

Susan J. Borghoff, M.S.P.H., Ph.D., DABT

VICE PRESIDENT, 21CT PRINCIPAL SCIENTIST

CONTACT INFORMATION

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PROFESSIONAL PROFILE

Dr. Susan Borghoff is based in Research Triangle Park, NC. She is a board-certified toxicologist with over 35 years of experience designing and managing diverse research programs and regulator-driven *in vitro* and *in vivo* toxicology studies for commercial and federal clients. Her work has also included scientific and technical oversight, data analysis and interpretation, and preparation of both Final GLP Study Reports for regulatory submission and peer-reviewed scientific manuscripts. Dr. Borghoff has directed scientific research to support dose-response and hazard identification for incorporation into human health risk assessments.

Dr. Borghoff is a recognized expert on mechanisms by which agents cause toxicity, modulate endocrine pathways, and cause cancer in rodents, and the relevance of these responses for assessing human risk. Most recently, she has been involved in evaluating mechanistic data combined with experimental cancer animal studies to assess the potential of selected sweeteners to cause cancer in humans. Her research also has focused on understanding the metabolism and pharmacokinetics of agents using physiologically based pharmacokinetic (PBPK) models for extrapolation of risk to humans. She has been involved with critical reviews of toxicity and carcinogenicity studies, advising on specific study designs to fill data gaps for understanding modes/mechanisms of action, and as an Expert Panel member for scientific oversight of industry-sponsored toxicity and carcinogenicity testing programs. Dr. Borghoff's experience also includes implementing a program to conduct GLP regulatory studies associated with the US Environmental Protection Agency (USEPA)-mandated Endocrine Disruption Screening Program (EDSP) and co-chairing a workshop titled, "Lessons Learned, Challenges, and Opportunities: The US Endocrine Disruptor Screening Program," held in Research Triangle Park in 2013, to assess this program.









Dr. Borghoff has served as a reviewer on a number of panels and working groups for international organizations, including USEPA, National Cancer Institute, International Programme on Chemical Safety (IPCS), European Centre for Ecotoxicology and Toxicology of Chemicals (ECETOC), and three Monograph Working Groups (vol. 73, 77, and 82) of the International Agency for Cancer Research (IARC), in addition to acting as Observer for Monograph vol.115, Some Industrial Chemicals. She was also an observer at the advisory group meeting to recommend priorities for the IARC Monographs during 2020–2024 (March 2019). She has been a reviewer for various grants through the National Institutes of Health (NIH), the National Institute of Environmental Health Sciences (NIEHS), and USEPA.

Dr. Borghoff received the Frank R. Blood Award in 1994 for the best paper of the year published in a Society of Toxicology (SOT) research journal, and an SOT Risk Assessment Specialty Section Award in 2000. She is a member of SOT's North Carolina Chapter, and at the national level, has served on SOT Council, as well as various committees (e.g., Program, Awards, Education). Dr. Borghoff was a member of SOT's TSCA Reform Task Force and was the Audit Committee Co-Chair. She has also served as an Associate Editor for *Toxicological Sciences*, along with other editorial positions for several scientific journals, including the editorial board of the *International Journal of Toxicology*.

Dr. Borghoff received her Ph.D. and MSPH in Environmental Sciences and Engineering from the University of North Carolina, and a B.S. in Chemistry from East Stroudsburg University in Pennsylvania. Dr. Borghoff became a Diplomate of the American Board of Toxicology in 1994.

EDUCATION AND DEGREES EARNED

Doctor of Philosophy (PhD) Environmental Sciences and Engineering University of North Carolina

Master of Science in Public Health (MSPH) Environmental Sciences and Engineering University of North Carolina

Bachelor of Science (BS) Chemistry (magna cum laude) East Stroudsburg University

CERTIFICATIONS

Diplomate, American Board of Toxicology (Recertification: 1999, 2004, 2009, 2014, 2020, 2025)

