REFERENCES

Acconcia, F., Pallottini, V., & Marino, M. (2015). Molecular mechanisms of action of BPA. *Dose-Response*, *13*(4). https://doi.org/10.1177/1559325815610582

Adolphs, R. (2013). The biology of fear. In *Current Biology* (Vol. 23, Issue 2, p. R79). NIH Public Access. https://doi.org/10.1016/j.cub.2012.11.055

Ahsan, N., Ullah, H., Ullah, W., & Jahan, S. (2018). Comparative effects of Bisphenol S and Bisphenol A on the development of female reproductive system in rats; a neonatal exposure study. *Chemosphere*, *197*, 336–343. https://doi.org/10.1016/j.chemosphere.2017.12.118

Allard, P., & Colaiácovo, M. P. (2011). Bisphenol A. In *Reproductive and Developmental Toxicology* (pp. 673–686). Elsevier Inc. https://doi.org/10.1016/B978-0-12-382032-7.10050-5

Amraje, F. F. M., Köylü, A., Dizakar, S. Ö. A., Türkoğlu, I., & Ömeroğlu, S. (2018). Bisphenol A and male reproductive system. In *Gazi Medical Journal* (Vol. 29, Issue 3, pp. 271–275). Gazi Universitesi. https://doi.org/10.12996/gmj.2018.76

Arco, A. Del, & Mora, F. (2009). Neurotransmitters and prefrontal cortex-limbic system interactions: Implications for plasticity and psychiatric disorders. *Journal of Neural Transmission*, *116*(8), 941–952. https://doi.org/10.1007/s00702-009-0243-8

Ardy, C., Hamid, I. S., Yudahana, A., Widjiati, Purnama, M. T. E., & Fikri, F. (2020). The Effect of Electric Cigarette Exposure on The Histopathology of Pulmo in Albino Rats (Rattus norvegicus). *Jurnal Medik Veteriner*. https://doi.org/10.20473/jmv.vol3.iss1.2020.38-44

Arnold, S. M., Clark, K. E., Staples, C. A., Klecka, G. M., Dimond, S. S., Caspers, N., & Hentges, S. G. (2013). Relevance of drinking water as a source of human exposure to bisphenol A. *Journal of Exposure Science and Environmental Epidemiology*, *23*(2), 137–144. https://doi.org/10.1038/jes.2012.66

Arredondo, S. B., Valenzuela-Bezanilla, D., Mardones, M. D., & Varela-Nallar, L. (2020). Role of Wnt Signaling in Adult Hippocampal Neurogenesis in Health and Disease. In *Frontiers in Cell and Developmental Biology* (Vol. 8, p. 860). Frontiers Media S.A. https://doi.org/10.3389/fcell.2020.00860

Asimakopoulos, A. G., Xue, J., De Carvalho, B. P., Iyer, A., Abualnaja, K. O., Yaghmoor, S. S., Kumosani, T. A., & Kannan, K. (2016). Urinary biomarkers of exposure to 57 xenobiotics and its association with oxidative stress in a population in Jeddah, Saudi Arabia. *Environmental Research*, *150*, 573–581. https://doi.org/10.1016/j.envres.2015.11.029

Aslanpour, S., Han, S., Schuurmans, C., & Kurrasch, D. M. (2020). Neurog2 acts as a classical proneural gene in the ventromedial hypothalamus and is required for the early phase of neurogenesis. *Journal of Neuroscience*, *40*(18), 3549–3563. https://doi.org/10.1523/JNEUROSCI.2610-19.2020

Auld, D. S., Coassin, P. A., Coussens, N. P., Hensley, P., Klumpp-Thomas, C., Michael, S., Sittampalam, G. S., Trask, O. J., Wagner, B. K., Weidner, J. R., Wildey, M. J., & Dahlin, J. L. (2004). Microplate Selection and Recommended Practices in High-throughput Screening and Quantitative

Biology. In *Assay Guidance Manual*. Eli Lilly & Company and the National Center for Advancing Translational Sciences. http://www.ncbi.nlm.nih.gov/pubmed/32520474

Avchalumov, Y., & Mandyam, C. D. (2021). Plasticity in the hippocampus, neurogenesis and drugs of abuse. In *Brain Sciences* (Vol. 11, Issue 3). MDPI AG. https://doi.org/10.3390/brainsci11030404

Awada, Z., Nasr, R., Akika, R., Cahais, V., Cuenin, C., Zhivagui, M., Herceg, Z., Ghantous, A., & Zgheib, N. K. (2019). DNA methylome-wide alterations associated with estrogen receptor-dependent effects of bisphenols in breast cancer. *Clinical Epigenetics*, *11*(1), 138. https://doi.org/10.1186/s13148-019-0725-y

Bárez-López, S., Obregon, M. J., Bernal, J., & Guadaño-Ferraz, A. (2018). Thyroid hormone economy in the perinatal mouse brain: Implications for cerebral cortex development. *Cerebral Cortex, 28*(5), 1783–1793. https://doi.org/10.1093/cercor/bhx088

Barkoski, J. M., Busgang, S. A., Bixby, M., Bennett, D., Schmidt, R. J., Barr, D. B., Panuwet, P., Gennings, C., & Hertz-Picciotto, I. (2019). Prenatal phenol and paraben exposures in relation to child neurodevelopment including autism spectrum disorders in the MARBLES study. *Environmental Research*, *179*. https://doi.org/10.1016/j.envres.2019.108719

Beg, M. A., & Sheikh, I. A. (2020). Endocrine disruption: Molecular interactions of environmental bisphenol contaminants with thyroid hormone receptor and thyroxine-binding globulin. *Toxicology* and Industrial Health, 36(5), 322–335. https://doi.org/10.1177/0748233720928165

Ben-Jonathan, N., & Hugo, E. R. (2016). Bisphenols come in different flavors: Is "S" better than "A"? In *Endocrinology* (Vol. 157, Issue 4, pp. 1321–1323). Endocrine Society. https://doi.org/10.1210/en.2016-1120

