

Editorial:

OXIDATIVE STRESS RESEARCH

R. Marchan

Leibniz Research Centre for Working Environment and Human Factors (*IfAdo*),
Ardeystrasse 67, 44139 Dortmund, Germany

E-mail: marchan@ifado.de, Telephone: +49 231-1084-213, Fax: +49 231-1084-403

Oxidative stress remains one of the most intensively studied and most frequently cited fields in toxicology. A perfect example is the conclusive review article by Cederbaum et al. (2009) illustrating how ethanol leads to increased generation of reactive oxygen species and depletion of antioxidants, key mechanisms that may finally lead to liver cirrhosis. Besides ethanol, several other chemicals were studied with regard to their ability to generate reactive oxygen species, including uranium, arsenic, aluminium perfluorooctanesulfonamides, benzimidazole anthelmics and the non-genotoxic liver carcinogen piperonyl butoxide. A second cutting-edge topic in the field of oxidative stress centres on anti-oxidative compounds, which may exert anti-carcinogenic activities. A frequently-cited example is the review article of Yang et al. (2009) on the activities of polyphenolic compounds. The authors attempt to differentiate “fact from fancy” concerning the hypothesis that drinking tea can protect from cancer.

Table 1: Oxidative stress research

Key message	Reference
Menadione is used as a vitamin K source in animal feed. However, menadione, in concentrations as low as 1.0 µM, decreased the proportion of blastocysts using a reactive oxygen species-dependent mechanism. Insulin-like growth factor-1 antagonized the effect of menadione, this time using a mechanism independent of ROS generation.	Moss et al., 2009
Aluminium, the most widely distributed metal in the environment, produces oxidative stress and impairs mitochondrial bioenergetics in neurons (review).	Kumar and Gill, 2009
Perfluorooctanesulfonamides are large scale industrial chemicals, but relatively little is known about their toxic mechanisms. This study demonstrates that N-ethyl perfluorooctanesulfonamidoethanol is metabolized to perfluorooctanesulfonate in rats which increased superoxide dismutase activity.	Xie et al., 2009
Arsenic enhances generation of reactive oxygen species and malondialdehyde in a human hepatocyte cell line, which coincided with an unexpected increase of intracellular glutathione.	Wang et al., 2009a
4-n-Nonylphenol augments vascular contractile responsiveness in rats by enhanced activity of vascular oxidant enzymes.	Hsieh et al., 2009
Oxidative damage and nitric oxide depletion by sub-chronic exposure to methylmercury are responsible for hypertension.	Grotto et al., 2009
Chronic ethanol consumption increased reactive oxygen species and depleted antioxidants.	Cederbaum et al., 2009; Hengstler et al., 2009a (editorial)

Table 1 (cont.): Oxidative stress research

Key message	Reference
Propyl gallate is a synthetic antioxidant widely used in processed food and cosmetics. However, in this study propyl gallate produced increased as well as decreased ROS levels depending on the incubation time.	Han et al., 2009
Oxidative DNA damage is repaired less in individuals with the Cys 326 Cys variant of the codon 326 nOGG1 polymorphism.	Ke et al., 2010
The ability to activate antioxidant defence systems determines the susceptibility of individual proximal tubular cells to lead acetate.	Wang et al., 2009b
Thermoluminescence assay quantifies oxidative stress with similar sensitivity as conventional markers, such as thiobarbituric acid reactive substances and malondialdehyde. An advantage is that cell and tissue material can be directly included into the assay without pre-processing or extraction procedures.	Schumann et al., 2009
Myeloperoxidase is a key enzyme in benzene metabolism generating the ultimate genotoxic intermediates.	Westphal et al., 2009
Alpha-tocopherol ameliorates the CCl ₄ -induced decrease in hepatic vitamin C concentration.	Iida et al., 2009
Uranium induces apoptosis in lung epithelial cells by generation of oxidative stress.	Periyakaruppan et al., 2009
Diphenyl ditelluride causes oxidative stress in the cerebral cortex of rats.	Stangherlin et al., 2009
The most sensitive indicators of the peroxidative damage induced by tert-butyl hydroperoxide in rat hepatocytes are changes in intracellular glutathione concentrations and compromised mitochondrial membrane potential.	Cervinková et al., 2009
This review discusses the antioxidative activities of polyphenolic compounds in tea.	Yang et al., 2009 (review); Hengstler et al., 2009b (editorial)
The antioxidative polyphenol, resveratrol shows a protective effect on lipopolysaccharide-induced acute phase response in rats.	Sebai et al., 2009
Administration of the benzimidazole, anthelmintic oxfendazole caused oxidative stress and tumour promotion in rat liver.	Dewa et al., 2009
Piperonyl butoxide induces oxidative stress in the livers of rats only when given above a threshold dose of 0.25 % in the diet of rats.	Muguruma et al., 2009
Mitochondrial uncoupling is a protective mechanism against acute iron overload.	Pardo Andreu et al., 2009
Exogenously administered metallothionein attenuates carmustine-induced oxidative stress and protects against pulmonary fibrosis in rats.	Helal and Helal, 2009

REFERENCES

- Cederbaum AI, Lu Y, Wu D. Role of oxidative stress in alcohol-induced liver injury. *Arch Toxicol* 2009;83:519-48.
- Cervinková Z, Kriváková P, Lábajová A, Rousar T, Lotková H, Kucera O et al. Mechanisms participating in oxidative damage of isolated rat hepatocytes. *Arch Toxicol* 2009;83:363-72.