PROFESSIONAL AFFILIATIONS

| 1984-Present | North Carolina Chapter of the Society of Toxicology |
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| 1986-Present | Society of Toxicology (Full Member over 35 yrs) |
| 2019-Present | American Society for Cellular and Computational Toxicology |
| 2015–2017 | Audit Committee; Co-chair 2015, Chair 2016 (appointed) |
| 2013–2016 | TSCA Reform Task Force (appointed) |



| 2009–2011 | Councilor (elected) |
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| 2005–2008 | Education Committee (elected) |
| 2002–2004 | Awards Committee (elected) |
| 1999–2004 | Program Committee (appointed) |
| 2002–2004 | Secretary/Treasurer of BMSS |
| 2007-Present | American College of Toxicology |
| 2013-Present | SETAC (Society of Environmental Toxicology and Chemistry) |
| 2018-Present | Toxicology Forum |
| 1992–2010 | International Society for the Study of Xenobiotics |
| 2003–2010 | RTP Drug Metabolism Discussion Group (Executive Board) |

AWARDS AND HONORS

| 2017 | Recognized by the Biological Modeling Specialty Section of the Society of Toxicology for publication of one of the top nine modeling papers in 2017: Borghoff SJ, Ring C, Banton M, Leavens TL. 2016. Physiologically based pharmacokinetic model for ethyl tertiary-butyl ether and tertiary-butyl alcohol in rats: Contribution of binding to α 2u-globulin in male rats and high-exposure nonlinear kinetics to toxicity and cancer outcomes. J Appl Toxicol 37(5):621–640, doi 10.1002/jat.3412. |
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| 2000 | Society of Toxicology Risk Assessment Specialty Section Award for outstanding presentation in risk assessment: "A physiologically-based pharmacokinetic model for methyl tert-butyl ether in humans" (Student Award—Amy Collins Licata). |
| 1994 | Frank R. Blood Award for Outstanding Paper published in SOT journal: Borghoff SJ, Lagarde WH. 1993. Assessment of binding of 2,4,4-trimethyl-2-pentanol to low-molecular weight proteins isolated from kidneys of male rats and humans. Toxicol Appl Pharmacol 119:228–235. |
| 1987 | National Kidney Foundation Research Fellowship, North Carolina Society of Toxicology—Study Travel Award |
| 1986 | National Society of Toxicology—Study Travel Award |
| 1982–1987 | Public Health Traineeship—University of North Carolina |

EDITORIAL AND REVIEWER ACTIVITIES

Associate Editor

Toxicological Sciences (2004–2008)—focus on reviews associated with PBPK models, pharmacokinetics, endocrine disruptors, kidney toxicity, and cancer



Editorial Board

International Journal of Toxicology (2014–2023) Chemico-Biological Interactions (2000–2008) Toxicological Sciences (2000–2003) Toxicology Letters (1995–2000)

Ad Hoc Reviewer

Regulatory Toxicology and Pharmacology
Environmental Health Perspectives
Toxicology and Applied Physiology
CRC Reviews in Toxicology
Journal of Toxicology and Environmental Health
Life Sciences Reviews
Toxicological Pathology
Risk Analysis

NATIONAL & INTERNATIONAL ADVISORY COMMITTEES, SCIENTIFIC MEETING ORGANIZER, AND INVITED SPEAKER

Chair and Speaker, Informational session: Five decades of evaluating the safety of aspartame consumption: Has anything changed? Society of Toxicology, March 2024.

International Agency for Research on Cancer (IARC) - Observer for Monograph on the Evaluation of Carcinogenic Risks to Humans, Volume 134; Aspartame, Methyleugenol, and Isoeugenol, Lyon, France, June 2023.

Session co-chair: "Advances in Thresholds of Toxicological Concern (TTC): Expansion of Databases to Support New Chemical Space." Also presented: "Integration of NAM data to identify chemical substances with endocrine disruptor properties under ECHA/EFSA guidance." 44th Annual Winter Meeting, the Toxicology Forum, Tysons, VA, January 2020.

44th Annual Summer Meeting of the Toxicology Forum (2018). Kinetically-Derived Maximum Dose: A Key Initiating Event Impacting Need for Mode of Action Investigations of High-Dose Specific Toxicity (Presenter).