Berg, D. A., Belnoue, L., Song, H., & Simon, A. (2013). Neurotransmitter-mediated control of neurogenesis in the adult vertebrate brain. In *Development (Cambridge)* (Vol. 140, Issue 12, pp. 2548–2561). Company of Biologists. https://doi.org/10.1242/dev.088005

Bernal, J. (2000). Thyroid Hormones in Brain Development and Function. In *Endotext*. MDText.com, Inc. http://www.ncbi.nlm.nih.gov/pubmed/25905404

Bhat, R. A., Kumar, D., Bhat, S. M., & Sofi, I. R. (2019). *Historical Perspective of Bisphenol A and Phthalates in the Environment and Their Health Effects* (pp. 246–262). https://doi.org/10.4018/978-1-5225-9452-9.ch013

Boudalia, S., & Oudir, M. (2016). Bisphenol-A: Legislation in industrials countries and in Algeria. In *Research Journal of Environmental Toxicology* (Vol. 10, Issue 3, pp. 189–192). Academic Journals Inc. https://doi.org/10.3923/rjet.2016.189.192

Caligioni, C. S. (2009). Assessing reproductive status/stages in mice. *Current Protocols in Neuroscience, APPENDIX*(SUPPL. 48), Appendix. https://doi.org/10.1002/0471142301.nsa04is48

Camacho, L., & Pogribny, I. P. (2017). Epigenetic Effects of Bisphenol A (BPA): A Literature Review in the Context of Human Dietary Exposure. In *Handbook of Nutrition, Diet, and Epigenetics* (pp. 1–20). Springer International Publishing. https://doi.org/10.1007/978-3-319-31143-2_32-1

Cano-Nicolau, J., Vaillant, C., Pellegrini, E., Charlier, T. D., Kah, O., & Coumailleau, P. (2016). Estrogenic effects of several BPA analogs in the developing zebrafish brain. *Frontiers in Neuroscience*, *10*(MAR), 112. https://doi.org/10.3389/fnins.2016.00112

Cantonwine, D. E., Hauser, R., & Meeker, J. D. (2013). Bisphenol A and human reproductive health. In *Expert Review of Obstetrics and Gynecology* (Vol. 8, Issue 4, pp. 329–335). NIH Public Access. https://doi.org/10.1586/17474108.2013.811939

Cao, Jie, Guo, L. H., Wan, B., & Wei, Y. (2011). In vitro fluorescence displacement investigation of thyroxine transport disruption by bisphenol A. *Journal of Environmental Sciences*, *23*(2), 315–321. https://doi.org/10.1016/S1001-0742(10)60408-1

Cao, Jinyan, Rebuli, M. E., Rogers, J., Todd, K. L., Leyrer, S. M., Ferguson, S. A., & Patisaul, H. B. (2013). Prenatal bisphenol a exposure alters sex-specific estrogen receptor expression in the neonatal rat hypothalamus and amygdala. *Toxicological Sciences*, *133*(1), 157–173. https://doi.org/10.1093/toxsci/kft035

Cao, S., Wang, S., Zhao, Y., Wang, L., Ma, Y., Schäffer, A., & Ji, R. (2020). Fate of bisphenol S (BPS) and characterization of non-extractable residues in soil: Insights into persistence of BPS. *Environment International*, *143*, 105908. https://doi.org/10.1016/j.envint.2020.105908

Castelli, M. P., Casti, A., Casu, A., Frau, R., Bortolato, M., Spiga, S., & Ennas, M. G. (2013). Regional distribution of 5α-reductase type 2 in the adult rat brain: An immunohistochemical analysis. *Psychoneuroendocrinology*, *38*(2), 281–293. https://doi.org/10.1016/j.psyneuen.2012.06.008

Castro, B., Sánchez, P., Miranda, M. T., Torres, J. M., & Ortega, E. (2015). Identification of dopamine- and serotonin-related genes modulated by bisphenol A in the prefrontal cortex of male rats. *Chemosphere*, *139*, 235–239. https://doi.org/10.1016/j.chemosphere.2015.06.061

Castro, B., Sánchez, P., Torres, J. M., & Ortega, E. (2015). Bisphenol A, bisphenol F and bisphenol S affect differently 5α-reductase expression and dopamine-serotonin systems in the prefrontal cortex of juvenile female rats. *Environmental Research*, 142, 281–287. https://doi.org/10.1016/j.envres.2015.07.001

Catanese, M. C., & Vandenberg, L. N. (2017). Bisphenol S (BPS) alters maternal behavior and brain in mice exposed during pregnancy/lactation and their daughters. *Endocrinology*, *158*(3), 516–530. https://doi.org/10.1210/en.2016-1723

Chen, C., Ma, Q., Chen, X., Zhong, M., Deng, P., Zhu, G., Zhang, Y., Zhang, L., Yang, Z., Zhang, K., Guo, L., Wang, L., Yu, Z., & Zhou, Z. (2015). Thyroid Hormone-Otx2 Signaling is Required for Embryonic Ventral Midbrain Neural Stem Cells Differentiated into Dopamine Neurons. *Stem Cells and Development*, *24*(15), 1751–1765. https://doi.org/10.1089/scd.2014.0489

Chen, C., Zhou, Z., Zhong, M., Zhang, Y., Li, M., Zhang, L., Qu, M., Yang, J., Wang, Y., & Yu, Z. (2012). Thyroid hormone promotes neuronal differentiation of embryonic neural stem cells by inhibiting STAT3 Signaling through TRa1. *Stem Cells and Development*, *21*(14), 2667–2681. https://doi.org/10.1089/scd.2012.0023

Chen, D., Kannan, K., Tan, H., Zheng, Z., Feng, Y. L., Wu, Y., & Widelka, M. (2016). Bisphenol Analogues Other Than BPA: Environmental Occurrence, Human Exposure, and Toxicity - A Review. In *Environmental Science and Technology* (Vol. 50, Issue 11, pp. 5438–5453). American Chemical Society. https://doi.org/10.1021/acs.est.5b05387

