i>Pseudokirchneriella subcapitata</i>	96 h EC50	0.010	AgNO ₃	mineral medium	Kennedy et al. 2010

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
<i>Ochromonas danica</i>	48 h EC50	0.005	not specified	complex medium	Miao et al. 2010
<i>Chattonella marina</i>	1 h EC50	0.59**	AgNO ₃	complex medium	He et al. 2012
<i>Chlamydomonas reinhardtii</i>	24 h EC50	0.0016	AgNO ₃	mineral medium	Hiriart-Baer et al. 2006
<i>Pseudokirchneriella subcapitata</i>	24 h EC50	0.0024	AgNO ₃	mineral medium	Hiriart-Baer et al. 2006
<i>Chlamydomonas reinhardtii</i>	6 h EC50	0.0013	AgNO ₃	mineral medium	Lee et al. 2005
<i>Pseudokirchneriella subcapitata</i>	6 h EC50	0.0028	AgNO ₃	mineral medium	Lee et al. 2005
Fish: Median (min-max) L(E)C50		0.058 (0.005-0.15)			
<i>Pimephales promelas</i>	96 h LC50	0.005	AgNO ₃	water	Hoheisel et al. 2012
<i>Danio rerio</i>	96 h LC50	0.08	AgNO ₃	water	Wang et al. 2012a
<i>Oryzias latipes</i>	72 h LC50	0.15	AgNO ₃	water	Kwok et al. 2012
<i>Oryzias latipes</i>	96 h LC50	0.04	AgNO ₃	water	Chae et al. 2009
Nematode: Median (min-max) L(E)C50		4.77 (0.06-22)			
<i>Caenorhabditis elegans</i>	24 h LC50	22.0	AgNO ₃	complex medium	Kim et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	6.36	AgNO ₃	mineral medium	Yang et al. 2012
<i>Caenorhabditis elegans</i>	72 h EC50	0.06	AgNO ₃	mineral medium	Yang et al. 2012

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
<i>Caenorhabditis elegans</i>	72 h EC50	3.17	AgNO ₃	mineral medium	Meyer et al. 2010
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		3.3 (0.004-108)			
<i>Escherichia coli</i>	Overnight MIC	0.5	AgNO ₃	complex medium	Greulich et al. 2012
<i>Escherichia coli</i>	Overnight MIC	3.3	AgNO ₃	complex medium	Greulich et al. 2012
<i>Staphylococcus aureus</i> (2 different isolates)	Overnight MIC	0.5-2.5	AgNO ₃	complex medium	Greulich et al. 2012
<i>Escherichia coli</i>	8 h MIC	4.9	AgNO ₃	complex medium	Suresh et al. 2010
<i>Bacillus subtilis</i>	8 h MIC	3.2	AgNO ₃	complex medium	Suresh et al. 2010
<i>Shewanella oneidensis</i>	8 h MIC	5.2	AgNO ₃	complex medium	Suresh et al. 2010
<i>Pseudomonas aeruginosa</i> (2 different isolates)	Overnight MIC	1.1-110	AgNO ₃	complex medium	Shakibaie and Harati 2004
<i>Legionella pneumophila</i>	12 h MIC	0.04	AgCl	not specified	Lin et al. 1996
<i>Enterococcus faecalis</i>	24 h MIC	6.8	AgNO ₃	complex medium	Panáček et al. 2006
<i>Staphylococcus aureus</i>	24 h MIC	6.8	AgNO ₃	complex medium	Panáček et al. 2006
<i>Escherichia coli</i>	24 h MIC	1.7	AgNO ₃	complex medium	Panáček et al. 2006
<i>Pseudomonas aeruginosa</i>	24 h MIC	0.9	AgNO ₃	complex medium	Panáček et al. 2006
<i>Pseudomonas aeruginosa</i>	24 h MIC	0.8	AgNO ₃	complex medium	Panáček et al. 2006

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
<i>Staphylococcus epidermidis</i>	24 h MIC	0.8	AgNO ₃	complex medium	Panáček et al. 2006
<i>Staphylococcus aureus</i>	24 h MIC	6.8	AgNO ₃	complex medium	Panáček et al. 2006
<i>Enterococcus faecium</i>	24 h MIC	3.4	AgNO ₃	complex medium	Panáček et al. 2006
<i>Klebsiella pneumonia</i>	24 h MIC	3.4	AgNO ₃	complex medium	Panáček et al. 2006
<i>Legionella pneumophila</i>	24 h MIC	0.06	AgNO ₃	water	Unger and Lück 2012
<i>Staphylococcus aureus</i>	24 h MIC	0.06	AgNO ₃	water	Unger and Lück 2012
<i>Pseudomonas aeruginosa</i>	24 h MIC	0.02	AgNO ₃	water	Unger and Lück 2012
<i>Escherichia coli</i>	24 h MIC	0.004	AgNO ₃	water	Unger and Lück 2012
<i>Staphylococcus aureus</i> (2 different isolates)	20 h MIC	12.5	AgNO ₃	complex medium	Zhao et al. 1998
<i>Escherichia coli</i>	20 h MIC	12.5	AgNO ₃	complex medium	Zhao et al. 1998
<i>Pseudomonas aeruginosa</i>	20 h MIC	12.5	AgNO ₃	complex medium	Zhao et al. 1998
Yeast/fungi: Median (min-max) L(E)C50		2.16 (0.22-10.0)			
<i>Saccharomyces cerevisiae</i>	IC30	5.04	AgNO ₃	complex medium	Niazi et al. 2011
<i>Saccharomyces cerevisiae</i>	MIC	10.0**	AgNO ₃	complex medium	Niazi et al. 2011
<i>Saccharomyces cerevisiae</i>	24 h IC50	2.16	AgNO ₃	complex medium	Jin et al. 2008

