

# Exposure, metabolism, and health effects of arsenic in residents from arsenic-contaminated groundwater areas of Vietnam and Cambodia: A review

Agusa T., Kunito T., Kubota R., Inoue S., Fujihara J., Minh T.B., Ha N.N., Tu N.P.C., Trang P.T.K., Chamnan C., Takeshita H., Iwata H., Tuyen B.C., Viet P.H., Tana T.S., Tanabe S.

Dept of Legal Medicine, Shimane University Faculty of Medicine, humo, Japan; Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan; Dept of Environmental Sciences, Faculty of Science, Shinshu University, Matsumoto, Japan; National Institute of Health Sciences, Setagaya-ku, Tokyo, Japan; Center for Environmental Technol and Sustainable Development, Hanoi University of Science, Hanoi, Viet Nam; Dept of Life Environmental Conservation, Faculty of Agriculture, Ehime University, Matsuyama, Japan; Inland Fisheries Research and Development Institute, Dept of Fisheries, Phnom Penh, Cambodia; Research Institute for BioTechnol and Environment, Nong Lam University, ThuDuc District, Ho Chi Minh City, Viet Nam; Social and Cultural Observation Unit, Office of the Council of Ministers, Phnom Penh, Cambodia

**Abstract:** In this review, we summarize the current knowledge on exposure, metabolism, and health effects of arsenic (As) in residents from As-contaminated groundwater areas of Vietnam and Cambodia based on our findings from 2000 and other studies. The health effects of As in humans include severe gastrointestinal disorders, hepatic and renal failure, cardiovascular disturbances, skin pigmentation, hyperkeratosis, and cancers in the lung, bladder, liver, kidney, and skin. Arsenic contamination in groundwater is widely present at Vietnam and Cambodia and the highest As levels are frequently found in groundwater from Cambodia. Sand filter system can reduce As concentration in raw groundwater. The results of hair and urine analyses indicate that residents from these As-contaminated areas are exposed to As. In general, sex, age, body mass index, and As exposure level are significantly associated with As metabolism. Genetic polymorphisms in arsenic (+III) methyltransferase and glutathione-5-transferase isoforms may be influenced As metabolism and accumulation in a Vietnamese population. It is suggested oxidative DNA damage is caused by exposure to As in groundwater from residents in Cambodia. An epidemiologic study on an association of As exposure with human health effects is required in these areas. © 2010 Freund Publishing House Limited.

**Author Keywords:** Arsenic; Cambodia; Groundwater; Human; Vietnam

Year: 2010

Source title: Reviews on Environmental Health

Volume: 25

Issue: 3

Page : 193-220

Link: Scopus Link

Correspondence Address: Tanabe, S.; Center for Marine Environmental Studies (CMES), Ehime University, Bunkyo-cho 2-5, Matsuyama 790-8577, Japan; email: shinsuke@agr.ehime-u.ac.jp

ISSN: 487554

CODEN: REVHA

Language of Original Document: English

Abbreviated Source Title: Reviews on Environmental Health

Document Type: Review

Source: Scopus

Authors with affiliations:

- Agusa, T., Dept of Legal Medicine, Shimane University Faculty of Medicine, humo, Japan, Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan
- Kunito, T., Dept of Environmental Sciences, Faculty of Science, Shinshu University, Matsumoto, Japan
- Kubota, R., National Institute of Health Sciences, Setagaya-ku, Tokyo, Japan
- Inoue, S., Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan
- Fujihara, J., Dept of Legal Medicine, Shimane University Faculty of Medicine, humo, Japan
- Minh, T.B., Center for Environmental Technol and Sustainable Development, Hanoi University of Science, Hanoi, Viet Nam
- Ha, N.N., Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan
- Tu, N.P.C., Dept of Life Environmental Conservation, Faculty of Agriculture, Ehime University, Matsuyama, Japan
- Trang, P.T.K., Center for Environmental Technol and Sustainable Development, Hanoi University of Science, Hanoi, Viet Nam
- Chamnan, C., Inland Fisheries Research and Development Institute, Dept of Fisheries, Phnom Penh, Cambodia
- Takeshita, H., Dept of Legal Medicine, Shimane University Faculty of Medicine, humo, Japan
- Iwata, H., Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan
- Tuyen, B.C., Research Institute for BioTechnol and Environment, Nong Lam University, ThuDuc District, Ho Chi Minh City, Viet Nam
- Viet, P.H., Center for Environmental Technol and Sustainable Development, Hanoi University of Science, Hanoi, Viet Nam
- Tana, T.S., Social and Cultural Observation Unit, Office of the Council of Ministers, Phnom Penh, Cambodia
- Tanabe, S., Center for Marine Environmental Studies, Ehime University, Matsuyama, Japan