Chen, F., Ying, G. G., Kong, L. X., Wang, L., Zhao, J. L., Zhou, L. J., & Zhang, L. J. (2011). Distribution and accumulation of endocrine-disrupting chemicals and pharmaceuticals in wastewater irrigated soils in Hebei, China. *Environmental Pollution*, *159*(6), 1490–1498. https://doi.org/10.1016/j.envpol.2011.03.016

Chen, S., Chang, Q., Yin, K., He, Q., Deng, Y., Chen, B., Liu, C., Wang, Y., & Wang, L. (2017). Rapid Analysis of Bisphenol A and Its Analogues in Food Packaging Products by Paper Spray Ionization Mass Spectrometry. *Journal of Agricultural and Food Chemistry*, *65*(23), 4859–4865. https://doi.org/10.1021/acs.jafc.7b02061

Cho, J. H., Kim, A. H., Lee, S., Lee, Y., Lee, W. J., Chang, S. C., & Lee, J. (2018). Sensitive neurotoxicity assessment of bisphenol A using double immunocytochemistry of DCX and MAP2. *Archives of Pharmacal Research*, *41*(11), 1098–1107. https://doi.org/10.1007/s12272-018-1077-4

Choi, S., Kim, M. J., Park, Y. J., Kim, S., Choi, K., Cheon, G. J., Cho, Y. H., Jeon, H. L., Yoo, J., & Park, J. (2020). Thyroxine-binding globulin, peripheral deiodinase activity, and thyroid autoantibody status in association of phthalates and phenolic compounds with thyroid hormones in adult population. *Environment International*, *140*, 105783. https://doi.org/10.1016/j.envint.2020.105783

Choi, Y. J., & Lee, L. S. (2017). Aerobic Soil Biodegradation of Bisphenol (BPA) Alternatives Bisphenol S and Bisphenol AF Compared to BPA. *Environmental Science and Technology*, *51*(23), 13698–13704. https://doi.org/10.1021/acs.est.7b03889

Claxton, L. (2007). A review of conflict of interest, competing interest, and bias for toxicologists. *Toxicology and Industrial Health*, *23*(10), 557–571. https://doi.org/10.1177/0748233708089046

Corbel, T., Gayrard, V., Puel, S., Lacroix, M. Z., Berrebi, A., Gil, S., Viguié, C., Toutain, P. L., & Picard-Hagen, N. (2014). Bidirectional placental transfer of Bisphenol A and its main metabolite, Bisphenol A-Glucuronide, in the isolated perfused human placenta. *Reproductive Toxicology*, *47*, 51–58. https://doi.org/10.1016/j.reprotox.2014.06.001

Corrales, J., Kristofco, L. A., Baylor Steele, W., Yates, B. S., Breed, C. S., Spencer Williams, E., & Brooks, B. W. (2015). Global assessment of bisphenol a in the environment: Review and analysis of its occurrence and bioaccumulation. *Dose-Response*, *13*(3). https://doi.org/10.1177/1559325815598308

Coumailleau, P., Pellegrini, E., Adrio, F., Diotel, N., Cano-Nicolau, J., Nasri, A., Vaillant, C., & Kah, O. (2015). Aromatase, estrogen receptors and brain development in fish and amphibians. In *Biochimica*

et Biophysica Acta - Gene Regulatory Mechanisms (Vol. 1849, Issue 2, pp. 152–162). https://doi.org/10.1016/j.bbagrm.2014.07.002

Cristine Viecelli, N., Perazzoli Baldasso, R., do Nascimento Filho, I., & Lusa Manfredini, K. (2014). Occurrence of Bisphenol A in soil and leachate of a municipal landfill: effect of the sample acidification. *Scientia Cum Industria*, 1, 10–14. https://doi.org/10.18226/23185279.v2iss1p10

Czeh, M., Gressens, P., & Kaindl, A. M. (2011). The yin and yang of microglia. In *Developmental Neuroscience* (Vol. 33, Issues 3–4, pp. 199–209). Karger Publishers. https://doi.org/10.1159/000328989

da Silva, B. S., Pietrobon, C. B., Bertasso, I. M., Lopes, B. P., Carvalho, J. C., Peixoto-Silva, N., Santos, T. R., Claudio-Neto, S., Manhães, A. C., Oliveira, E., de Moura, E. G., & Lisboa, P. C. (2019). Short and long-term effects of bisphenol S (BPS) exposure during pregnancy and lactation on plasma lipids, hormones, and behavior in rats. *Environmental Pollution*, *250*, 312–322. https://doi.org/10.1016/j.envpol.2019.03.100

Da Silva, M. M., Gonçalves, C. F. L., Miranda-Alves, L., Fortunato, R. S., Carvalho, D. P., & Ferreira, A. C. F. (2019). Inhibition of Type 1 Iodothyronine Deiodinase by Bisphenol A. *Hormone and Metabolic Research*, *51*(10), 671–677. https://doi.org/10.1055/a-0919-3879

Danzl, E., Sei, K., Soda, S., Ike, M., & Fujita, M. (2009). Biodegradation of bisphenol A, bisphenol F and bisphenol S in seawater. *International Journal of Environmental Research and Public Health*, *6*(4), 1472–1484. https://doi.org/10.3390/ijerph6041472

Denver, R. J., Hu, F., Scanlan, T. S., & Furlow, J. D. (2009). Thyroid hormone receptor subtype specificity for hormone-dependent neurogenesis in Xenopus laevis. *Developmental Biology*, *326*(1), 155–168. https://doi.org/10.1016/j.ydbio.2008.11.005