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
<i>Saccharomyces cerevisiae</i>	96 h IC50	0.22	AgNO ₃	complex medium	Ershov et al. 1997
<i>Saccharomyces cerevisiae</i>	96 h MIC	0.54	AgNO ₃	complex medium	Ershov et al. 1997
Mammalian cells in vitro: Median (min-max) L(E)C50		2.0 (0.62-14)			
Human mesenchymal stem cells	24 h IC50	2.0	AgC ₂ H ₃ O ₂	complex medium	Greulich et al. 2012
human peripheral blood mononuclear cells	24 h IC50	2.0	AgC ₂ H ₃ O ₂	complex medium	Greulich et al. 2012
Murine osteoblasts	IC50 (time not specified)	1	AgNO ₃	not specified	Albers et al. 2012
Murine osteoclasts	IC50 (time not specified)	14	AgNO ₃	not specified	Albers et al. 2012
A549 cells	24 h IC50	5.5	AgNO ₃	complex medium	Foldbjerg et al. 2011
THP-1 cells	24 h EC50	0.62	AgNO ₃	complex medium	Foldbjerg et al. 2009
HepG2 cells	28 h IC50	1.4	AgNO ₃	complex medium	Kim et al. 2009b
HepG2 cells	28 h IC50	1.8	AgNO ₃	complex medium	Kim et al. 2009b
HepG2 cells	28 h IC50	0.8	AgNO ₃	complex medium	Kim et al. 2009b
A549 cells	24 h EC50	2.3	AgNO ₃	complex medium	Liu et al. 2010
HepG2 cells	24 h EC50	0.7	AgNO ₃	complex medium	Liu et al. 2010
MCF-7 cells	24 h EC50	1.2	AgNO ₃	complex medium	Liu et al. 2010

Species	Exposure time and endpoint	EC50 on metal basis, mg Ag/L	Tested as (compound)	Test medium*	Reference
SGC-7901 cells	24 h EC50	2.1	AgNO ₃	complex medium	Liu et al. 2010
primary human endometrial epithelial cells	24 h TC50	3.9	Ag ₂ SO ₄	complex medium	Wu et al. 2012
Caco-2 cells (proliferating)	24 h IC50	1.5**	AgNO ₃	complex medium	Böhmert et al. 2012
Caco-2 cells (proliferating)	48 h IC50	1.5**	AgNO ₃	complex medium	Böhmert et al. 2012
Caco-2 cells (differentiated)	24 h IC50	7.5**	AgNO ₃	complex medium	Böhmert et al. 2012
Caco-2 cells (differentiated)	48 h IC50	3.5**	AgNO ₃	complex medium	Böhmert et al. 2012
<i>Vibrio fischeri</i> : Median (min-max) EC50		5.7 (1.9-9.5)			
<i>Vibrio fischeri</i>	30 min EC50	1.9	AgNO ₃	mineral medium	Binaeian et al. 2012
<i>Vibrio fischeri</i>	5 min EC50	9.5	AgNO ₃	mineral medium	Hsieh et al. 2004
Protozoa : Median (min-max) L(E)C50		1.50 (1.46-1.8)			
<i>Tetrahymena thermophila</i>	24 h EC50	1.5	AgNO ₃	water	Juganson et al. 2013
<i>Tetrahymena thermophila</i>	2 h EC50	1.8	AgNO ₃	water	Juganson et al. 2013
<i>Tetrahymena pyriformis</i>	24 h IC60	1.46**	AgNO ₃	complex medium	Shi et al. 2012

*Mineral medium - without organic components; Complex medium - with organic components; ** estimated value

Table S5 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **CuO nanoparticles** for various organisms

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Cu/L	Primary size (method used for determination), coating	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		2.1 (0.08-12.3)	1.7 (0.06-9.8)			
<i>Daphnia magna</i>	24 h EC50	0.10	0.078	<100 nm (provider), uncoated	mineral medium	Jo et al. 2012
<i>Daphnia magna</i>	48 h LC50	3.2	2.56	30 nm (provider), uncoated	mineral medium	Heinlaan et al. 2008
<i>Daphnia magna</i>	48 h EC50	3.3	2.6	30 nm (provider), uncoated	mineral medium	Blinova et al. 2010
<i>Thamnocephalus platyurus</i>	24 h LC50	12.3	9.80	30 nm (provider), uncoated	mineral medium	Manusadžianas et al. 2012
<i>Daphnia magna</i>	48 h EC50	5.9	4.72	<50 nm (provider), uncoated	mineral medium	Sovova et al. 2009
<i>Daphnia pulex</i>	48 h LC50	0.08	0.06	15-45 nm (provider), uncoated	mineral medium	Griffitt et al. 2008
<i>Daphnia magna</i>	48 h LC50	0.99	0.79	40 nm (provider), uncoated	mineral medium	Zhao et al. 2012
<i>Daphnia magna</i>	48 h EC50	0.46	0.37	40 nm (provider), uncoated	mineral medium	Zhao et al. 2012
Algae: Median (min-max) L(E)C50		2.80 (0.68-47.0)	2.24 (0.54-37.55)			
<i>Pseudokirchneriella subcapitata</i>	96 h EC50	0.68	0.54	15-45 nm (provider), uncoated	mineral medium	Griffitt et al. 2008
<i>Nitellopsis obtusa</i>	96 h LC50	2.80	2.24	< 50 nm (mean ~30 nm) (provider), uncoated	mineral medium	Manusadžianas et al. 2012
<i>Chlorella spp.</i>	30 min IC50	47.0	37.55	< 50 nm (mean ~30 nm) (provider), uncoated	mineral medium	Manusadžianas et al. 2012
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.89	0.71	mean 30 nm (provider), uncoated	complex medium	Aruoja et al. 2009

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Cu/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Chlamydomonas reinhardtii</i>	6 h EC50	19.0**	15.18**	30-40 nm (DLS), poly(styrene-co-butyl acrylate)	mineral medium	Perreault et al. 2012
Fish: Median (min-max) L(E)C50		100	79.9			
<i>Cyprinus carpio</i>	96 h EC50	100**	79.9**	20-40 nm (TEM)	water	Zhao et al. 2011
Nematode: Median (min-max) L(E)C50		not analyzed				
not analyzed	not analyzed	not analyzed	not analyzed	not analyzed	not analyzed	not analyzed
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		200 (20-280)	160 (16-224)			
<i>Escherichia coli</i> (4 different strains)	MIC	140-280	112-224	9.2 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Bacillus subtilis</i>	MIC	20	16	9.2 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Staphylococcus aureus</i> (3 different strains)	MIC	140	112	9.2 nm (TEM)	complex medium	Ruparelia et al. 2008
<i>Prevotella intermedia</i> , <i>Fusobacterium nucleatum</i> , <i>Porphyromonas gingivalis</i> , <i>Aggregatibacter actinomycetemcomitans</i>	48 h MIC	250	200	12-50 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Escherichia coli</i>	20 h MIC	25	20	6 nm (TEM)	complex medium	Hassan et al. 2012
Yeast/fungi: Median (min-max) L(E)C50		17.1 (4.80-643)	13.6 (3.83-514)			
<i>Saccharomyces cerevisiae</i>	24 h IC50	643	514	30 nm (provider)	complex medium	Kasemets et al. 2013
<i>Saccharomyces cerevisiae</i>	24 h IC50	4.80	3.83	30 nm (provider)	water	Kasemets et al. 2013
<i>Saccharomyces cerevisiae</i>	8 h EC50	20.7	16.5	30 nm (provider)	complex medium	Kasemets et al. 2009