42nd Annual Winter Meeting of the Toxicology Forum (2018). Key Characteristics of Carcinogens: Refining Approaches to Systematic Evaluations of Mechanistic Data (Session Chair); Identification of Mechanistic Data According to Key Characteristics of Carcinogens (KCC) and Review of Studies for Reliability, Relevance, and Activity (Presenter).

American College of Toxicology, Advanced Comprehensive Toxicology Class—Lecture on Kidney Toxicology, August 2017.

42nd Annual Summer Meeting of the Toxicology Forum (2016). Update: Organization and Review of Mechanistic Data According to "Ten Key Characteristics of Carcinogens" (TKCC) (Presenter).

International Agency for Research on Cancer (IARC) — Observer for Monograph on the Evaluation of Carcinogenic Risks to Humans, Volume 115: Some Industrial Chemicals. Lyon, France, 2–9 February 2016.

American College of Toxicology Annual Meeting, November 2013, San Antonio, TX

Session Co-Chair of "Prospective and Retrospective Approaches to Characterize Potential Neurotoxicity"



Invited Speaker of session titled, "Endocrine Disruption Screening — Status of the USEPA and EU programs: State of the Science and Slate for the Future"

Co-Organizer and Session Chair Symposium titled, "Population Diversity Mouse Models and their Use in Toxicology." Toxicology Forum, Aspen, CO, July 2013.

Co-Organizer and Session Chair of a Workshop titled, "Lessons Learned, Challenges, and Opportunities: The US Endocrine Disruptor Screening Program." April 22–23, 2013, Research Triangle Park, NC.

Invited Speaker, Toxicology Forum Meeting, January 2012, "Using a Rat MTBE PBPK Model as a Tool to Evaluate: Mode-of-Action and Route to Route Extrapolation."

Grant reviewer, "Engineered Nanomaterials: Linking Physical and Chemical Properties to Biology (U19)," NIEHS (2010).

Invited Speaker — REACH Symposium, "Fulfilling Technical Requirements." Society of Toxicology presentation (2008).

Ad hoc member of the Voluntary Children's Chemical Evaluation Program (VCCEP) panel to review the assessment of ethylbenzene prepared by the American Chemistry Council (February 2007).

Reviewer on NIEHS Special Emphasis Panel for Absorption, Distribution, Metabolism and Excretion (ADME) Chemical Disposition in Mammals (RFP NIH ES-06-01) (August 17–18, 2006).

USEPA Science Advisory Board — Environmental Health Committee, Consultant on the Trichloroethylene Health Risk Assessment: Synthesis and Characterization Review Panel (June 2002).

International Agency for Research on Cancer (IARC) — Working Group Member for Monograph on the Evaluation of Carcinogenic Risks to Humans, Vol. 82, "Traditional Herbal Medicines, Some Mycotoxins, Naphthalene and Styrene" (2002).

International Agency for Research on Cancer (IARC) — Working Group Member for Monograph on the Evaluation of Carcinogenic Risks to Humans, Vol. 77, "Some Industrial Chemicals" (2000).

European Centre for Ecotoxicology and Toxicology of Chemicals (ECETOC) — Corresponding member on the MTBE Risk Assessment Document (2000).

USEPA. Invited participant and presenter, Workshop on MTBE Pharmacokinetic-Based Extrapolation (May 2000).

Invited participant in a meeting of the ECETOC Task Force on MTBE Risk Assessment and the Finnish Competent Authorities preparing the European Union's Risk Assessment of MTBE (May 2000).

Invited Speaker, Novartis Crop Protection AG, Basel Switzerland, "Criteria for Evaluating the Ability of a Chemical to Cause α2u-Globulin-Mediated Renal Tumors in Male Rats," (November 1999)

Society of Toxicology Annual Meeting — Organizer, Co-Chair, and participant of the symposium titled, "Aliphatic Ethers as Fuel Oxygenates: Health Effects and Regulatory Issues" (March 1999).

Reviewer of NIEHS Superfund Basic Research Program Grant (October 1999).