Derakhshan, A., Philips, E. M., Ghassabian, A., Santos, S., Asimakopoulos, A. G., Kannan, K., Kortenkamp, A., Jaddoe, V. W. V., Trasande, L., Peeters, R. P., & Korevaar, T. I. M. (2021). Association of urinary bisphenols during pregnancy with maternal, cord blood and childhood thyroid function. *Environment International*, *146*, 106160. https://doi.org/10.1016/j.envint.2020.106160

Desai, M., Ferrini, M. G., Han, G., Jellyman, J. K., & Ross, M. G. (2018). In vivo maternal and in vitro BPA exposure effects on hypothalamic neurogenesis and appetite regulators. *Environmental Research*, *164*, 45–52. https://doi.org/10.1016/j.envres.2018.02.011

Ding, B. (2015). Gene expression in maturing neurons: regulatory mechanisms and related neurodevelopmental disorders. *Sheng Li Xue Bao : [Acta Physiologica Sinica]*, *67*(2), 113–133. https://europepmc.org/article/med/25896042

Diotel, N., Vaillant, C., Kah, O., & Pellegrini, E. (2016). Mapping of brain lipid binding protein (Blbp) in the brain of adult zebrafish, co-expression with aromatase B and links with proliferation. *Gene Expression Patterns*, *20*(1), 42–54. https://doi.org/10.1016/j.gep.2015.11.003

Dodson, R. E., Boronow, K. E., Susmann, H., Udesky, J. O., Rodgers, K. M., Weller, D., Woudneh, M., Brody, J. G., & Rudel, R. A. (2020). Consumer behavior and exposure to parabens, bisphenols, triclosan, dichlorophenols, and benzophenone-3: Results from a crowdsourced biomonitoring study. *International Journal of Hygiene and Environmental Health*, 230, 113624. https://doi.org/10.1016/j.ijheh.2020.113624

Dong, H., Yauk, C. L., Rowan-Carroll, A., You, S. H., Zoeller, R. T., Lambert, I., & Wade, M. G. (2009). Identification of thyroid hormone receptor binding sites and target genes using ChIP-on-chip in developing mouse cerebellum. *PLoS ONE*, *4*(2), 4610. https://doi.org/10.1371/journal.pone.0004610

Dong, H., You, S. H., Williams, A., Wade, M. G., Yauk, C. L., & Thomas Zoeller, R. (2015). Transient Maternal Hypothyroxinemia Potentiates the Transcriptional Response to Exogenous Thyroid Hormone in the Fetal Cerebral Cortex before the Onset of Fetal Thyroid Function: A Messenger and MicroRNA Profiling Study. *Cerebral Cortex*, *25*(7), 1735–1745. https://doi.org/10.1093/cercor/bht364

Drobná, Z., Henriksen, A. D., Wolstenholme, J. T., Montiel, C., Lambeth, P. S., Shang, S., Harris, E. P., Zhou, C., Flaws, J. A., Adli, M., & Rissman, E. F. (2018). Transgenerational effects of bisphenol a on gene expression and DNA methylation of imprinted genes in brain. *Endocrinology*, *159*(1), 132–144. https://doi.org/10.1210/en.2017-00730

Dupuis, A., Migeot, V., Cariot, A., Albouy-Llaty, M., Legube, B., & Rabouan, S. (2012). Quantification of bisphenol A, 353-nonylphenol and their chlorinated derivatives in drinking water treatment plants. *Environmental Science and Pollution Research*, *19*(9), 4193–4205. https://doi.org/10.1007/s11356-012-0972-3

ECHA. (2022). The use of bisphenol A and its alternatives in thermal paper in the EU during 2014 - 2022. *ECHA, June 2020*. https://doi.org/10.2823/592282

Elena, U., & Mona Elena, P. (2020). Safety aspects related to the Bisphenol A migration process in packed meat and milk products-a review USAMVB Timisoara "YOUNG PEOPLE AND MULTIDISCIPLINARY RESEARCH IN APPLIED LIFE SCIENCES."

Escrivá, L., Hanberg, A., Zilliacus, J., & Beronius, A. (2019). Assessment of the endocrine disrupting properties of Bisphenol AF according to the EU criteria and ECHA/EFSA guidance. *EFSA Journal*, *17*(S2). https://doi.org/10.2903/j.efsa.2019.e170914

Escrivá, L., Hessel, E., Gustafsson, S., van Spronsen, R., Svanberg, M., & Beronius, A. (2020). A validated search filter for the identification of endocrine disruptors based on the ECHA/EFSA guidance recommendations. *Environment International, 142,* 105828. https://doi.org/10.1016/j.envint.2020.105828

Essner, R. A., Smith, A. G., Jamnik, A. A., Ryba, A. R., Trutner, Z. D., & Carter, M. E. (2017). AgRP neurons can increase food intake during conditions of appetite suppression and inhibit anorexigenic parabrachial neurons. *Journal of Neuroscience*, *37*(36), 8678–8687. https://doi.org/10.1523/JNEUROSCI.0798-17.2017

Fan, Z., Hu, J., An, W., & Yang, M. (2013). Detection and occurrence of chlorinated byproducts of bisphenol a, nonylphenol, and estrogens in drinking water of China: Comparison to the parent compounds. *Environmental Science and Technology*, *47*(19), 10841–10850. https://doi.org/10.1021/es401504a

Fiege, H., Voges, H.-W., Hamamoto, T., Umemura, S., Iwata, T., Miki, H., Fujita, Y., Buysch, H.-J., Garbe, D., & Paulus, W. (2000). Phenol Derivatives. In *Ullmann's Encyclopedia of Industrial Chemistry*. Wiley-VCH Verlag GmbH & Co. KGaA. https://doi.org/10.1002/14356007.a19_313

FitzGerald, R., Loveren, H. Van, Civitella, C., Castoldi, A. F., & Bernasconi, G. (2020). Assessment of new information on Bisphenol S (BPS) submitted in response to the Decision 1 under REACH Regulation (EC) No 1907/2006. *EFSA Supporting Publications*, *17*(4), 1844E. https://doi.org/10.2903/sp.efsa.2020.en-1844