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Cu/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Saccharomyces cerevisiae</i>	24 h EC50	13.4	10.7	30 nm (provider)	complex medium	Kasemets et al. 2009
Mammalian cells in vitro: Median (min-max) L(E)C50		25 (13-100)	20 (10.4-80)			
A549 cells	24 h IC50	15	12	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	24 h IC50	13	10.4	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	24 h IC50	17	13.6	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	24 h IC50	20	16	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	18 h IC50	25	20	20-40 nm (TEM)	complex medium	Karlsson et al. 2008
RAW 264.7	24 h IC50	16.5	13.2	12.8±3.4 nm (TEM)	complex medium	Zhang et al. 2012a
RAW 264.7	24 h IC50	18	14.5	12.8±3.4 nm (TEM)	complex medium	Zhang et al. 2012a
BEAS-2B cells	24 h IC50	30	24	12.8±3.4 nm (TEM)	complex medium	Zhang et al. 2012a
BEAS-2B cells	24 h IC50	30	24	12.8±3.4 nm (TEM)	complex medium	Zhang et al. 2012a
HepG2 cells	24 h IC50	25	20	Rod-shaped: 10±3 nm × 74±17 nm (TEM)	complex medium	Piret et al. 2012a
HepG2 cells	24 h IC50	25	20	40±16 nm (TEM)	complex medium	Piret et al. 2012a
Caco-2 cells	24 h IC50	37.5	30	Rod-shaped: 10±3 nm × 74±17 nm (TEM)	complex medium	Piret et al. 2012b
Caco-2 cells	24 h IC50	75	60	40±16 nm (TEM)	complex medium	Piret et al. 2012b

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Cu/L	Primary size (method used for determination), coating	Test medium*	Reference
immortalized human keratinocytes	24 h IC50	17.5	14	Rod-shaped: 10±3 nm × 74±17 nm (TEM)	complex medium	Piret et al. 2012b
immortalized human sebocytes	24 h IC50	37.5	30	Rod-shaped: 10±3 nm × 74±17 nm (TEM)	complex medium	Piret et al. 2012b
A549 cells	24 h IC50	75	60	Rod-shaped: 10±3 nm × 74±17 nm (TEM)	complex medium	Piret et al. 2012b
A549 cells	24 h IC50	100	80	40±16 nm (TEM)	complex medium	Piret et al. 2012b
A549 cells	24 h IC50	37.5	30	50-60 nm (TEM)	complex medium	Ahamed et al. 2010
A549 cells	24 h IC50	37.5	30	50-60 nm (TEM)	complex medium	Ahamed et al. 2010
A549 cells	24 h IC50	15	12	50-60 nm (TEM)	complex medium	Ahamed et al. 2010
A549 cells	24 h IC50	15	12	20-40 nm (TEM)	complex medium	Wang et al. 2012b
<i>Vibrio fischeri</i> : Median (min-max) L(E)C50		73.6 (12-204)	58.8 (9.8-163)			
<i>Vibrio fischeri</i>	30 min EC50	79	63	30 nm (provider), uncoated	mineral medium	Heinlaan et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	68	54	30 nm (provider), uncoated	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	204	163	30 nm (provider), uncoated	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	12	9.8	<50 nm (provider), uncoated	mineral medium	Sovova et al. 2009
Protozoa : Median (min-max) L(E)C50		124.5 (0.98-161.5)	99.5 (0.78-129.0)			
<i>Paramecium multimicronucleatum</i>	48 h LC50	0.98	0.78	13.3 ± 3.9 nm (TEM), uncoated	mineral medium	Li et al. 2012a

Species	Exposure time and endpoint	EC50 on compound basis, mg/L	EC50 on metal basis, mg Cu/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Tetrahymena thermophila</i>	24 h EC50	80.0	63.9	mean 30 nm (provider), uncoated	mineral medium	Mortimer et al. 2011
<i>Tetrahymena thermophila</i>	4 h EC50	159.0	127.0	mean 30 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	4 h EC50	161.5	129.0	mean 30 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	122.5	97.9	mean 30 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	126.4	101.0	mean 30 nm (provider), uncoated	mineral medium	Mortimer et al. 2010

DLS – dynamic light scattering; TEM – transmission electron microscopy

*Mineral medium - without organic components; Complex medium - with organic components; ** estimated value

Table S6 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **copper salt** for various organisms

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		0.02 (0.004-0.07)			
<i>Daphnia magna</i>	24 h EC50	0.03	CuCl ₂ *2H ₂ O	mineral medium	Jo et al. 2012
<i>Daphnia magna</i>	48 h LC50	0.07	CuSO ₄	mineral medium	Heinlaan et al. 2008
<i>Daphnia magna</i>	48 h EC50	0.07	CuSO ₄	mineral medium	Blinova et al. 2010
<i>Daphnia similis</i>	48 h EC50	0.04	CuSO ₄	mineral medium	de Oliveira-Filho et al. 2004
<i>Daphnia pulex</i>	48 h LC50	0.01	CuCl ₂	mineral medium	Griffitt et al. 2008
<i>Ceriodaphnia dubia</i>	48 h LC50	0.004	CuSO ₄	mineral medium	Harmon et al. 2003
<i>Daphnia ambigua</i>	48 h LC50	0.007	CuSO ₄	mineral medium	Harmon et al. 2003
<i>Daphnia magna</i>	48 h LC50	0.019	CuSO ₄	mineral medium	Mastin and Rodgers 2000
Algae: Median (min-max) L(E)C50		0.075 (0.0044-13.0)			
<i>Nitellopsis obtusa</i>	96 h LC50	0.11	Cu(NO ₃) ₂	mineral medium	Manusadzianas et al. 2012
<i>Chlorella spp.</i>	30 min IC50	13.0	CuSO ₄	mineral medium	Manusadzianas et al. 2012
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.020	CuSO ₄	complex medium	Aruoja et al. 2009
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.017	CuSO ₄	complex medium	Heijerick et al. 2005