References:

- Goering, P.L., Aposhian, H.V., Mass, M.J., Cebrin, M., Beck, B.D., Waalkes, M.P., The enigma of arsenic carcinogenesis: Role of metabolism (1999) *Toxicol Sci*, 49 (1), pp. 5-14
- Gorby, M.S., Arsenic in human medicine (1994) *Arsenic in the Environment, Part II: Human Health and Ecosystem Effects*, pp. 1-16. , Nriagu JO, ed. Wiley, New York
- (2004) Guidelines for Drinking Water Quality, pp. 1-515. , World Health Organization (WHO) Third ed. WHO, Geneva
- Mandai, B.K., Suzuki, K.T., Arsenic round the world: A review (2002) *Talanta*, 58 (1), pp. 201-235
- Nordstrom, D.K., Public health. Worldwide occurrences of arsenic in ground water (2002) *Science*, 296 (5576), pp. 2143-2145
- Smedley, P.L., Kinniburgh, D.G., A review of the source, behaviour and distribution of arsenic in natural waters (2002) *Appl Geochem*, 17 (5), pp. 517-568
- Berg, M., Tran, H.C., Nguyen, T.C., Pham, H.V., Schertenleib, R., Giger, W., Arsenic contamination of groundwater and drinking water in Vietnam: A human health threat (2001) *Environ Sci Technol*, 35 (13), pp. 2621-2626
- Nga, T.T.V., Inoue, M., Khatiwada, N.R., Takizawa, S., Heavy metal tracers for the analysis of groundwater contamination: Case study in Hanoi City, Vietnam (2003) *Water Science and Technology: Water Supply*, 3 (1-2), pp. 343-350
- Trang, P.T.K., Berg, M., Viet, P.H., Van Mui, N., Van Meer Jr., D., Bacterial bioassay for rapid and accurate analysis of arsenic in highly variable groundwater samples (2005) *Environ Sci Technol*, 39 (19), pp. 7625-7630
- Berg, M., Luzzi, S., Trang, P.T.K., Viet, P.H., Giger, W., Stuifbergen, D., Arsenic removal from groundwater by household sand