International Agency for Research on Cancer (IARC) — Working Group Member for Monograph on the Evaluation of Carcinogenic Risks to Humans, Vol. 73, "Some Agents that Cause Tumors of the Kidney, or Urinary Bladder in Rodents and some other Substances" (1998).

Invited Toxicology Scholar Colloquium Speaker, Center for Biochemical Toxicology, University of Connecticut, "The use of mechanistic information in risk assessment-chemical induced α 2u-globulin mediated renal tumors." May 1998.



Invited Participant — Finnish Institute of Occupational Heath, Helsinki, Finland, "Mechanism of MTBE-induced renal tumors in male rats." October 1998.

Invited Participant – Office of Environmental Health Hazard Assessment, Sacramento, CA, "Mechanism of MTBE-induced tumors in rodents." July 22, 1998.

European Centre for Ecotoxicology and Toxicology of Chemicals (ECETOC) — Member of the task force preparing a document on Toxicological Mechanisms (1997–2001).

International Program on Chemical Safety, Methyl Tertiary Butyl Ether, Sponsored by Health Canada. (March 1997).

Workshop to Provide Guidance on the Feasibility and Design of Epidemiologic Studies among Populations Exposed to MTBE. Sponsored by the USEPA, Panelist (April 4–5, 1995).

Industry Briefing on Oxyfuel Studies. USEPA, Washington D.C., Presenter (April 1995).

Potential Acute Health Effects of Oxygenates in Fuels: A Workshop of the Health Effects Institute Oxygenates Review, Chicago, IL. Presenter — Pharmacokinetics of MTBE Across Species (July 1995).

Oxyfuels Information Needs. USEPA, Washington D.C., Presenter (September 1995).

PUBLICATIONS

Nelms MD, Antonijevic T, Ring C, Harris DL, Bever RJ, Lynn SG, ... **Borghoff S**, et al. 2024. Chemistry domain of applicability evaluation against existing estrogen receptor high-throughput assay-based activity models. Front Toxicol 6:https://doi.org/10.3389/ftox.2024.1346767.

Bever RJ, Edwards SW, Antonijevic T, Nelms MD, Ring C, Harris D, ... **Borghoff S**, Markey KJ. 2024. Optimizing androgen receptor prioritization using high-throughput assay-based activity models. Front Toxicol 6:https://www.frontiersin.org/articles/10.3389/ftox.2024.1347364/full.

Henderson RG, Welsh BT, Rogers JM, **Borghoff SJ**, Trexler KR, Bonn-Miller MO, Lefever TW. 2023. Reproductive and developmental toxicity evaluation of cannabidiol. Food Chem Toxicol 176:113786. doi: 10.1016/j.fct.2023.113786.

Henderson RG, Lefever TW, Heintz MM, Trexler KR, **Borghoff SJ**, Bonn-Miller MO. 2023. Oral toxicity evaluation of cannabidiol. Food Chem Toxicol 176:113778. doi: 10.1016/j.fct.2023.113778.

Borghoff SJ, Cohen SS, Jiang X, Lea IA, Klaren WD, Chappell GA, Britt JK, Rivera BN, Choksi NY, Wikoff DS. 2023. Updated systematic assessment of human, animal and mechanistic evidence demonstrates lack of human carcinogenicity with consumption of aspartame. Food Chem Toxicol 172:113549. doi: 10.1016/j.fct.2022.113549.

Goyak KO, Sarang SS, Franzen A, **Borghoff SJ**, Ryman-Rasmussen JP. 2022. Adverse outcome pathway (AOP): α 2u-globulin nephropathy and kidney tumors in male rats. Crit Rev Toxicol 52(5):345-357. doi: 10.1080/10408444.2022.2082269.

Lea IA, Pham LL, Antonijevic T, Thompson C, **Borghoff SJ**. 2022. Assessment of the applicability of the threshold of toxicological concern for per- and polyfluoroalkyl substances. Regul Toxicol Pharmacol 133:105190. doi: 10.1016/j.yrtph.2022.105190.