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.25	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.16	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.15	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.097	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.084	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.11	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.065	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.11	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.053	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.034	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.037	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.19	CuSO ₄	natural water	Heijerick et al. 2005
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.032	CuSO ₄	natural water	Heijerick et al. 2005
<i>Chlorella sp.</i>	48 h IC50	0.0044	CuNO ₃	mineral medium	Apte et al. 2005
<i>Chlorella sp.</i>	48 h EC50	0.0057	CuSO ₄ *5H ₂ O	mineral medium	Franklin et al. 2002

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Chlorella sp.</i>	72 h EC50	0.0070	CuSO ₄ *5H ₂ O	mineral medium	Franklin et al. 2002
Fish: Median (min-max) L(E)C50		0.28 (0.01-7.50)			
<i>Oncorhynchus mykiss</i>	72 h LC50	0.10	CuSO ₄ *5H ₂ O	water	Shaw et al. 2012
<i>Oncorhynchus mykiss</i>	96 h LC50	0.04	CuSO ₄ *5H ₂ O	water	Little et al. 2012
<i>Oncorhynchus mykiss</i>	96 h LC50	0.03	CuSO ₄ *5H ₂ O	water	Little et al. 2012
<i>Danio rerio</i>	96 h LC50	0.15	CuSO ₄ *5H ₂ O	water	Alsop and Wood 2011
<i>Danio rerio</i>	96 h LC50	0.21	CuSO ₄ *5H ₂ O	water	Alsop and Wood 2011
<i>Danio rerio</i>	96 h LC50	0.01	CuSO ₄ *5H ₂ O	water	Alsop and Wood 2011
<i>Oryzias melastigma</i>	96 h LC50	7.30	CuSO ₄ *5H ₂ O	water	Bao et al. 2011
<i>Synechogobius hasta</i>	96 h LC50	0.77	CuSO ₄	water	Chen et al. 2010
<i>Capoeta fusca</i>	96 h LC50	1.10	CuCl ₂	water	Ebrahimpour et al. 2010
<i>Capoeta fusca</i>	96 h LC50	5.40	CuCl ₂	water	Ebrahimpour et al. 2010
<i>Capoeta fusca</i>	96 h LC50	7.50	CuCl ₂	water	Ebrahimpour et al. 2010
<i>Pimephales promelas</i>	96 h LC50	0.28	CuSO ₄	water	Lewis and Keller 2009
<i>Solea senegalensis</i>	96 h LC50	0.32	CuSO ₄ *5H ₂ O	water	Oliva et al. 2009

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Poecilia reticulata</i>	24 h LC50	1.17	CuSO ₄ *5H ₂ O	water	Park and Heo 2009
<i>Oncorhynchus mykiss</i>	96 h LC50	0.21	Cu(NO ₃) ₂ *2H ₂ O	water	De Boeck et al. 2004
<i>Cyprinus carpio</i>	96 h LC50	0.66	Cu(NO ₃) ₂ *2H ₂ O	water	De Boeck et al. 2004
<i>Carassius auratus gibelio</i>	96 h LC50	1.40	Cu(NO ₃) ₂ *2H ₂ O	water	De Boeck et al. 2004
<i>Danio rerio</i>	72 h LC50	0.13	CuSO ₄	water	de Oliveira-Filho et al. 2004
<i>Danio rerio</i>	96 h LC50	0.09	CuSO ₄	water	de Oliveira-Filho et al. 2004
Nematode: Median (min-max) L(E)C50		19.4 (1.27-101)			
<i>Caenorhabditis elegans</i>	48 h EC50	20.3	CuCl ₂ *2H ₂ O	mineral medium	Harrington et al. 2012
<i>Caenorhabditis elegans</i>	24 h EC50	101	CuSO ₄	complex medium	Calafato et al. 2008
<i>Caenorhabditis elegans</i>	48 h EC50	29.9	CuSO ₄	complex medium	Calafato et al. 2008
<i>Caenorhabditis elegans</i>	72 h EC50	18.4	CuSO ₄	complex medium	Calafato et al. 2008
<i>Caenorhabditis elegans</i>	96 h EC50	1.27	CuSO ₄	complex medium	Calafato et al. 2008
<i>Caenorhabditis elegans</i>	120 h EC50	11.7	CuCl ₂ *2H ₂ O	complex medium	Jonker et al. 2004
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		32 (0.4-640)			
<i>Pseudomonas aeruginosa</i> (2 different isolates)	MIC	64-640	CuSO ₄	complex medium	Shakibaie and Harati 2004

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Legionella pneumophila</i>	12 h MIC	0.4	CuCl ₂	not specified	Lin et al. 1996
<i>Pseudomonas aeruginosa</i>	56 h MIC	128	CuSO ₄	complex medium	Teitzel and Parsek 2003
<i>Staphylococcus aureus</i>	24 h MIC	12.8	CuSO ₄	complex medium	Baker et al. 2010
<i>Escherichia coli</i>	16 h MIC	83.2	CuSO ₄	complex medium	Grass and Rensing 2001
<i>Pseudomonas putida</i>	MIC	192	not specified	not specified	Chen et al. 2006
<i>Bacillus circulans</i>	72 h MIC	160	CuSO ₄	complex medium	Yilmaz 2003
<i>Streptomyces sp.</i>	MIC	32	Cu(NO ₃) ₂	mineral medium	Majzlik et al. 2011
<i>Staphylococcus aureus</i> (2 different isolates)	20 h MIC	12.5	Cu(NO ₃) ₂	complex medium	Zhao et al. 1998
<i>Escherichia coli</i>	20 h MIC	25	Cu(NO ₃) ₂	complex medium	Zhao et al. 1998
<i>Pseudomonas aeruginosa</i>	20 h MIC	12.5	Cu(NO ₃) ₂	complex medium	Zhao et al. 1998
Yeast/fungi: Median (min-max) L(E)C50		11.1 (0.82-516)			
<i>Saccharomyces cerevisiae</i>	8 h EC50	11.4	CuSO ₄	complex medium	Kasemets et al. 2009
<i>Saccharomyces cerevisiae</i>	24 h EC50	10.9	CuSO ₄	complex medium	Kasemets et al. 2009
<i>Saccharomyces cerevisiae</i>	24 h IC50	516	CuSO ₄	complex medium	Kasemets et al. 2013
<i>Saccharomyces cerevisiae</i>	24 h IC50	0.82	CuSO ₄	water	Kasemets et al. 2013