- filters: Comparative field study, model calculations, and health benefits (2006) *Environ Sci Technol*, 40 (17), pp. 5567-5573
- Shinkai, Y., Duong, V.T., Sumi, D., Canh, D., Kumagai, Y., Arsenic and other metal contamination of groundwater in the Mekong River Delta, Vietnam (2007) *Journal of Health Science*, 53 (3), pp. 344-346. , [http://jhs.pharm.or.jp/53\(3\)/53\\_344.pdf](http://jhs.pharm.or.jp/53(3)/53_344.pdf), DOI 10.1248/jhs.53.344
  - Buschmann, J., Berg, M., Stengel, C., Winkel, L., Sampson, M.L., Trang, P.T.K., Contamination of drinking water resources in the Mekong delta floodplains: Arsenic and other trace metals pose serious health risks to population (2008) *Environ Int*, 34 (6), pp. 756-764
  - Norrman, J., Sparrenbom, C.J., Berg, M., Nhan, D.D., Nhan, P.Q., Rosqvist, H., Arsenic mobilisation in a new well field for drinking water production along the Red River, Nam Du, Hanoi (2008) *Appl Geochem*, 23 (11), pp. 3127-3142
  - Nguyen, V.A., Bang, S., Viet, P.H., Kim, K.W., Contamination of groundwater and risk assessment for arsenic exposure in Ha Nam province, Vietnam (2009) *Environ Int*, 35 (3), pp. 466-472
  - Poiya, D.A., Gault, A.G., Diebe, N., Feldman, P., Rosenboom, J.W., Gilligan, E., Arsenic hazard in shallow Cambodian groundwaters (2005) *Mineral Mag*, 69 (5), pp. 807-823
  - Berg, M., Stengel, C., Pham, T.K., Pham, H.V., Sampson, M.L., Leng, M., Magnitude of arsenic pollution in the Mekong and Red River Deltas Cambodia and Vietnam (2007) *Sci Total Environ*, 372 (2-3), pp. 413-425
  - Buschmann, J., Berg, M., Stengel, C., Sampson, M.L., Arsenic and manganese contamination of drinking water resources in Cambodia: Coincidence of risk areas with low relief topography (2007) *Environ Sci Technol*, 41 (7), pp. 2146-2152
  - Gault, A.G., Rowland, H.A.L., Charnock, J.M., Woelius, R.A., Gomez-Morilla, I., Vong, S., Arsenic in hair and nails of individuals exposed to arsenicrich groundwaters in Kandal province, Cambodia (2008) *Sci Total Environ*, 393 (1), pp. 168-176
  - Kocar, B.D., Polizzotto, M.L., Benner, S.G., Ying, S.C., Ung, M., Ouch, K., Integrated biogeochemical and hydrologic processes driving arsenic release from shallow sediments to groundwaters of the Mekong delta (2008) *Appl Geochem*, 23 (11), pp. 3059-3071
  - Sthiannopkao, S., Kim, K.W., Sotham, S., Choup, S., Arsenic and manganese in tube well waters of Prey Veng and Kandal provinces, Cambodia (2008) *Appl Geochem*, 23 (5), pp. 1086-1093
  - Rowland, H.A.L., Gault, A.G., Lythgoe, P., Polya, D.A., Geochemistry of aquifer sediments and arsenicrich groundwaters from Kandal Province, Cambodia (2008) *Appl Geochem*, 23 (11), pp. 3029-3046
  - Mazumder, D.N.G., Majumdar, K.K., Santra, S.C., Kol, H., Vicheth, C., Occurrence of arsenicosis in a rural village of Cambodia (2009) *J Environ Sci Health A Tox Hazard Subst Environ Eng*, 44 (5), pp. 480-487
  - Luu, T.T.G., Sthiannopkao, S., Kim, K.W., Arsenic and other trace elements contamination in groundwater and a risk assessment study for the residents in the Kandal Province of Cambodia (2009) *Environ Int*, 35 (3), pp. 455-460
  - Agusa, T., Kunito, T., Kubota, R., Monirith, I., Tanabe, S., Tana, T.S., Arsenic pollution in Cambodia (2002) *Biomed Res Trace Elem*, 13 (4), pp. 254-255. , in Japanese
  - Agusa, T., Kunito, T., Fujihara, J., Kubota, R., Minh, T.B., Trang, P.T.K., Contamination by trace elements in groundwater of Vietnam (2004) *Biomed Res Trace Elem*, 15 (4), pp. 339-341
  - Agusa, T., Inoue, S., Kunito, T., Kubota, R., Minh, T.B., Trang, P.T.K., Widely-distributed arsenic pollution in groundwater in the Red River Delta, Vietnam (2005) *Biomed Res Trace Elem*, 16 (4), pp. 296-298
  - Agusa, T., Kunito, T., Fujihara, J., Kubota, R., Minh, T.B., Trang, P.T.K., Contamination by arsenic and other trace elements in tube-well water and its risk assessment to humans in Hanoi, Vietnam (2006) *Environ Pollut*, 139 (1), pp. 95-106
  - Agusa, T., Kubota, R., Kunito, T., Minh, T.B., Trang, P.T.K., Chamnan, C., Arsenic pollution in groundwater of Vietnam and Cambodia: A review (2007) *Biomed Res Trace Elem*, 18 (1), pp. 35-47
  - Agusa, T., Kunito, T., Minh, T.B., Trang, P.T.K., Iwata, H., Viet, P.H., Relationship of urinary arsenic metabolites to intake