Gentry R, Greene T, Chappell G, Lea I, **Borghoff S**, Yang C, Rathman J, Ribeiro JV, Hobocienski B, Mostrag A, Rodricks J, Clewell H. 2021. Integration of evidence to evaluate the potential for neurobehavioral effects following exposure to USFDA-approved food colors. Food Chem Toxicol 151:112097. doi: 10.1016/j.fct.2021.112097.



Gentry R, Rodricks J, Clewell H, Greene T, Chappell G, Lea I, **Borghoff S**, Yang C, Rathman J, Ribeiro JV, Hobocienski B, Mostrag A. 2021. RE: Response to the Office of Environmental Health Hazard Assessment on comments related to Gentry et al. (2021). Food Chem Toxicol 152:112202. doi: 10.1016/j.fct.2021.112202.

Black MB, Andersen ME, Pendse SN, **Borghoff SJ**, Streicker M, McMullen PD. 2021. RNA-sequencing (transcriptomic) data collected in liver and lung of male and female B6C3F1 mice exposed to various dose levels of 4-methylimidazole for 2, 5, or 28 days. Data in Brief 38:107420. doi: 10.1016/j.dib.2021.107420.

Borghoff SJ, Fitch SE, Black MVB, McMullen PD, Andersen ME, Chappell GA. 2021. A systematic approach to evaluate plausible modes of [action] for mouse lung tumors in mice exposed to 4-methylimidozole. Regul Toxicol Pharmacol 124:104977. doi: 10.1016/j.vrtph.2021.104977.

Chappell GA, Heintz MM, **Borghoff SJ**, Doepker CL, Wikoff DS. 2021. Lack of potential carcinogenicity for steviol glycosides — Systematic evaluation and integration of mechanistic data into the totality of evidence. Food Chem Toxicol 150:112045. doi: 10.1016/j.fct.2021.112045.

Chappell GA, Britt JK, **Borghoff, SJ**. 2020. Systematic assessment of mechanistic data for FDA-certified food colors and neurodevelopmental processes. Food Chem Toxicol 140:111310. doi: 10.1016/j.fct.2020.111310.

Chappell GA, Wikoff DS, Doepker CL, **Borghoff SJ**. 2020. Lack of potential carcinogenicity for acesulfame potassium — Systematic evaluation and integration of mechanistic data into the totality of the evidence. Food Chem Toxicol 141:111375. doi: 10.1016/j.fct.2020.111375.

Pham LL, **Borghoff SJ**, Thompson CM. 2020. Comparison of threshold of toxicological concern (TTC) values to oral reference dose (RfD) values. Regul Toxicol Pharmacol 113:104651. doi: 10.1016/j.yrtph.2020.104651.

Chappell GA, **Borghoff SJ**, Pham L, Doepker CL, Wikoff DS. 2019. Lack of potential carcinogenicity for sucralose — Systematic evaluation and integration of mechanistic data into the totality of the evidence. Food Chem Toxicol 135:110898. doi: 10.1016/j.fct.2019.110898.

Wikoff DS, Chappell GA, Fitch S, Doepker CL, **Borghoff SJ**. 2019. Lack of potential carcinogenicity for aspartame – Systematic evaluation and integration of mechanistic data into the totality of the evidence. Food Chem Toxicol 135:110866. doi: 10.1016/j.fct.2019.110866.

van de Ligt J, **Borghoff SJ**, Yoon M, Ferguson LJ, DeMaio W, McClanahan RH. 2019. Nondetectable or minimal detectable residue levels of N-(n-butyl) thiophosphoric triamide in bovine tissues and milk from a 28-d NBPT dosing study. Transl Anim Sci 3(4): 1606-1616. doi: 10.1093/tas/txz153.

Wikoff DS, Rager JE, Chappell GA, Fitch S, Haws L, **Borghoff SJ**. 2018. A framework for systematic evaluation and quantitative integration of mechanistic data in assessments of potential human carcinogens. Toxicol Sci 167(2):322–335. doi: 10.1093/toxsci/kfy279.

Borghoff SJ, Fitch S, Rager JE, Huggett D. 2018. A hypothesis-driven weight-of-evidence analysis to evaluate potential endocrine activity of perfluorohexanoic acid. Regul Toxicol Pharmacol 99:168–181.