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
Mammalian cells in vitro: Median (min-max) L(E)C50		53 (5.6-80)			
primary human endometrial epithelial cells	24 h TC50	10.8	CuCl ₂	complex medium	Wu et al. 2012
L929 cells	24 h IC50	46	CuCl ₂	complex medium	Cao et al. 2012
A549	24 h IC50	>40	CuCl ₂	complex medium	Karlsson et al. 2008
HGF cells	24 h IC50	71	CuCl ₂	complex medium	Contreras et al. 2010
MC3T3-E1	24 h IC50	71	CuCl ₂	complex medium	Contreras et al. 2010
Bel-7402 cells	24 h IC50	5.6	CuCl ₂	complex medium	Zhang et al. 2007
HK-2 cells	24 h IC50	24	CuCl ₂	complex medium	Zhang et al. 2007
HepG2 cells	24 h IC50	80	CuCl ₂	complex medium	Piret et al. 2012a
Caco-2 cells	24 h IC50	60	CuCl ₂	complex medium	Piret et al. 2012b
MDA-MB468 cells	48 h IC50	63.5	CuCl ₂	complex medium	Panjehpour et al. 2010
Vibrio fischeri: Median (min-max) L(E)C50		0.8 (0.2-0.8)			
<i>Vibrio fischeri</i>	30 min EC50	0.6	CuSO ₄	mineral medium	Heinlaan et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	0.8	CuSO ₄	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	0.8	CuSO ₄	mineral medium	Mortimer et al. 2008

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Vibrio fischeri</i>	15 min EC50	0.78	not specified	mineral medium	McCloskey et al. 1996
<i>Vibrio fischeri</i>	30 min EC50	0.30	CuSO ₄	mineral medium	Kurvet et al. 2011
<i>Vibrio fischeri</i>	15 min EC50	0.8	Cu(NO ₃) ₂	mineral medium	Hsieh et al. 2004
<i>Vibrio fischeri</i>	15 min EC50	0.20	CuCl ₂ *2H ₂ O	mineral medium	Kungolos et al. 2006
Protozoa: Median (min-max) L(E)C50		0.43 (0.010-1.70)			
<i>Paramecium multimicronucleatum</i>	48 h LC50	0.020	CuCl ₂	mineral medium	Li et al. 2012a
<i>Tetrahymena thermophila</i>	24 h EC50	1.60	CuSO ₄	mineral medium	Mortimer et al. 2011
<i>Tetrahymena thermophila</i>	4 h EC50	1.10	CuSO ₄	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	4 h EC50	1.10	CuSO ₄	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	1.40	CuSO ₄	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	1.70	CuSO ₄	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	0.40	CuSO ₄	complex medium	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	0.46	CuSO ₄	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	0.27	CuSO ₄	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h LC50	0.47	CuSO ₄ *5H ₂ O	mineral medium	Gallego et al. 2007

Species	Exposure time and endpoint	EC50 on metal basis, mg Cu/L	Tested as (compound)	Test medium*	Reference
<i>Dexiotricha granulosa</i>	24 h LC50	0.17	CuCl ₂ *2H ₂ O	mineral medium	Madoni and Romeo 2006
<i>Colpidium colpoda</i>	24 h LC50	0.050	CuCl ₂ *2H ₂ O	mineral medium	Madoni and Romeo 2006
<i>Euplotes aediculatus</i>	24 h LC50	0.010	CuCl ₂ *2H ₂ O	mineral medium	Madoni and Romeo 2006
<i>Halteria grandinella</i>	24 h LC50	0.010	CuCl ₂ *2H ₂ O	mineral medium	Madoni and Romeo 2006

*Mineral medium - without organic components; Complex medium - with organic components

Table S7 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **ZnO nanoparticles** for various organisms

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		2.3 (0.62-22)	1.9 (0.5-17.7)			
<i>Daphnia magna</i>	48 h LC50	8.8	7.1	50-70 nm (provider), uncoated	mineral medium	Heinlaan et al. 2008
<i>Daphnia magna</i>	48 h EC50	0.6	0.5	20 nm (provider), uncoated	mineral medium	Zhu et al. 2009
<i>Daphnia magna</i>	48 h LC50	1.5	1.2	20 nm (provider), uncoated	mineral medium	Zhu et al. 2009
<i>Daphnia magna</i>	48 h EC50	3.2	2.6	50-70 nm (provider), uncoated	mineral medium	Blinova et al. 2010
<i>Daphnia magna</i>	48 h EC50	2.6	2.1	<100 nm (provider), uncoated	mineral medium	Sovova et al. 2009
<i>Daphnia magna</i>	48 h, LC50	1.5	1.2	40 nm (provider), uncoated	mineral medium	Zhao et al. 2012
<i>Daphnia magna</i>	48 h EC50	0.8	0.6	40 nm (provider), uncoated	mineral medium	Zhao et al. 2012
<i>Daphnia magna</i>	24 h LC50	22.0	17.7	20 nm (provider), uncoated	mineral medium	Poynton et al. 2011
<i>Daphnia magna</i>	24 h EC50	2.6	2.1	50-70 nm (provider), uncoated	mineral medium	Naddafi et al. 2011
<i>Daphnia magna</i>	48 h EC50	2.1	1.7	50-70 nm (provider), uncoated	mineral medium	Naddafi et al. 2011
Algae: Median (min-max) L(E)C50		0.085 (0.052-4.56)	0.068 (0.042-3.66)			
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.052	0.042	50-70 nm (provider), uncoated	mineral medium	Aruoja et al. 2009