- estimates in residents of the Red River Delta, Vietnam (2009) *Environ Pollut.*, 157 (2), pp. 396-403
- Agusa, T., Iwata, H., Fujihara, J., Kunito, T., Takeshita, H., Minh, T.B., Genetic polymorphisms in AS3MT and arsenic metabolism in residents of the Red River Delta, Vietnam (2009) *Toxicol Appl Pharmacol*, 236 (2), pp. 131-141
  - Agusa, T., Inoue, S., Kunito, T., Minh, T.B., Ha, N.N., Tu, N.P.C., Human exposure to arsenic from groundwater in the Red River and the Mekong River Deltas in Vietnam (2009) *Int J Environ Stud*, 66 (1), pp. 49-57
  - Kubota, R., Kunito, T., Agusa, T., Fujihara, J., Monirith, I., Iwata, H., Urinary 8-hydroxy-2' deoxyguanosine in inhabitants chronically exposed to arsenic in groundwater in Cambodia (2006) *J Environ Monit*, 8 (2), pp. 293-299
  - Iwata, H., Kim, E.Y., Yamauchi, M., Inoue, S., Agusa, T., Tanabe, S., Chemical contamination in aquatic ecosystems (2007) *Yakugaku Zasshi-J Pharmaceut Soc Jpn*, 127 (3), pp. 417-428
  - Yamaguchi, S., Miura, C., Ito, A., Agusa, T., Iwata, H., Tanabe, S., Tuyen, B.C., Miura, T., Effects of lead, molybdenum, rubidium, arsenic and organochlorines on spermatogenesis in fish: Monitoring at Mekong Delta area and in vitro experiment (2007) *Aquatic Toxicology*, 83 (1), pp. 43-51. , DOI 10.1016/j.aquatox.2007.03.010, PII S0166445X07001105
  - Yamaguchi, S., Celino, F.T., Ito, A., Agusa, T., Tanabe, S., Tuyen, B.C., Effects of arsenic on gonadal development in freshwater crab, *Somanniathelphusa pax*, in Vietnam and *Geothelphusa dehaani* in Japan (2008) *Ecotoxicology*, 17 (8), pp. 772-780
  - Agusa, T., Iwata, H., Fujihara, J., Kunito, T., Takeshita, H., Minh, T.B., Genetic polymorphisms in glutathione S-transferase (GST) superfamily and arsenic metabolism in residents of the Red River Delta, Vietnam (2010) *Toxicol Appl Pharmacol*, 242 (3), pp. 352-362
  - Tanabe, S., Environ Specimen Bank in Ehime University (es-BANK), Japan for global monitoring (2006) *J Environ Monit*, 8 (8), pp. 782-790
  - Harvey, C.F., Ashfaque, K.N., Yu, W., Badruzzaman, A.B.M., Ali, Ma., Oates, P.M., Groundwater dynamics and arsenic contamination in Bangladesh (2006) *Chem Geol*, 228 (1-3), pp. 112-136. , Special Issue
  - Harvey, C.F., Beckie, R.D., Arsenic: Its biogeochemistry and transport in groundwater (2005) *Met Ions Biol Syst*, 44, pp. 145-169
  - McArthur, J.M., Banerjee, D.M., Hudson-Edwards, K.A., Mishra, R., Purohit, R., Ravenscroft, P., Natural organic matter in sedimentary basins and its relation to arsenic in anoxic ground water: The example of West Bengal and its worldwide implications (2004) *Appl Geochem*, 19 (8), pp. 1255-1293
  - Meng, X., Korfiatis, G.P., Christodoulatos, C., Bang, S., Treatment of arsenic in Bangladesh well water using a household co-precipitation and filtration system (2001) *Water Res*, 35 (12), pp. 2805-2810
  - Meng, X., Korfiatis, G.P., Bang, S., Bang, K.W., Combined effects of anions on arsenic removal by iron hydroxides (2002) *Toxicology Letters*, 133 (1), pp. 103-111. , DOI 10.1016/S0378-4274(02)00080-2, PII S0378427402000802
  - Meharg, A.A., Rahman, M.M., Arsenic contamination of Bangladesh paddy field soils: Implications for rice contribution to arsenic consumption (2003) *Environ Sci Technol*, 37 (2), pp. 229-234
  - Abedin, M.J., Cresser, M.S., Meharg, A.A., Feldmann, J., Cotter-Howells, J., Arsenic accumulation and metabolism in rice (*Oryza sativa L.*) (2002) *Environ Sci Technol*, 36 (5), pp. 962-968
  - Williams, P.N., Price, A.H., Raab, A., Hossain, S.A., Feldmann, J., Meharg, A.A., Variation in arsenic speciation and concentration in paddy rice related to dietary exposure (2005) *Environ Sci Technol*, 39 (15), pp. 5531-5540
  - Schoof, R.A., Yost, L.J., Eickhoff, J., Crecelius, E.A., Cragn, D.W., Meacher, D.M., A market basket survey of inorganic arsenic in food (1999) *Food Chem Toxicol*, 37 (8), pp. 839-846
  - Meharg, A.A., Lombi, E., Williams, P.N., Scheckel, K.G., Feldmann, J., Raab, A., Speciation and localization of arsenic in white and brown rice grains (2008) *Environ Sci Technol*, 42 (4), pp. 1051-1057