Food Standards Australia New Zealand. (2015). *Regulation and monitoring of BPA*. https://www.foodstandards.gov.au/consumer/chemicals/bpa/pages/regulationandmonitor5377.asp x

Frankland, P. W., & Josselyn, S. A. (2016). Hippocampal neurogenesis and memory clearance. In *Neuropsychopharmacology* (Vol. 41, Issue 1, pp. 382–383). Nature Publishing Group. https://doi.org/10.1038/npp.2015.243

Frankowski, R., Zgoła-Grześkowiak, A., Smułek, W., & Grześkowiak, T. (2020). Removal of Bisphenol A and Its Potential Substitutes by Biodegradation. *Applied Biochemistry and Biotechnology*, *191*(3), 1100–1110. https://doi.org/10.1007/s12010-020-03247-4

Fraser, T. W. K., Khezri, A., Jusdado, J. G. H., Lewandowska-Sabat, A. M., Henry, T., & Ropstad, E. (2017). Toxicant induced behavioural aberrations in larval zebrafish are dependent on minor methodological alterations. *Toxicology Letters*, 276, 62–68. https://doi.org/10.1016/j.toxlet.2017.05.021

Freitas, J., Cano, P., Craig-Veit, C., Goodson, M. L., David Furlow, J., & Murk, A. J. (2011). Detection of thyroid hormone receptor disruptors by a novel stable in vitro reporter gene assay. *Toxicology in Vitro*, *25*(1), 257–266. https://doi.org/10.1016/j.tiv.2010.08.013

Fu, J., Guo, Y., Yang, L., Han, J., & Zhou, B. (2020). Nano-TiO2 enhanced bioaccumulation and developmental neurotoxicity of bisphenol a in zebrafish larvae. *Environmental Research*, *187*, 109682–109682. https://doi.org/10.1016/j.envres.2020.109682

Fu, P., & Kawamura, K. (2010). Ubiquity of bisphenol A in the atmosphere. *Environmental Pollution*, *158*(10), 3138–3143. https://doi.org/10.1016/j.envpol.2010.06.040

Geens, T., Goeyens, L., & Covaci, A. (2011). Are potential sources for human exposure to bisphenol-A overlooked? In *International Journal of Hygiene and Environmental Health* (Vol. 214, Issue 5, pp. 339–347). https://doi.org/10.1016/j.ijheh.2011.04.005

Geneaid. (2008). GENEzol[™] Reagent For research use only. *GENEzol[™] Reagent[™] Reagent*, 1–4. www.geneaid.com

Gentilcore, D., Porreca, I., Rizzo, F., Ganbaatar, E., Carchia, E., Mallardo, M., De Felice, M., & Ambrosino, C. (2013). Bisphenol A interferes with thyroid specific gene expression. *Toxicology*, *304*, 21–31. https://doi.org/10.1016/j.tox.2012.12.001

Gerona, R. R., Woodruff, T. J., Dickenson, C. A., Pan, J., Schwartz, J. M., Sen, S., Friesen, M. W., Fujimoto, V. Y., & Hunt, P. A. (2013). Bisphenol-A (BPA), BPA glucuronide, and BPA sulfate in midgestation umbilical cord serum in a northern and central california population. *Environmental Science and Technology*, *47*(21), 12477–12485. https://doi.org/10.1021/es402764d

Gibson, R., Durán-Álvarez, J. C., Estrada, K. L., Chávez, A., & Jiménez Cisneros, B. (2010). Accumulation and leaching potential of some pharmaceuticals and potential endocrine disruptors in soils irrigated with wastewater in the Tula Valley, Mexico. *Chemosphere*, *81*(11), 1437–1445. https://doi.org/10.1016/j.chemosphere.2010.09.006

Gilbert, M. E., Rovet, J., Chen, Z., & Koibuchi, N. (2012). Developmental thyroid hormone disruption: Prevalence, environmental contaminants and neurodevelopmental consequences. *NeuroToxicology*, *33*(4), 842–852. https://doi.org/10.1016/j.neuro.2011.11.005

Goasdoué, K., Miller, S. M., Colditz, P. B., & Björkman, S. T. (2017). Review: The blood-brain barrier; protecting the developing fetal brain. In *Placenta* (Vol. 54, pp. 111–116). W.B. Saunders Ltd. https://doi.org/10.1016/j.placenta.2016.12.005

González, N., Cunha, S. C., Ferreira, R., Fernandes, J. O., Marquès, M., Nadal, M., & Domingo, J. L. (2020). Concentrations of nine bisphenol analogues in food purchased from Catalonia (Spain): Comparison of canned and non-canned foodstuffs. *Food and Chemical Toxicology*, *136*, 110992. https://doi.org/10.1016/j.fct.2019.110992

Goodman, J. E., & Peterson, M. K. (2014). Bisphenol A. In *Encyclopedia of Toxicology: Third Edition* (pp. 514–518). Elsevier. https://doi.org/10.1016/B978-0-12-386454-3.00366-3

Graziani, N. S., Carreras, H., & Wannaz, E. (2019). Atmospheric levels of BPA associated with particulate matter in an urban environment. *Heliyon*, *5*(4). https://doi.org/10.1016/j.heliyon.2019.e01419

Grignard, E., Lapenna, S., & Bremer, S. (2012). Weak estrogenic transcriptional activities of Bisphenol A and Bisphenol S. *Toxicology in Vitro*, *26*(5), 727–731. https://doi.org/10.1016/j.tiv.2012.03.013

Guo, Y., Chen, L., Wu, J., Hua, J., Yang, L., Wang, Q., Zhang, W., Lee, J. S., & Zhou, B. (2019). Parental co-exposure to bisphenol A and nano-TiO 2 causes thyroid endocrine disruption and developmental neurotoxicity in zebrafish offspring. *Science of the Total Environment*, *650*, 557–565. https://doi.org/10.1016/j.scitotenv.2018.09.007