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Pseudokirchneriella subcapitata</i>	72 h IC50	0.085	0.068	30 nm (provider), uncoated	complex medium	Franklin et al. 2007
<i>Pseudokirchneriella subcapitata</i>	72 h IC50	0.061	0.049	30 nm (provider), dispersed with Teric N30 (nonylphenol ethoxylate)	complex medium	Franklin et al. 2007
<i>Thalassiosira pseudonana</i>	96 h IC50	4.56	3.66	26.2 ± 5.1 nm (TEM), uncoated	mineral medium	Wong et al. 2010
<i>Skeletonema costatum</i>	96 h IC50	2.36	1.90	26.2 ± 5.1 nm (TEM), uncoated	mineral medium	Wong et al. 2010
Fish: Median (min-max) L(E)C50		3.02 (1.79-4.92)	2.42 (1.44-3.95)			
<i>Danio rerio</i>	96 h LC50	4.92	3.95	30 nm	water	Xiong et al. 2011
<i>Danio rerio</i>	96 h LC50	3.97	3.19	30 nm	water	Yu et al. 2011
<i>Danio rerio</i>	96 h LC50	1.79	1.44	20 nm	water	Zhu et al. 2008
<i>Danio rerio</i>	84 h EC50	2.07	1.66	20 nm	water	Zhu et al. 2008
Nematode: Median (min-max) L(E)C50		39.2 (2.20-982)	31.5 (1.77-789)			
<i>Caenorhabditis elegans</i>	24 h LC50	21.2	17.0	40-100 nm	mineral medium	Ma et al. 2011
<i>Caenorhabditis elegans</i>	4 h EC50	982	789	2-6 nm	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	24 h LC50	790	635	2-6 nm	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	72 h EC50	57.3	46.0	2-6 nm	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	24 h LC50	2.20	1.77	20 nm	water	Wang et al. 2009

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Caenorhabditis elegans</i>	24 h LC50	4.90	3.94	20 nm	mineral medium	Wang et al. 2009
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		500 (50-1000)	400 (40-800)			
<i>Campylobacter jejuni</i>	16 h MIC	50	40	30 nm (TEM)	complex medium	Xie et al. 2011
<i>Escherichia coli</i>	16 h MIC	800	640	30 nm (TEM)	complex medium	Xie et al. 2011
<i>Escherichia coli</i>	16-20 h MIC	400	320	47 nm (TEM)	complex medium	Padmavathy and Vijayaraghavan 2008
<i>Staphylococcus aureus</i>	24 h MIC	500	400	3 nm (TEM)	complex medium	Emami-Karvani and Chehrizi 2011
<i>Escherichia coli</i>	24 h MIC	1000	800	3 nm (TEM)	complex medium	Emami-Karvani and Chehrizi 2011
<i>Campylobacter jejuni</i>	24 h MIC	50	40	30 nm (TEM)	complex medium	Xie et al. 2011
<i>Prevotella intermedia</i>	48 h MIC	1000	800	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Porphyromonas gingivalis</i>	48 h MIC	250	200	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Fusobacterium nucleatum</i>	48 h MIC	250	200	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Aggregatibacter actinomycetemcomitans</i>	48 h MIC	250	200	20 nm (TEM)	complex medium	Vargas-Reus et al. 2012
<i>Staphylococcus aureus</i>	Overnight MIC	1000	800	19 nm (TEM)	complex medium	Ansari et al. 2012
<i>Escherichia coli</i>	Overnight MIC	500	400	19 nm (TEM)	complex medium	Ansari et al. 2012
<i>Klebsiella pneumoniae</i>	Overnight MIC	500	400	19 nm (TEM)	complex medium	Ansari et al. 2012

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Escherichia coli</i>	Overnight MIC	972	777	70 nm (TEM)	complex medium	Liu et al. 2009
<i>Streptococcus mutants</i>	24 h MIC	500	400	>50 nm (TEM)	not specified	Hernández-Sierra et al. 2008
Yeast/fungi: Median (min-max) L(E)C50		121 (40.0-488)	97.2 (32.1-392)			
<i>Aspergillus niger</i>	24 h EC50	50.0	40.2	30-40 nm	complex medium	Patra et al. 2012
<i>Fusarium oxysporum</i>	120 h EC50	50.0	40.2	30-40 nm	complex medium	Patra et al. 2012
<i>Penicillium expansum</i>	12 d EC50	244	196	70 nm	complex medium	He et al. 2011
<i>Botrytis cinerea</i>	12 d EC50	488	392	70 nm	complex medium	He et al. 2011
<i>Candida albicans</i>	24 h EC50	40.0	32.1	6.8 -11.6 nm	mineral medium	Lipovsky et al. 2011
<i>Saccharomyces cerevisiae</i>	8 h EC50	121	97.2	50-70nm (provider)	complex medium	Kasemets et al. 2009
<i>Saccharomyces cerevisiae</i>	24 h EC50	131	105	50-70 nm (provider)	complex medium	Kasemets et al. 2009
Mammalian cells in vitro: Median (min-max) L(E)C50		43.4 (4.5-75)	34.9 (3.6-60.2)			
HeLa cells	72 h IC50	45.7	36.7	100 nm (TEM)	not specified	Li et al. 2012b
A549 cells	24 h IC50	52	41.8	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	24 h IC50	54	43.4	<50 nm (TEM)	complex medium	Seiffert et al. 2012
A549 cells	24 h IC50	45	36.1	<50 nm (TEM)	complex medium	Seiffert et al. 2012