- Phuong, T.D., Chuong, P.V., Khiem, D.T., Kokot, S., Elemental content of Vietnamese rice. Part 1. Sampling, analysis and comparison with previous studies (1999) *Analyst*, 124 (4), pp. 553-560
- Watanabe, C., Kawata, A., Sudo, N., Sekiyama, M., Inaoka, T., Bae, M., Water intake in an Asian population living in arsenic-contaminated area (2004) *Toxicol Appl Pharmacol*, 198 (3), pp. 272-282
- (2005) Food Security Statistics, , [http://www.fao.org/faostat/foodsecurity/index\\_en.htm](http://www.fao.org/faostat/foodsecurity/index_en.htm)
- Evaluation of ce
- tain food additives and contaminants (1989) 33rd Report of the Joint FAO/WHO Expert Committee on Food Additives, , World Health Organization (WHO) Technical Report Series 776. WHO, Geneva
- Arnold Jr., H.L., Odom, R.B., James, W.D., (1990) Andrew's Diseases of the Skin: Clinical Dermatology, , 8th ed. WB Saunders Company, Philadelphia
- Hopenhayn-Rich, C., Biggs, M.L., Kalman, D.A., Moore, L.E., Smith, A.H., Arsenic methylation patterns before and after changing from high to lower concentrations of arsenic in drinking water (1996) *Environmental Health Perspectives*, 104 (11), pp. 1200-1207
- Stýblo, M., Drobna, Z., Jaspers, I., Lin, S., Thomas, D.J., The role of biomethylation in toxicity and carcinogenicity of arsenic: A research update (2002) *Environ Health Perspect*, 110 (SUPPL. 5), pp. 767-771
- Challenger, F., Biological methylation (1945) *Chem Rev*, 36 (3), pp. 315-361
- Cullen, W.R., Reimer, K.J., Arsenic speciation in the environment (1989) *Chem Rev*, 89 (4), pp. 713-764
- Hayakawa, T., Kobayashi, Y., Cui, X., Hirano, S., A new metabolic pathway of arsenite: Arsenic-glutathione complexes are substrates for human arsenic methyltransferase Cyt19 (2005) *Arch Toxicol*, 79 (4), pp. 183-191
- Naranmandura, H., Suzuki, N., Suzuki, K.T., Trivalent arsenicals are bound to proteins during reductive methylation (2006) *Chemical Research in Toxicology*, 19 (8), pp. 1010-1018. , DOI 10.1021/tx060053f
- Zakharyan, R.A., Tsaprailis, G., Chowdhury, U.K., Hernandez, A., Aposhian, H.V., Interactions of sodium selenite, glutathione, arsenic species, and omega class human glutathione transferase (2005) *Chem Res Toxicol*, 18 (8), pp. 1287-1295
- Lin, S., Shi, Q., Brent Nix, F., Styblo, M., Beck, M.A., Herbin-Davis, K.M., A novel S-adenosyl-Lmethionine:arsenic(III) methyltransferase from rat liver cytosol (2002) *J Biol Chem*, 277 (13), pp. 10795-10803
- Wood, T.C., Salavagionne, O.E., Mukherjee, B., Wang, L., Klumpp, A.F., Thomae, B.A., Eckloff, B.W., Weinshilboum, R.M., Human arsenic methyltransferase (AS3MT) pharmacogenetics: Gene resequencing and functional genomics studies (2006) *Journal of Biological Chemistry*, 281 (11), pp. 7364-7373. , <http://www.jbc.org/cgi/reprint/281/11/7364>, DOI 10.1074/jbc.M512227200
- Vahter, M., Methylation of inorganic arsenic in different mammalian species and population groups (1999) *Sci Prog*, 82
- Del Razo, L.M., Garcia-Vargas, G.G., Vargas, H., Albores, A., Gonsebatt, M.E., Montero, R., Altered profile of urinary arsenic metabolites in adults with chronic arsenicism (1997) A Pilot Study. *Arch Toxicol*, 71 (4), pp. 211-217
- Chowdhury, U.K., Rahman, M.M., Sengupta, M.K., Lodh, D., Chanda, C.R., Roy, S., Pattern of excretion of arsenic compounds [arsenite, arsenate, MMA(V), DMA(V)] in urine of children compared to adults from an arsenic exposed area in Bangladesh (2003) *J Environ Sci Health A Tox Hazard Subst Environ Eng*, 38 (1), pp. 87-113
- Meza, M.M., Yu, L., Rodriguez, Y.Y., Guild, M., Thompson, D., Gandolfi, A.J., Developmentally restricted genetic determinants of human arsenic metabolism: Association between urinary methylated arsenic and CYT19 polymorphisms in children (2005) *Environ Health Perspect*, 113 (6), pp. 775-781
- Chung, J.S., Kalman, D.A., Moore, L.E., Kosnett, M.J., Arroyo, A.P., Beeris, M., Family correlations of arsenic methylation patterns in children and parents exposed to high concentrations of arsenic in drinking water (2002) *Environ Health Perspect*, 110 (7), pp. 729-733