Gupta, R. K., & Gupta, R. C. (2017). Placental toxicity. In *Reproductive and Developmental Toxicology* (pp. 1301–1325). Elsevier. https://doi.org/10.1016/B978-0-12-804239-7.00068-8

Gyimah, E., Xu, H., Dong, X., Qiu, X., Zhang, Z., Bu, Y., & Akoto, O. (2020). Developmental neurotoxicity of low concentrations of bisphenol A and S exposure in zebrafish. *Chemosphere*, *262*. https://doi.org/10.1016/j.chemosphere.2020.128045

Higgins, J. P. T., & Green, S. (2019). Cochrane Handbook for Systematic Reviews of Interventions. In *Cochrane Handbook for Systematic Reviews of Interventions*. https://doi.org/10.1002/9781119536604

Howdeshell, K. L. (2002). A model of the development of the brain as a construct of the thyroid system. In *Environmental Health Perspectives* (Vol. 110, Issue SUPPL. 3, pp. 337–348). Public Health Services, US Dept of Health and Human Services. https://doi.org/10.1289/ehp.02110s3337

Huang, W., Zhao, C., Zhong, H., Zhang, S., Xia, Y., & Cai, Z. (2019). Bisphenol S induced epigenetic and transcriptional changes in human breast cancer cell line MCF-7. *Environmental Pollution*, *246*, 697–703. https://doi.org/10.1016/j.envpol.2018.12.084

Huang, Y. Q., Wong, C. K. C., Zheng, J. S., Bouwman, H., Barra, R., Wahlström, B., Neretin, L., & Wong, M. H. (2012). Bisphenol A (BPA) in China: A review of sources, environmental levels, and potential human health impacts. *Environment International*, 42(1), 91–99. https://doi.org/10.1016/j.envint.2011.04.010

Hufnagel, R. B., Le, T. T., Riesenberg, A. L., & Brown, N. L. (2010). Neurog2 controls the leading edge of neurogenesis in the mammalian retina. *Developmental Biology*, *340*(2), 490–503. https://doi.org/10.1016/j.ydbio.2010.02.002

Huo, X., Chen, D., He, Y., Zhu, W., Zhou, W., & Zhang, J. (2015). Bisphenol-a and female infertility: A possible role of gene-environment interactions. In *International Journal of Environmental Research and Public Health* (Vol. 12, Issue 9, pp. 11101–11116). MDPI AG. https://doi.org/10.3390/ijerph120911101

Ideta-Otsuka, M., Igarashi, K., Narita, M., & Hirabayashi, Y. (2017). Epigenetic toxicity of environmental chemicals upon exposure during development - Bisphenol A and valproic acid may have epigenetic effects. *Food and Chemical Toxicology*, *109*, 812–816. https://doi.org/10.1016/j.fct.2017.09.014

Ike, M., Chen, M. Y., Danzl, E., Sei, K., & Fujita, M. (2006). Biodegradation of a variety of bisphenols under aerobic and anaerobic conditions. *Water Science and Technology*, *53*(6), 153–159. https://doi.org/10.2166/wst.2006.189

Inadera, H. (2015). Neurological effects of bisphenol A and its analogues. In *International Journal of Medical Sciences* (Vol. 12, Issue 12, pp. 926–936). Ivyspring International Publisher. https://doi.org/10.7150/ijms.13267

Industry Experts. (2017). Bisphenol-A – A Global Market Overview. *Industry Experts*, 1–194. www.hexion.com

Itoh, K., Yaoi, T., & Fushiki, S. (2012). Bisphenol A, an endocrine-disrupting chemical, and brain development. *Neuropathology*, *32*(4), 447–457. https://doi.org/10.1111/j.1440-1789.2011.01287.x

Iwamuro, S., Yamada, M., Kato, M., & Kikuyama, S. (2006). Effects of bisphenol A on thyroid hormone-dependent up-regulation of thyroid hormone receptor α and β and down-regulation of retinoid X receptor γ in Xenopus tail culture. *Life Sciences*, *79*(23), 2165–2171. https://doi.org/10.1016/j.lfs.2006.07.013

Jansson, L. C., & Åkerman, K. E. (2014). The role of glutamate and its receptors in the proliferation, migration, differentiation and survival of neural progenitor cells. *Journal of Neural Transmission*, *121*(8), 819–836. https://doi.org/10.1007/s00702-014-1174-6

Jiang, Y., Li, J., Xu, S., Zhou, Y., Zhao, H., Li, Y., Xiong, C., Sun, X., Liu, H., Liu, W., Peng, Y., Hu, C., Cai, Z., & Xia, W. (2020). Prenatal exposure to bisphenol A and its alternatives and child neurodevelopment at 2 years. *Journal of Hazardous Materials, 388*. https://doi.org/10.1016/j.jhazmat.2019.121774

Jin, F. L., Li, X., & Park, S. J. (2015). Synthesis and application of epoxy resins: A review. In *Journal* of *Industrial and Engineering Chemistry* (Vol. 29, pp. 1–11). Korean Society of Industrial Engineering Chemistry. https://doi.org/10.1016/j.jiec.2015.03.026

Jin, H., Xie, J., Mao, L., Zhao, M., Bai, X., Wen, J., Shen, T., & Wu, P. (2020). Bisphenol analogue concentrations in human breast milk and their associations with postnatal infant growth. *Environmental Pollution*, *259*, 113779. https://doi.org/10.1016/j.envpol.2019.113779

Jin, H., & Zhu, L. (2016). Occurrence and partitioning of bisphenol analogues in water and sediment from Liaohe River Basin and Taihu Lake, China. *Water Research*, *103*, 343–351. https://doi.org/10.1016/j.watres.2016.07.059