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
A549 cells	24 h IC50	56	45	<50 nm (TEM)	complex medium	Seiffert et al. 2012
CCL-131 cells	24 h IC50	75	60.2	50-70 nm (provider)	complex medium	Jeng and Swanson 2006
CCL-131 cells	24 h IC50	13.5	10.8	50-70 nm (provider)	complex medium	Jeng and Swanson 2006
skin fibroblasts	24 h IC50	49	39.4	50-70 nm (provider)	complex medium	Dechsakulthorn et al. 2007
cultured rat alveolar epithelial cell monolayers	24 h IC50	13.7	17	20 nm (provider)	serum-free	Kim et al. 2010
cultured rat alveolar epithelial cell monolayers	24 h IC50	22.5	28	20 nm (provider)	complex medium	Kim et al. 2010
Ana-1 cells	24 h IC50	44.6	35.8	342 nm (TEM)	complex medium	Song et al. 2010
Ana-1 cells	24 h IC50	44.0	35.3	108 nm (TEM)	complex medium	Song et al. 2010
Ana-1 cells	24 h IC50	40.3	32.4	70.89 nm (TEM)	complex medium	Song et al. 2010
Ana-1 cells	24 h IC50	30.9	24.8	19 nm (TEM)	complex medium	Song et al. 2010
RAW 264.7 cells	24 h IC50	21.7	17.4	22.6 ± 5.1 nm (TEM)	complex medium	Rallo et al. 2012
RAW 264.7 cells	24 h IC50	10.1	8.1	22.6 ± 5.1 nm (TEM)	complex medium	Rallo et al. 2012
BEAS-2B cells	24 h IC50	6.5	5.2	22.6 ± 5.1 nm (TEM)	complex medium	Rallo et al. 2012
BEAS-2B cells	24 h IC50	4.5	3.6	22.6 ± 5.1 nm (TEM)	complex medium	Rallo et al. 2012
MH-S cells	24 h IC50	48.5	39.0	342 nm (TEM)	complex medium	Zhang et al. 2012b

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
MH-S cells	24 h IC50	47.4	38.0	108 nm (TEM)	complex medium	Zhang et al. 2012b
MH-S cells	24 h IC50	45.4	36.5	71 nm (TEM)	complex medium	Zhang et al. 2012b
MH-S cells	24 h IC50	26.7	21.5	19 nm (TEM)	complex medium	Zhang et al. 2012b
A549 cells	24 h IC50	13.6	10.9	70 ± 13 nm (TEM)	complex medium	Lin et al. 2009
A549 cells	24 h IC50	14.2	11.4	420 ± 269 nm (TEM)	complex medium	Lin et al. 2009
HEK293 cells	24 h IC50	21.47	17.2	n.a.	complex medium	Dua et al. 2011
<i>Vibrio fischeri</i> : Median (min-max) L(E)C50		4.3 (1.8-7.3)	3.5 (1.4-5.8)			
<i>Vibrio fischeri</i>	30 min EC50	1.8	1.4	50-70 nm (provider), uncoated	mineral medium	Heinlaan et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	4.8	3.9	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	3.8	3.0	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	7.3	5.8	<100 nm (provider), uncoated	mineral medium	Sovova et al. 2009
Protozoa : Median (min-max) L(E)C50		11.70 (5.35-573.8)	9.40 (4.30-461.0)			
<i>Paramecium multimicronucleatum</i>	48 h LC50	573.8	461.0	9.8 ± 5.0 nm (TEM), uncoated	mineral medium	Li et al. 2012a
<i>Tetrahymena thermophila</i>	4 h EC50	5.35	4.30	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	4 h EC50	6.22	5.00	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2010

Species	Exposure time and endpoint	EC50 on compound basis, mg ZnO/L	EC50 on metal basis, mg Zn/L	Primary size (method used for determination), coating	Test medium*	Reference
<i>Tetrahymena thermophila</i>	24 h EC50	8.46	6.80	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	10.3	8.30	50-70 nm (provider), uncoated	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	11.7	9.40	50-70 nm (provider), uncoated	complex medium	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	20.4	16.4	50-70 nm (provider), uncoated	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	15.4	12.4	50-70 nm (provider), uncoated	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	33.0	26.5	50-70 nm (provider), uncoated	natural water	Blinova et al. 2010

n.a.-not analysed; TEM – transmission electron microscopy

*Mineral medium - without organic components; Complex medium - with organic components

Table S8 The L(E)C50 values from individual studies and median L(E)C50 values (minimum-maximum value) for **zinc salt** for various organisms

Species	Exposure time and endpoint	EC50 on metal basis, mg Zn/L	Tested as (compound)	Test medium*	Reference
Crustaceans: Median (min-max) L(E)C50		1.3 (0.4-1.8)			
<i>Daphnia magna</i>	48 h LC50	1.38	ZnSO ₄ *7H ₂ O	mineral medium	Heinlaan et al. 2008
<i>Daphnia magna</i>	48 h EC50	1.40	ZnSO ₄ *7H ₂ O	mineral medium	Blinova et al. 2010
<i>Daphnia magna</i>	24 h LC50	1.28	ZnSO ₄ *7H ₂ O	mineral medium	Poynton et al. 2011
<i>Daphnia magna</i>	48 h LC50	0.87	ZnCl ₂	mineral medium	Wang and Guan 2010
<i>Daphnia magna</i>	24 h LC50	1.80	ZnSO ₄ *7H ₂ O	mineral medium	Teodorovic et al. 2009
<i>Daphnia magna</i>	48 h LC50	0.41	ZnSO ₄ *7H ₂ O	mineral medium	Teodorovic et al. 2009
Algae: Median (min-max) L(E)C50		0.088 (0.042-3.48)			
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.042	ZnSO ₄ *7H ₂ O	mineral medium	Aruoja et al. 2009
<i>Pseudokirchneriella subcapitata</i>	72 h IC50	0.061	ZnCl ₂	complex medium	Franklin et al. 2007
<i>Thalassiosira pseudonana</i>	96 h IC50	3.48	ZnSO ₄ *7H ₂ O	complex medium	Wong et al. 2010
<i>Skeletonema costatum</i>	96 h IC50	1.23	ZnSO ₄ *7H ₂ O	complex medium	Wong et al. 2010
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.056	ZnSO ₄ *7H ₂ O	complex medium	Graff et al. 2003
<i>Pseudokirchneriella subcapitata</i>	72 h EC50	0.20	ZnSO ₄ *7H ₂ O	natural water	Graff et al. 2003