- Chiou, H.Y., Hsueh, Y.M., Hsieh, L.L., Hsu, L.I., YiHsiang, H., Hsieh, F.I., Arsenic methylation capacity, body retention, and null genotypes of glutathione S-transferase M1 and T1 among current arsenic-exposed residents in Taiwan (1997) *Mutat Res*, 386 (3), pp. 197-207
- Loffredo, C.A., Aposhian, H.V., Cebrian, M.E., Yamauchi, H., Silbergeld, E.K., Variability in human metabolism of arsenic (2003) *Environ Res*, 92 (2), pp. 85-91
- Watanabe, C., Inaoka, T., Kadono, T., Nagano, M., Nakamura, S., Ushijima, K., Murayama, N., Ohtsuka, R., Males in rural Bangladeshi communities are more susceptible to chronic arsenic poisoning than females: Analyses based on urinary arsenic (2001) *Environmental Health Perspectives*, 109 (12), pp. 1265-1270
- Tseng, C.H., A review on Environ factors regulating arsenic methylation in humans (2009) *Toxicol Appl Pharmacol*, 235 (3), pp. 338-350
- Bailey, K.V., Ferro-Luzzi, A., Use of body mass index of adults in assessing individual and community nutritional status (1995) *Bull WHO*, 73, pp. 673-680
- Board, P.G., Coggan, M., Chelvanayagam, G., Easteal, S., Jermini, L.S., Schulte, G.K., Identification, characterization, and crystal structure of the omega class glutathione transferases (2000) *J Biol Chem*, 275 (32), p. 24. , 798-806
- Zakharyan, R.A., Aposhian, H.V., Enzymatic reduction of arsenic compounds in mammalian systems: The rate-limiting enzyme of rabbit liver arsenic biotransformation is MMA(V) reductase (1999) *Chem Res Toxicol*, 12 (12), pp. 1278-1283
- Zakharyan, R.A., Tsaprailis, G., Chowdhury, U.K., Hernandez, A., Aposhian, H.V., Interactions of sodium selenite, glutathione, arsenic species, and omega class human glutathione transferase (2005) *Chem Res Toxicol*, 18 (8), pp. 1287-1295
- Whitbread, A.K., Tetlow, N., Eyre, H.J., Sutherland, G.R., Board, P.G., Characterization of the human Omega class glutathione transferase genes and associated polymorphisms (2003) *Pharmacogenetics*, 13 (3), pp. 131-144
- Schmuck, E.M., Board, P.G., Whitbread, A.K., Tetlow, N., Cavanaugh, J.A., Blackburn, A.C., Characterization of the monomethylarsonate reductase and dehydroascorbate reductase activities of Omega class glutathione transferase variants: Implications for arsenic metabolism and the age-at-onset of Alzheimer's and Parkinson's diseases (2005) *Pharmacogenet Genomics*, 15 (7), pp. 493-501
- Tanaka-Kagawa, T., Jinno, H., Hasegawa, T., Makino, Y., Seko, Y., Hanioka, N., Functional characterization of two variant human GSTO 1-1s (Ala 140Asp and Thr217Asn) (2003) *Biochem Biophys Res Commun*, 301 (2), pp. 516-520
- Chung, C.J., Hsueh, Y.M., Bai, C.H., Huang, Y.K., Huang, Y.L., Yang, M.H., Polymorphisms in arsenic metabolism genes, urinary arsenic methylation profile and cancer (2009) *Cancer Causes Control*, 20 (9), pp. 1653-1661
- Hernandez, A., Xamena, N., Surralles, J., Sekaran, C., Tokunaga, H., Quinteros, D., Role of the Met(287)Thr polymorphism in the AS3MT gene on the metabolic arsenic profile (2008) *Mutat Res*, 637 (1-2), pp. 80-92
- Lindberg, A.-L., Kumar, R., Goessler, W., Thirumaran, R., Gurzau, E., Koppova, K., Rudnai, P., Vahter, M., Metabolism of low-dose inorganic arsenic in a central European population: Influence of sex and genetic polymorphisms (2007) *Environmental Health Perspectives*, 115 (7), pp. 1081-1086. , <http://www.ehponline.org/members/2007/10026/10026.pdf>, DOI 10.1289/ehp.10026
- Marnell, L.L., Garcia-Vargas, G.G., Chowdhury, U.K., Zakharyan, R.A., Walsh, B., Avram, M.D., Polymorphisms in the human monomethylarsonic acid (MMA V) reductase/hGSTO1 gene and changes in urinary arsenic profiles (2003) *Chem Res Toxicol*, 16 (12), pp. 1507-1513
- Paiva, L., Marcos, R., Creus, A., Coggan, M., Oakley, A.J., Board, P.G., Polymorphism of glutathione transferase Omega 1 in a population exposed to a high Environ arsenic burden (2008) *Pharmacogenet Genomics*, 18 (1), pp. 1-10
- Schläwicke Engström, K., Broberg, K., Concha, G., Nermell, B., Warholm, M., Vahter, M., Genetic polymorphisms influencing arsenic metabolism: Evidence from Argentina (2007) *Environ Health Perspect*, 115 (4), pp. 599-605