Harrill AH, **Borghoff S**, Zorrilla L, Blystone C, Kissling GE, Malarkey D, Shockley K, Travlos G, DeVito MJ. 2018. NTP Research Report on baseline characteristics of Diversity Outbred (J:DO) mice relevant to toxicology studies. National Toxicology Program Research Report 6, Research Triangle Park, NC. Available at: https://www.ncbi.nlm.nih.gov/books/NBK525004/.

Klaren WD, Ring C, Harris MA, Thompson CM, **Borghoff S**, Sipes NS, Hsieh J-H, Auerbach SS, Rager JE. 2018. Identifying attributes that influence *in vitro*-to-*in vivo* concordance by comparing *in vitro* Tox21 bioactivity versus *in vivo* DrugMatrix transcriptomic responses across 130 chemicals. Toxicol Sci 120:709–723. doi: 10.1093/toxsci/kfy220.



Proctor DM, Suh M, Chappell G, **Borghoff SJ**, Thompson CM, Wiench K, Finch L, Ellis-Hutchings R. 2018. An adverse outcome pathway (AOP) for forestomach tumors induced by non-genotoxic initiating events. Regul Toxicol Pharmacol 96:30–40. doi: 10.1016/j.yrtph.2018.04.016.

Suh M, Proctor DM, Chappell G, Rager JE, Thompson CM, **Borghoff S**, Finch L, Ellis-Hutchings R, Wiench K. 2018. A review of the genotoxic, mutagenic, and carcinogenic potentials of several lower acrylates. Toxicology 402–403:50–67. doi: 10.1016/j.tox.2018.04.006.

Thompson CT, Suh M, Chappell G, **Borghoff S**, Ellis-Hutchings R, Wiench K, Finch L, Proctor DM. 2018. Assessment of the mode of action underlying development of forestomach tumors in rodents following oral exposure to ethyl acrylate and relevance to humans. Regul Toxicol Pharmacol 96:178–189. doi: 10.1016/j.vrtph.2018.05.006.

Borghoff SJ, Ring C, Banton M, Leavens TL. 2016. Physiologically based pharmacokinetic model for ethyl tertiary-butyl ether and tertiary-butyl alcohol in rats: Contribution of binding to α 2u-globulin in male rats and high-exposure nonlinear kinetics to toxicity and cancer outcomes. J Appl Toxicol 37(5):621-640. doi 10.1002/jat.3412

Hughs BJ, Johnson T, Lynch A, **Borghoff SJ**, Green S, Mensing T, Sarang SS, LeBaron MJ. 2016. Methyl isobutyl ketone-induced hepatocellular carcinogenesis in B6C3F1 mice: A constitutive androstane receptor (CAR)-mediated mode of action. Regul Toxicol Pharmacol 81:421–429.

Wikoff DS, Rager JE, Haws LC, **Borghoff SJ**. 2016. A high dose mode of action for Tetrabromobisphenol A-induced uterine adenocarcinomas in Wistar Han rats: A critical evaluation of key events in an adverse outcome pathway framework. Regul Toxicol Pharmacol 77:143–159. doi: 10.1016/j.yrtph.2016.01.018.

Borghoff SJ, Wikoff D, Harvey S, Haws L. 2016. Dose- and time-dependent changes in tissue levels of tetrabromobisphenol A (TBBPA) and its sulfate and glucuronide conjugates following repeated administration to female Wistar Han Rats. Toxicol Rep 155:104–108. doi:10.1016/j.toxrep.2016.01.007.

Borghoff SJ, Poet TS, Green S, Davis J, Hughes B, Mensing T, Sarang SS, Lynch AM, Hard GC. 2015. Methyl isobutyl ketone exposure-related increases in specific measures of α 2u-globulin (α 2u) nephropathy in male rats along with in vitro evidence of reversible protein binding. Toxicology 333:1–13.

Wikoff D, Thompson C, Perry C, White M, **Borghoff S**, Fitzgerald L, Haws LC. 2015. Development of toxicity values and exposure estimates for tetrabromobisphenol A (TBBPA): Application in a margin of exposure assessment. J Appl Toxicol 35(11):1292–1308. doi: 10.1002/jat.3132.