Species	Exposure time and endpoint	EC50 on metal basis, mg Zn/L	Tested as (compound)	Test medium*	Reference
<i>Chlorella sp.</i>	48 h EC50	0.085	ZnSO ₄ *7H ₂ O	mineral medium	Franklin et al. 2002
<i>Chlorella sp.</i>	72 h EC50	0.092	ZnSO ₄ *7H ₂ O	mineral medium	Franklin et al. 2002
Fish: Median (min-max) L(E)C50		7.48 (2.54-8.06)			
<i>Danio rerio</i>	96 h LC50	8.06	ZnSO ₄ *7H ₂ O	water	Xiong et al. 2011
<i>Danio rerio</i>	96 h LC50	7.48	ZnSO ₄ *7H ₂ O	water	Yu et al. 2011
<i>Danio rerio</i>	96 h LC50	2.54	ZnSO ₄ *7H ₂ O	water	Alsop and Wood 2011
Nematode: Median (min-max) L(E)C50		49.1 (1.39-884)			
<i>Caenorhabditis elegans</i>	48 h EC50	39.2	ZnCl ₂	mineral medium	Harrington et al. 2012
<i>Caenorhabditis elegans</i>	7 d EC50	10.5	ZnCl ₂	mineral medium	Murphy et al. 2011
<i>Caenorhabditis elegans</i>	24 h LC50	884	ZnCl ₂	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	4 h EC50	546	ZnCl ₂	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	72 h EC50	59.0	ZnCl ₂	mineral medium	Ma et al. 2009
<i>Caenorhabditis elegans</i>	24 h LC50	1.39	ZnCl ₂	water	Wang et al. 2009
Bacteria: Median (min-max) minimal inhibitory concentration (MIC)		30 (12.5-1430)			
<i>Pseudomonas aeruginosa</i>	56 h MIC	520	ZnSO ₄	complex medium	Teitzel and Parsek 2003

Species	Exposure time and endpoint	EC50 on metal basis, mg Zn/L	Tested as (compound)	Test medium*	Reference
<i>Pseudomonas putida</i>	MIC	325	not reported	not reported	Chen et al. 2006
<i>Bacillus circulans</i>	72 h MIC	1430	ZnCl ₂	complex medium	Yilmaz 2003
<i>Helicobacter pylori</i>	16 h MIC	100	ZnCl ₂	complex medium	Mobley et al. 1999
<i>Streptomyces sp.</i>	MIC	30	ZnCl ₂	mineral medium	Majzlik et al. 2011
<i>Staphylococcus aureus</i> (2 different isolates)	20 h MIC	12.5	Zn(NO ₃) ₂	complex medium	Zhao et al. 1998
<i>Escherichia coli</i>	20 h MIC	25	Zn(NO ₃) ₂	complex medium	Zhao et al. 1998
<i>Pseudomonas aeruginosa</i>	20 h MIC	12.5	Zn(NO ₃) ₂	complex medium	Zhao et al. 1998
Yeast/fungi: Median (min-max) L(E)C50		78.2 (75.3-81.2)			
<i>Saccharomyces cerevisiae</i>	8 h EC50	75.3	ZnSO ₄ *7H ₂ O	complex medium	Kasemets et al. 2009
<i>Saccharomyces cerevisiae</i>	24 h EC50	81.2	ZnSO ₄ *7H ₂ O	complex medium	Kasemets et al. 2009
Mammalian cells in vitro: Median (min-max) L(E)C50		9.8 (5.9-15.7)			
Ana-1 cells	24 h IC50	13.33	ZnCl ₂	complex medium	Song et al. 2010
primary human endometrial epithelial cells	24 h TC50	14.1	ZnCl ₂	complex medium	Wu et al. 2012
PC12 cells	24 h IC50	8.2	ZnSO ₄ *7H ₂ O	first 2 h without serum	Pavlica et al. 2009
PC12 cells	24 h IC50	9.5	ZnSO ₄ *7H ₂ O	first 2 h without serum	Pavlica et al. 2009

Species	Exposure time and endpoint	EC50 on metal basis, mg Zn/L	Tested as (compound)	Test medium*	Reference
PC12 cells	24 h IC50	5.9	ZnSO ₄ *7H ₂ O	first 2 h without serum	Pavlica et al. 2009
PC12 cells	24 h IC50	12.1	ZnSO ₄	first 2 h without serum	Pavlica et al. 2009
PC12 cells	24 h IC50	15.7	ZnSO ₄	first 2 h without serum	Pavlica et al. 2009
PC12 cells	24 h IC50	12.1	ZnSO ₄	first 2 h without serum	Pavlica et al. 2009
B16 mouse melanoma cells	24 h IC50	8.2	ZnCl ₂	complex medium	Borovanský and Riley 1989
HeLa cells	24 h IC50	9.8	ZnCl ₂	complex medium	Borovanský and Riley 1989
I-221 epithelial cells	24 h IC50	9.8	ZnCl ₂	complex medium	Borovanský and Riley 1989
<i>Vibrio fischeri</i> : Median (min-max) L(E)C50		3.2 (0.25-9.3)			
<i>Vibrio fischeri</i>	30 min EC50	0.25	ZnSO ₄ *7H ₂ O	mineral medium	Heinlaan et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	3.8	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	30 min EC50	3.2	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2008
<i>Vibrio fischeri</i>	15 min EC50	2.28	not specified	mineral medium	McCloskey et al. 1996
<i>Vibrio fischeri</i>	30 min EC50	4.64	ZnSO ₄ *7H ₂ O	mineral medium	Teodorovic et al. 2009
<i>Vibrio fischeri</i>	30 min EC50	9.27	ZnSO ₄ *7H ₂ O	mineral medium	Kurvet et al. 2011
<i>Vibrio fischeri</i>	15 min EC50	1.49	ZnCl ₂	mineral medium	Kungolos et al., 2006

Species	Exposure time and endpoint	EC50 on metal basis, mg Zn/L	Tested as (compound)	Test medium*	Reference
Protozoa: Median (min-max) L(E)C50:		7.00 (3.58-175.2)			
<i>Paramecium multimicronucleatum</i>	48 h LC50	175.2	ZnCl ₂	mineral medium	Li et al. 2012a
<i>Tetrahymena thermophila</i>	4 h EC50	4.50	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	4 h EC50	5.20	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	6.70	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	7.00	ZnSO ₄ *7H ₂ O	mineral medium	Mortimer et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	7.10	ZnSO ₄ *7H ₂ O	complex medium	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	21.1	ZnSO ₄ *7H ₂ O	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h EC50	18.6	ZnSO ₄ *7H ₂ O	natural water	Blinova et al. 2010
<i>Tetrahymena thermophila</i>	24 h LC50	3.58	ZnSO ₄ *7H ₂ O	mineral medium	Gallego et al. 2007

*Mineral medium - without organic components; Complex medium - with organic components

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