Jocsak, G., Kiss, D. S., Toth, I., Goszleth, G., Bartha, T., Frenyo, L. V., Horvath, T. L., & Zsarnovszky, A. (2016). Comparison of individual and combined effects of four endocrine disruptors on estrogen receptor beta transcription in cerebellar cell culture: The modulatory role of estradiol and triiodo-thyronine. *International Journal of Environmental Research and Public Health*, *13*(6), 1–14. https://doi.org/10.3390/ijerph13060619

Joëls, M. (2018). Corticosteroids and the brain. In *Journal of Endocrinology* (Vol. 238, Issue 3, pp. R121–R130). BioScientifica Ltd. https://doi.org/10.1530/JOE-18-0226

Jurkowski, M. P., Bettio, L., K. Woo, E., Patten, A., Yau, S. Y., & Gil-Mohapel, J. (2020). Beyond the Hippocampus and the SVZ: Adult Neurogenesis Throughout the Brain. In *Frontiers in Cellular Neuroscience* (Vol. 14, p. 293). Frontiers Media S.A. https://doi.org/10.3389/fncel.2020.576444

Kapoor, R., Fanibunda, S. E., Desouza, L. A., Guha, S. K., & Vaidya, V. A. (2015). Perspectives on thyroid hormone action in adult neurogenesis. In *Journal of Neurochemistry* (Vol. 133, Issue 5, pp. 599–616). Blackwell Publishing Ltd. https://doi.org/10.1111/jnc.13093

Kellermeyer, L., Harnke, B., & Knight, S. (2018). Covidence and Rayyan. *Journal of the Medical Library Association*, *106*(4), 580. https://doi.org/10.5195/jmla.2018.513

Khmiri, I., Côté, J., Mantha, M., Khemiri, R., Lacroix, M., Gely, C., Toutain, P. L., Picard-Hagen, N., Gayrard, V., & Bouchard, M. (2020). Toxicokinetics of bisphenol-S and its glucuronide in plasma and urine following oral and dermal exposure in volunteers for the interpretation of biomonitoring data. *Environment International*, *138*, 105644. https://doi.org/10.1016/j.envint.2020.105644

Kim, J. D., Leyva, S., & Diano, S. (2014). Hormonal regulation of the hypothalamic melanocortin system. In *Frontiers in Physiology* (Vol. 5, Issue Nov, p. 480). Frontiers Research Foundation. https://doi.org/10.3389/fphys.2014.00480

Kim, J. J., Kumar, S., Kumar, V., Lee, Y. M., Kim, Y. S., & Kumar, V. (2020). Bisphenols as a legacy pollutant, and their effects on organ vulnerability. In *International Journal of Environmental Research and Public Health* (Vol. 17, Issue 1). MDPI AG. https://doi.org/10.3390/ijerph17010112

Kim, K. Y., Lee, E., & Kim, Y. (2019). The association between bisphenol a exposure and obesity in children—a systematic review with meta-analysis. *International Journal of Environmental Research and Public Health*, *16*(14). https://doi.org/10.3390/ijerph16142521

Kinch, C. D., Ibhazehiebo, K., Jeong, J. H., Habibi, H. R., & Kurrasch, D. M. (2015). Low-dose exposure to bisphenol a and replacement bisphenol S induces precocious hypothalamic neurogenesis in embryonic zebrafish. *Proceedings of the National Academy of Sciences of the United States of America*, *112*(5), 1475–1480. https://doi.org/10.1073/pnas.1417731112

Kishi, R. (2020). Impacts of Developmental Exposure to Environmental Chemicals on Human Health with Global Perspectives (R. Kishi & P. Grandjean (eds.); pp. 3–22). Springer Singapore. https://doi.org/10.1007/978-981-15-0520-1_1

Kleywegt, S., Pileggi, V., Yang, P., Hao, C., Zhao, X., Rocks, C., Thach, S., Cheung, P., & Whitehead, B. (2011). Pharmaceuticals, hormones and bisphenol A in untreated source and finished drinking water in Ontario, Canada - Occurrence and treatment efficiency. *Science of the Total Environment, 409*(8), 1481–1488. https://doi.org/10.1016/j.scitotenv.2011.01.010

Komada, M., Asai, Y., Morii, M., Matsuki, M., Sato, M., & Nagao, T. (2012). Maternal bisphenol A oral dosing relates to the acceleration of neurogenesis in the developing neocortex of mouse fetuses. *Toxicology*, *295*(1–3), 31–38. https://doi.org/10.1016/j.tox.2012.02.013

Krishnan, A. V., Stathis, P., Permuth, S. F., Tokes, L., & Feldman, D. (1993). Bisphenol-a: An estrogenic substance is released from polycarbonate flasks during autoclaving. *Endocrinology*, *132*(6), 2279–2286. https://doi.org/10.1210/endo.132.6.8504731

Kundakovic, M., & Champagne, F. A. (2011). Epigenetic perspective on the developmental effects of bisphenol A. In *Brain, Behavior, and Immunity* (Vol. 25, Issue 6, pp. 1084–1093). https://doi.org/10.1016/j.bbi.2011.02.005

Kundakovic, M., Gudsnuk, K., Franks, B., Madrid, J., Miller, R. L., Perera, F. P., & Champagne, F. A. (2013). Sex-specific epigenetic disruption and behavioral changes following low-dose in utero bisphenol a exposure. *Proceedings of the National Academy of Sciences of the United States of America*, *110*(24), 9956–9961. https://doi.org/10.1073/pnas.1214056110

Kunz, N., Camm, E. J., Somm, E., Lodygensky, G., Darbre, S., Aubert, M. L., Hüppi, P. S., Sizonenko, S. V., & Gruetter, R. (2011). Developmental and metabolic brain alterations in rats exposed to bisphenol A during gestation and lactation. *International Journal of Developmental Neuroscience*, *29*(1), 37–43. https://doi.org/10.1016/j.ijdevneu.2010.09.009

