

Guest editorial:

GEBEL-CRITERIA FOR RISK ASSESSMENT IN NANOTOXICOLOGY

Seddik Hammad^{1*}, Ahmed M. Abdou¹, Mosaab A. Omar²

¹ Department of Forensic Medicine and Veterinary Toxicology, Faculty of Veterinary Medicine, South Valley University, 83523 Qena/Egypt

² Department of Parasitology, Faculty of Veterinary Medicine, South Valley University, 83523 Qena/Egypt

* Corresponding author: Seddik Hammad. E-mail: seddik.hammad@vet.svu.edu.eg

Risk assessment of nanomaterials represents one of the cutting-edge topics in toxicology (Zhao et al., 2014; Lucafo et al., 2013; Fadeel et al., 2013; Kroll et al., 2012; Kim et al., 2012; Hadrup et al., 2012; Lange and Obendorf, 2012; Xu et al., 2013). Numerous nanomaterial containing consumer products have already been introduced on the market, including textiles, sunscreens, paints, car tyres and electronics (Bolt et al., 2012; Kumar and Dhawan, 2013; Schluesener and Schluesener, 2013; Marchan, 2012). The number of manuscripts published in this field is high, with more 500 publications focusing on nanosafety-associated topics per year (Bolt et al., 2012; Schäfer et al., 2013; Horie et al., 2013; Klein et al., 2012; Hoelting et al., 2013; Silva et al., 2014). It has become clear that a detailed physical and chemical characterization of nanomaterials is required for risk evaluation (Park et al., 2013; Xiong et al., 2013; Zhao et al., 2013; Couto et al., 2014). Among the current challenges are the methodological requirements in exposure monitoring (Babič et al., 2014; Lainé et al., 2014; Su et al., 2014; Bruchet et al., 2013). Because of the enormous variability and the rapid development of novel materials it has become difficult for regulators to keep pace and maintain overview. In this complex situation the Advisory Board of the German Society of Toxicology introduced the Gebel-criteria, a novel concept of risk assessment in nanotoxicology (Gebel et al., 2014). Accord-

ing to this concept, a first step in evaluating novel nanomaterials should be to check whether they belong to one of the three following categories: **Category 1**: Nanoparticles for which toxicity is mediated by the specific chemical properties of its components, such as relaxed ions. Nanomaterials belonging to this category must be evaluated on a case-by-case basis. **Category 2** are rigid biopersistent respirable fibrous nanomaterials. They may cause lung cancer and mesotheliomas, if they show a high aspect ratio. In this case they will act similarly as carcinogenic asbestos fibres. **Category 3** are respirable granular biodurable particles. After inhalation they may cause inflammation and finally lung cancer. It should be considered that nanomaterials of categories 2 and 3 are of relevance only after inhalation (Gebel et al., 2014). Considering the complex situation in current nanotoxicology the introduction of the three ‘Gebel-criteria’ will facilitate risk assessment in future.

REFERENCES

- Babič M, Schmiedtová M, Poledne R, Herynek V, Horák D. In vivo monitoring of rat macrophages labeled with poly(l-lysine)-iron oxide nanoparticles. J Biomed Mater Res B Appl Biomater. 2014 Oct 6. doi: 10.1002/jbm.b.33292. [Epub ahead of print]
- Bolt HM, Marchan R, Hengstler JG. Nanotoxicology and oxidative stress control: cutting-edge topics in toxicology. Arch Toxicol. 2012;86:1629-35.