- Kile, M.L., Houseman, E.A., Rodrigues, E., Smith, T.J., Quamruzzaman, Q., Rahman, M., Toenail arsenic concentrations, GSTT1 gene polymorphisms, and arsenic exposure from drinking water (2005) *Cancer Epidemiol Biomarkers Prev*, 14 (10), pp. 2419-2426
- Lin, G.F., Du, H., Chen, J.G., Lu, H.C., Kai, J.X., Zhou, Y.S., Glutathione S-transferases M1 and T1 polymorphisms and arsenic content in hair and urine in two ethnic clans exposed to indoor combustion of high arsenic coal in Southwest Guizhou, China (2007) *Arch Toxicol*, 81 (8), pp. 545-551
- Marcos, R., Martinez, V., Hernandez, A., Creus, A., Sekaran, C., Tokunaga, H., Metabolic profile in workers occupationally exposed to arsenic: Role of GST polymorphisms (2006) *J Occup Environ Med*, 48 (3), pp. 334-341
- McCarty, K.M., Chen, Y.C., Quamruzzaman, Q., Rahman, M., Mahiuddin, G., Hsueh, Y.M., Arsenic methylation, GSTT1, GSTM1, GSTP1 polymorphisms, and skin lesions (2007) *Environ Health Perspect*, 115 (3), pp. 341-345
- Steinmaus, C., Moore, L.E., Shipp, M., Kalman, D., Rey, O.A., Biggs, M.L., Hopenhayn, C., Smith, A.H., Genetic polymorphisms in MTHFR 677 and 1298, GSTM1 and T1, and metabolism of arsenic (2007) *Journal of Toxicology and Environmental Health - Part A: Current Issues*, 70 (2), pp. 159-170. , DOI 10.1080/15287390600755240, PII 769176708
- Zhong, S.L., Zhou, S.F., Chen, X., Chan, S.Y., Chan, E., Ng, K.Y., Relationship between genotype and enzyme activity of glutathione S-transferases M1 and P1 in Chinese (2006) *Eur J Pharmaceut Sci*, 28 (1-2), pp. 77-85
- Mukherjee, B., Salavaggione, O.E., Pelleymounter, L.L., Moon, I., Eckloff, B.W., Schaid, D.J., Glutathione S-transferase omega 1 and omega 2 pharmacogenomics (2006) *Drug Metabol Dispos*, 34 (7), pp. 1237-1246
- An, Y., Gao, Z., Wang, Z., Yang, S., Liang, J., Feng, Y., Immunohistochemical analysis of oxidative DNA damage in arsenic-related human skin samples from arsenic-contaminated area of China (2004) *Cancer Lett*, 214 (1), pp. 11-18
- Bau, D.-T., Wang, T.-S., Chung, C.-H., Wang, A.S.S., Jan, K.-Y., Oxidative DNA adducts and DNA-protein cross-links are the major DNA lesions induced by arsenite (2002) *Environ Health Perspect*, 110 (SUPPL. 5), pp. 753-756
- Kitchin, K.T., Ahmad, S., Oxidative stress as a possible mode of action for arsenic carcinogenesis (2003) *Toxicol Lett*, 137 (1-2), pp. 3-13
- Rossman, T.G., Mechanism of arsenic carcinogenesis: An integrated approach (2003) *Mutat Res*, 533 (1-2), pp. 37-65
- Floyd, R.A., The role of 8-hydroxyguanine in carcinogenesis (1990) *Carcinogenesis*, 11 (9), pp. 1447-1450
- Kasai, H., Analysis of a form of oxidative DNA damage, 8-hydroxy-2'-deoxyguanosine, as a marker of cellular oxidative stress during carcinogenesis (1997) *Mutat Res*, 387 (3), pp. 147-163
- Yamauchi, H., Aminaka, Y., Yoshida, K., Sun, G., Pi, J., Waalkes, M.P., Evaluation of DNA damage in patients with arsenic poisoning: Urinary 8-hydroxydeoxyguanine (2004) *Toxicol Appl Pharmacol*, 198 (3), pp. 291-296
- Fujino, Y., Guo, X., Liu, J., Matthews, I.P., Shirane, K., Wu, K., Chronic arsenic exposure and urinary 8-Hydroxy-2-deoxyguanosine in an arsenicaffected area in Inner Mongolia, China (2005) *J Expo Anal Environ Epidemiol*, 15 (2), pp. 147-152
- Matsui, M., Nishigori, C., Toyokuni, S., Takada, J., Akaboshi, M., Ishikawa, M., Imamura, S., Miyachi, Y., The role of oxidative DNA damage in human arsenic carcinogenesis: Detection of 8-hydroxy-2'-deoxyguanosine in arsenic-related Bowen's disease (1999) *Journal of Investigative Dermatology*, 113 (1), pp. 26-31. , DOI 10.1046/j.1523-1747.1999.00630.x
- <http://www.patient.co.uk/doctor/Arsenic-Poisoning.htm>Dang, M.N., Nguyen, K.H., Chander, B., Nguyen, Q.H., (2004) The Adverse Effects of Arsenic on Population Health in Selected Communities of Ha Nam Province, , Workshop of Science and Technol Relating to Arsenic Contamination. Hanoi, Vietnam
- Mazumder, D.N.G., Majumdar, K.K., Santra, S.C., Kol, H., Vicheth, C., Occurrence of arsenicosis in a rural village of Cambodia (2009) *J Environ Sci Health A Tox Hazard Subst Environ Eng*, 44 (5), pp. 480-487
- Stanger, G., A palaeo-hydrogeological model for arsenic contamination in southern and south-east Asia (2005) *Environ*