Dewa Y, Nishimura J, Muguruma M, Jin M, Kawai M, Saegusa Y et al. Involvement of oxidative stress in hepatocellular tumor-promoting activity of oxfendazole in rats. *Arch Toxicol* 2009;83:503-11.

Grotto D, Barcelos GR, Valentini J, Antunes LM, Angeli JP, Garcia SC et al. Low levels of methylmercury induce DNA damage in rats: protective effects of selenium. *Arch Toxicol* 2009;83:249-5.

Han YH, Moon HJ, You BR, Park WH. The anti-apoptotic effects of caspase inhibitors on propyl gallate-treated HeLa cells in relation to reactive oxygen species and glutathione levels. *Arch Toxicol* 2009;83:825-33.

Helal GK, Helal OK. Metallothionein attenuates carmustine-induced oxidative stress and protects against pulmonary fibrosis in rats. *Arch Toxicol* 2009;83:87-94.

Hengstler JG, Godoy P, Stewart JD, Bolt HM. Alcohol-induced liver injury: how a small molecule overwhelms one of the cell types with the best regeneration capacity of the human body. *Arch Toxicol* 2009a;83:513-4.

Hengstler JG, Marchan R, Bolt HM. Can drinking tea prevent cancer? A controversy revisited. *Arch Toxicol* 2009b;83:1-2.

Hsieh CY, Miaw CL, Hsieh CC, Tseng HC, Yang YH, Yen CH. Effects of chronic 4-nonylphenol treatment on aortic vasoconstriction and vasorelaxation in rats. *Arch Toxicol* 2009;83:941-6.

Iida C, Fujii K, Koga E, Washino Y, Kitamura Y, Ichi I et al. Effect of alpha-tocopherol on carbon tetrachloride intoxication in the rat liver. *Arch Toxicol* 2009;83:477-83.

Ke Y, Duan X, Wen F, Xu X, Tao G, Zhou L et al. Association of melamine exposure with urinary stone and oxidative DNA damage in infants. *Arch Toxicol* 2010;84:301-7.

Kumar V, Gill KD. Aluminium neurotoxicity: neurobehavioural and oxidative aspects. *Arch Toxicol* 2009;83:965-78.

Moss JI, Pontes E, Hansen PJ. Insulin-like growth factor-1 protects preimplantation embryos from anti-developmental actions of menadione. *Arch Toxicol* 2009;83:1001-7.

Muguruma M, Kawai M, Dewa Y, Nishimura J, Saegusa Y, Yasuno H et al. Threshold dose of piperonyl butoxide that induces reactive oxygen species-mediated hepatocarcinogenesis in rats. *Arch Toxicol* 2009;83:183-93.

Pardo Andreu GL, Inada NM, Vercesi AE, Curti C. Uncoupling and oxidative stress in liver mitochondria isolated from rats with acute iron overload. *Arch Toxicol* 2009;83:47-53.

Periyakaruppan A, Sarkar S, Ravichandran P, Sadanandan B, Sharma CS, Ramesh V et al. Uranium induces apoptosis in lung epithelial cells. *Arch Toxicol* 2009;83:595-600.

Schumann A, Bauer A, Hermes M, Gilbert M, Hengstler JG, Wilhelm C. A rapid and easy to handle thermoluminescence based technique for evaluation of carbon tetrachloride-induced oxidative stress on rat hepatocytes. *Arch Toxicol* 2009;83:709-20.

Sebai H, Ben-Attia M, Sani M, Aouani E, Ghanem-Boughanmi N. Protective effect of resveratrol in endotoxemia-induced acute phase response in rats. *Arch Toxicol* 2009;83:335-40.

Stangherlin EC, Ardais AP, Rocha JB, Nogueira CW. Exposure to diphenyl ditelluride, via maternal milk, causes oxidative stress in cerebral cortex, hippocampus and striatum of young rats. *Arch Toxicol* 2009; 83:485-91.

Wang Y, Xu Y, Wang H, Xue P, Li X, Li B et al. Arsenic induces mitochondria-dependent apoptosis by reactive oxygen species generation rather than glutathione depletion in Chang human hepatocytes. *Arch Toxicol* 2009a;83:899-908.

Wang L, Wang H, Hu M, Cao J, Chen D, Liu Z. Oxidative stress and apoptotic changes in primary cultures of rat proximal tubular cells exposed to lead. *Arch Toxicol* 2009b;83:417-27.

Westphal GA, Büniger J, Lichey N, Taeger D, Mönnich A, Hallier E. The benzene metabolite para-benzoquinone is genotoxic in human, phorbol-12-acetate-13-myristate induced, peripheral blood mononuclear cells at low concentrations. *Arch Toxicol* 2009;83:721-9.

Xie W, Wu Q, Kania-Korwel I, Tharappel JC, Telu S, Coleman MC et al. Subacute exposure to N-ethyl perfluorooctanesulfonamidoethanol results in the formation of perfluorooctanesulfonate and alters superoxide dismutase activity in female rats. *Arch Toxicol* 2009;83:909-24.

Yang CS, Lambert JD, Sang S. Antioxidative and anti-carcinogenic activities of tea polyphenols. *Arch Toxicol* 2009;83:11-21.