- Bruchet A, Charles P, Janex Habibi ML, Glucina K. Monitoring and treatment of selected nanoparticles. *Water Sci Technol.* 2013;68:1454-60.
- Couto D, Freitas M, Vilas-Boas V, Dias I, Porto G, Lopez-Quintela MA, et al. Interaction of polyacrylic acid coated and non-coated iron oxide nanoparticles with human neutrophils. *Toxicol Lett.* 2014;225:57-65.
- Fadeel B, Alenius H, Savolainen K. *Nanotoxicology. Toxicology.* 2013;313:1-2.
- Gebel T, Foth H, Damm G, Freyberger A, Kramer PJ, Lilienblum W, et al. Manufactured nanomaterials: categorization and approaches to hazard assessment. *Arch Toxicol.* 2014 Oct 19. [Epub ahead of print]
- Hadrup N, Loeschner K, Bergström A, Wilcks A, Gao X, Vogel U, et al. Subacute oral toxicity investigation of nanoparticulate and ionic silver in rats. *Arch Toxicol.* 2012;86:543-51.
- Hoelting L, Scheinhardt B, Bondarenko O, Schildknecht S, Kapitzka M, Tanavde V, et al. A 3-dimensional human embryonic stem cell (hESC)-derived model to detect developmental neurotoxicity of nanoparticles. *Arch Toxicol.* 2013;87:721-33.
- Horie M, Kato H, Iwahashi H. Cellular effects of manufactured nanoparticles: effect of adsorption ability of nanoparticles. *Arch Toxicol.* 2013;87:771-81.
- Kim JE, Shin JY, Cho MH. Magnetic nanoparticles: an update of application for drug delivery and possible toxic effects. *Arch Toxicol.* 2012;86:685-700.
- Klein CL, Wiench K, Wiemann M, Ma-Hock L, van Ravenzwaay B, Landsiedel R. Hazard identification of inhaled nanomaterials: making use of short-term inhalation studies. *Arch Toxicol.* 2012;86:1137-51.
- Kroll A, Pillukat MH, Hahn D, Schneckeburger J. Interference of engineered nanoparticles with in vitro toxicity assays. *Arch Toxicol.* 2012;86:1123-36.
- Kumar A, Dhawan A. Genotoxic and carcinogenic potential of engineered nanoparticles: an update. *Arch Toxicol.* 2013;87:1883-900.
- Lainé AL, Gravier J, Henry M, Sancey L, Béjaud J, Pancani E, et al. Conventional versus stealth lipid nanoparticles: formulation and in vivo fate prediction through FRET monitoring. *J Control Release.* 2014;188:1-8.
- Lange LE, Obendorf SK. Effect of plasma etching on destructive adsorption properties of polypropylene fibers containing magnesium oxide nanoparticles. *Arch Environ Contam Toxicol.* 2012;62:185-94.
- Lucafò M, Gerdol M, Pallavicini A, Pacor S, Zorzet S, Da Ros T, et al. Profiling the molecular mechanism of fullerene cytotoxicity on tumor cells by RNA-seq. *Toxicology* 2013;314:183-92.
- Marchan R. A special issue on nanotoxicology. *EXCLI J* 2012;11:176-7.
- Park EJ, Shim HW, Lee GH, Kim JH, Kim DW. Comparison of toxicity between the different-type TiO₂ nanowires in vivo and in vitro. *Arch Toxicol.* 2013;87:1219-30.
- Schäfer B, Brocke JV, Epp A, Götz M, Herzberg F, Kneuer C, et al. State of the art in human risk assessment of silver compounds in consumer products: a conference report on silver and nanosilver held at the BfR in 2012. *Arch Toxicol.* 2013;87:2249-62.
- Schluesener JK, Schluesener HJ. Nanosilver: application and novel aspects of toxicology. *Arch Toxicol.* 2013;87:569-76.
- Silva T, Pokhrel LR, Dubey B, Tolaymat TM, Maier KJ, Liu X. Particle size, surface charge and concentration dependent ecotoxicity of three organo-coated silver nanoparticles: comparison between general linear model-predicted and observed toxicity. *Sci Total Environ.* 2014;468-469:968-76.
- Su CK, Hung CW, Sun YC. In vivo measurement of extravasation of silver nanoparticles into liver extracellular space by push-pull-based continuous monitoring system. *Toxicol Lett.* 2014;227:84-90.
- Xiong S, George S, Yu H, Damoiseaux R, France B, Ng KW, et al. Size influences the cytotoxicity of poly (lactic-co-glycolic acid) (PLGA) and titanium dioxide (TiO₂) nanoparticles. *Arch Toxicol.* 2013;87:1075-86.
- Xu J, Li Z, Xu P, Xiao L, Yang Z. Nanosized copper oxide induces apoptosis through oxidative stress in podocytes. *Arch Toxicol.* 2013;87:1067-73.
- Zhao B, Sun L, Zhang W, Wang Y, Zhu J, Zhu X, et al. Secretion of intestinal goblet cells: a novel excretion pathway of nanoparticles. *Nanomedicine* 2014;10:839-49.
- Zhao X, Ng S, Heng BC, Guo J, Ma L, Tan TT, et al. Cytotoxicity of hydroxyapatite nanoparticles is shape and cell dependent. *Arch Toxicol.* 2013;87:1037-52.