Geochem Health, 27 (4), pp. 359-368

- Diaz, O.P., Leyton, I., Munoz, O., Nunez, N., Devesa, V., Suner, Ma., Contribution of water, bread, and vegetables (raw and cooked) to dietary intake of inorganic arsenic in a rural village of Northern Chile (2004) *J Agric Food Chem*, 52 (6), pp. 1773-1779
- Del Razo, L.M., Garcia-Vargas, G.G., Garcia-Salcedo, J., Sanmiguel, M.F., Rivera, M., Hernandez, M.C., Arsenic levels in cooked food and assessment of adult dietary intake of arsenic in the Region Lagunera, Mexico (2002) *Food Chem Toxicol*, 40 (10), pp. 1423-1431
- Roychowdhury, T., Tokunaga, H., Ando, M., Survey of arsenic and other heavy metals in food composites and drinking water and estimation of dietary intake by the villagers from an arsenic-affected area of West Bengal, India (2003) *Science of the Total Environment*, 308 (1-3), pp. 15-35. , DOI 10.1016/S0048-9697(02)00612-5, PII S0048969702006125
- Kile, M.L., Houseman, E.A., Breton, C.V., Smith, T., Quamruzzaman, Q., Rahman, M., Dietary arsenic exposure in Bangladesh (2007) *Environ Health Perspect*, 115 (6), pp. 889-893