Faber W, Kirkpatrick D, Coder P, Li A, **Borghoff S**, Banton M. 2014. Subchronic, reproductive, and maternal toxicity studies with tertiary butyl acetate (TBAC). Regul Toxicol Pharmacol 68:332–342.

Juberg DR, **Borghoff SJ**, Becker RA, Casey W, Hartung T, Holsapple MP, Marty MS, Mihaich EM, Van Der Kraak G, Wade MG, et al. 2014. Lessons learned, challenges, and opportunities: The U.S. Endocrine Disuptor Screening Program. Altex 31(1):63–8.

Becker RA, Bergfelt DR, **Borghoff S**, Davis JP, Hamby BT, O'Connor, J.C., Kaplan, A.M., Sloan, C.S., Tyl, R.W., Wade, M., and Marty, M.S. 2012. Inter-laboratory study comparison of the 15-day intact adult male rat screening assay: Evaluation of an anti-thyroid chemical and a negative control chemical. Birth Defects Res (Part B) 95:63–78.

Borghoff SJ, Parkinson H, Leavens TL. 2010. Physiologically based pharmacokinetic rat model for methyl tertiary-butyl ether; comparison of selected dose metrics following various MTBE exposure scenarios used for toxicity and carcinogenicity evaluation. Toxicology 275(1-3):79–91.

Borghoff SJ, Hard GC, Berdasco NM, Gingell R, Green SM, Gulledge W. 2009. Methyl isobutyl ketone (MIBK) induction of α 2u-globulin nephropathy in male, but not female rats. Toxicology 258:131–138.



Leavens TL, **Borghoff SJ.** 2009. Physiologically based pharmacokinetic model of methyl tertiary butyl ether and tertiary butyl alcohol dosimetry in male rats based on binding to α 2u-globulin. Toxicol Sci 109(2):321–335.

Clewell RA, Kremer JJ, Williams CC, Campbell JL, Jr., Andersen ME, **Borghoff SJ**. 2009. Kinetics of selected dibutyl phthalate metabolites and fetal testosterone following repeated and single administration in pregnant rats. Toxicology 255(1–2):80–90.

Clewell RA, Kremer JJ, Williams CC, Campbell JL, Jr., Andersen ME, **Borghoff SJ**. 2008. Tissue exposures to free and glucuronidated monobutylphthalate in pregnant and fetal rat following exposure to di-n-butylphthalate: Evaluation with a PBPK model. Toxicol Sci 103(2):241–259.

Liao KH, Tan Y, Conolly RB, **Borghoff SJ**, Gargas ML, Andersen ME, Clewell HJ. 2007. Bayesian estimation of pharmacokinetic and pharmacodynamic parameters in a mode-of-action-based cancer risk assessment for chloroform. Risk Anal 27:1535–1551.

Ferguson LJ, Lebetkin EH, Lih FB, Tomer KB, Parkinson HD, **Borghoff SJ**., Burka LT. 2007. 14C-Labeled pulegone and metabolites binding to alph α 2u-globulin in kidneys of male F-344 rats. J Toxicol Environ Health A.70:1416–1423.

Gaido KW, Hensley JB, Liu D, Wallace DG, **Borghoff S**, Johnson KJ, Hall SJ, Boekelheide K. 2007. Fetal mouse phthalate exposure shows that gonocyte multinucleation is not associated with decreased testicular testosterone. Toxicol Sci 97(2):491–503.

Cruzan G, **Borghoff SJ**, de Peyster A, Hard GC, McClain M, McGregor DB, Thomas MG. 2007. Methyl tertiary-butyl ether mode-of-action for cancer endpoints in rodents. Regul Toxicol Pharmacol 47:156–165.

Soucy NV, Parkinson HD, Sochaski MA, **Borghoff SJ**. 2006. Kinetics of genistein and conjugated metabolites in pregnant Sprague-Dawley rats following single and repeated administration of genistein. Toxicol Sci 90:230–240.

Schlosser PM, **Borghoff SJ**, Coldham NG, David JA, Ghosh SK. 2006. Physiologically-based pharmacokinetic modeling of genistein in rats, Part I: Model development. Risk Anal 26:483–500.

Kremer JJ, Williams CW, Parkinson HP, Borghoff SJ. 2005. Pharmacokinetics of monobutylphthalate, the toxic metabolite of di-n-butylphthate, in pregnant rats. Toxicol Lett 159:144–153.

Bauer AK, Faiola B, Abernethy DJ, Marchan R, Pluta LJ, Wong VA, Roberts K., Jaiswal AK, Gonzalez FJ, Butterworth BE, **Borghoff SJ**, Parkinson HD, Everitt JI, Recio L. 2003. Genetic susceptibility to benzene-induced toxicity: Role of NAD(P)H oxidoreductase-1. Cancer Res 63:929–935.

Bauer AK, Faiola B, Abernethy DJ, Marchan R, Pluta LJ, Wong VA, Gonzalez FJ, Butterworth BE, **Borghoff SJ**, Everitt JI, Recio L. 2003. Male mice deficient in microsomal epoxide hydrolase are not susceptible to benzene-induced toxicity. Toxicol Sci 72:201–209.

Licata AC, Dekant W, Smith CE, **Borghoff SJ**. 2001. A physiologically-based pharmacokinetic model for methyl tert-butyl ether in humans: Implementing sensitivity and variability analysis. Toxicol Sci 62:191–204.

Borghoff SJ, Prescott JS, Poet TS, Janszen D, Wong B, Everitt JI. 2001. α 2u-Globulin nephropathy, renal cell proliferation and dosimetry of inhaled tertbutyl alcohol in male and female Fischer 344 rats. Toxicol Sci 61:176–186 (highlighted paper).

Williams TM, **Borghoff SJ**. 2001. Characterization of tert-butyl alcohol binding to alpha2u-globulin in F-344 rats. Toxicol Sci 62(2):228–235.

Williams TM, **Borghoff SJ**. 2000. Induction of testosterone biotransformation enzymes following oral administration of methy tert-butyl ether to male Sprague-Dawley rats. Toxicol Sci 57:147–155.



Williams TM, Cattley RC, **Borghoff SJ**. 2000. Alterations in endocrine responses in male Sprague- Dawley rats following oral administration of methyl tert-butyl ether. Toxicol Sci 54:168–176.

Prescott-Mathews JS, Poet TS, **Borghoff SJ**. 1999. Evaluation of the in vivo interaction of methyl tert-butyl ether with α2u-globulin in male F-344 rats. Toxicol Appl Pharmacol 157:60–67.

Collins AS, Sumner SCJ, **Borghoff SJ**, Medinsky MA. 1999. A physiological model for tert-amyl methyl ether and tert-amyl alcohol: Hypothesis testing of model structures. Toxicol Sci 49:15–28.

Poet TS, Valentine JL, **Borghoff SJ**. 1997. Pharmacokinetics of tertiary butyl alcohol in male and female Fischer 344 rats. Toxicol Lett 92:179–186.

Poet TS, **Borghoff SJ**. 1997. In vitro uptake of methyl tert-butyl ether in male rat kidney: Use of a two compartment model to describe protein interactions. Toxicol Appl Pharmacol 145:340–348.

Prescott-Mathews JS, Wolf DC, Wong BA, **Borghoff SJ**. 1997. Methyl tert-butyl ether causes **α**2uglobulin nephropathy and enhanced renal cell proliferation in male Fischer-344 rats. Toxicol Appl Pharmacol 143:301–314.

Dix KJ, Kedderis GL, **Borghoff SJ**. 1997. Vehicle-dependent oral absorption and target tissue dosimetry of chloroform in male rats and female mice. Toxicol Lett 91:197–209.

Borghoff SJ, Murphy JE, Medinsky MA. 1996. Development of a physiologically-based pharmacokinetic model for methyl tertiary-butyl ether and tertiary-butanol in male Fischer 344 rats. Fundam Appl Toxicol 30:264–275.

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