Lagarde, F., Beausoleil, C., Belcher, S. M., Belzunces, L. P., Emond, C., Guerbet, M., & Rousselle, C. (2015). Non-monotonic dose-response relationships and endocrine disruptors: A qualitative method of assessment -No section-. In *Environmental Health: A Global Access Science Source* (Vol. 14, Issue 1). BioMed Central Ltd. https://doi.org/10.1186/1476-069X-14-13

Landis, S. C., Amara, S. G., Asadullah, K., Austin, C. P., Blumenstein, R., Bradley, E. W., Crystal, R. G., Darnell, R. B., Ferrante, R. J., Fillit, H., Finkelstein, R., Fisher, M., Gendelman, H. E., Golub, R. M., Goudreau, J. L., Gross, R. A., Gubitz, A. K., Hesterlee, S. E., Howells, D. W., ... Silberberg, S. D. (2012). A call for transparent reporting to optimize the predictive value of preclinical research. In *Nature* (Vol. 490, Issue 7419, pp. 187–191). https://doi.org/10.1038/nature11556

Lee, J., Choi, K., Park, J., Moon, H. B., Choi, G., Lee, J. J., Suh, E., Kim, H. J., Eun, S. H., Kim, G. H., Cho, G. J., Kim, S. K., Kim, S., Kim, S. Y., Kim, S., Eom, S., Choi, S., Kim, Y. D., & Kim, S. (2018). Bisphenol A distribution in serum, urine, placenta, breast milk, and umbilical cord serum in a birth panel of mother–neonate pairs. *Science of the Total Environment, 626*, 1494–1501. https://doi.org/10.1016/j.scitotenv.2017.10.042

Lee, Sangwoo, Kim, C., Shin, H., Kho, Y., & Choi, K. (2019). Comparison of thyroid hormone disruption potentials by bisphenols A, S, F, and Z in embryo-larval zebrafish. *Chemosphere*, 221, 115–123. https://doi.org/10.1016/j.chemosphere.2019.01.019

Lee, Sunggyu, Liao, C., Song, G. J., Ra, K., Kannan, K., & Moon, H. B. (2015). Emission of bisphenol analogues including bisphenol A and bisphenol F from wastewater treatment plants in Korea. *Chemosphere*, *119*, 1000–1006. https://doi.org/10.1016/j.chemosphere.2014.09.011

Lehmler, H. J., Liu, B., Gadogbe, M., & Bao, W. (2018). Exposure to Bisphenol A, Bisphenol F, and Bisphenol S in U.S. Adults and Children: The National Health and Nutrition Examination Survey 2013-2014. *ACS Omega*, *3*(6), 6523–6532. https://doi.org/10.1021/acsomega.8b00824

Lenz, K. M., & Nelson, L. H. (2018). Microglia and beyond: Innate immune cells as regulators of brain development and behavioral function. In *Frontiers in Immunology* (Vol. 9, Issue APR, p. 1). Frontiers Media S.A. https://doi.org/10.3389/fimmu.2018.00698

Liang, X., Yin, N., Liang, S., Yang, R., Liu, S., Lu, Y., Jiang, L., Zhou, Q., Jiang, G., & Faiola, F. (2020). Bisphenol A and several derivatives exert neural toxicity in human neuron-like cells by decreasing neurite length. *Food and Chemical Toxicology*, *135*. https://doi.org/10.1016/j.fct.2019.111015

Liao, C., & Kannan, K. (2012). Determination of free and conjugated forms of bisphenol A in human urine and serum by liquid chromatography-tandem mass spectrometry. *Environmental Science and Technology*, *46*(9), 5003–5009. https://doi.org/10.1021/es300115a

Liao, C., & Kannan, K. (2014). A survey of alkylphenols, bisphenols, and triclosan in personal care products from China and the United States. *Archives of Environmental Contamination and Toxicology*, *67*(1), 50–59. https://doi.org/10.1007/s00244-014-0016-8

Liao, C., Liu, F., Moon, H. B., Yamashita, N., Yun, S., & Kannan, K. (2012). Bisphenol analogues in sediments from industrialized areas in the United States, Japan, and Korea: Spatial and temporal distributions. *Environmental Science and Technology*, *46*(21), 11558–11565. https://doi.org/10.1021/es303191g

Liberati, A., Altman, D. G., Tetzlaff, J., Mulrow, C., Gøtzsche, P. C., Ioannidis, J. P. A., Clarke, M., Devereaux, P. J., Kleijnen, J., & Moher, D. (2009). *Guidelines and Guidance The PRISMA Statement for Reporting Systematic Reviews and Meta-Analyses of Studies That Evaluate Health Care Interventions: Explanation and Elaboration*. https://doi.org/10.1371/journal.pmed.1000100

Lin, Z., Wang, L., Jia, Y., Zhang, Y., Dong, Q., & Huang, C. (2017). A Study on Environmental Bisphenol A Pollution in Plastics Industry Areas. *Water, Air, and Soil Pollution, 228*(3). https://doi.org/10.1007/s11270-017-3277-9

Little, A. G. (2016). A review of the peripheral levels of regulation by thyroid hormone. In *Journal of Comparative Physiology B: Biochemical, Systemic, and Environmental Physiology* (Vol. 186, Issue 6, pp. 677–688). Springer Verlag. https://doi.org/10.1007/s00360-016-0984-2

Liu, J., Li, J., Wu, Y., Zhao, Y., Luo, F., Li, S., Yang, L., Moez, E. K., Dinu, I., & Martin, J. W. (2017). Bisphenol A Metabolites and Bisphenol S in Paired Maternal and Cord Serum. *Environmental Science and Technology*, *51*(4), 2456–2463. https://doi.org/10.1021/acs.est.6b05718

Liu, M. L., Zang, T., Zou, Y., Chang, J. C., Gibson, J. R., Huber, K. M., & Zhang, C. L. (2013). Small molecules enable neurogenin 2 to efficiently convert human fibroblasts into cholinergic neurons. *Nature Communications*, *4*(1), 1–10. https://doi.org/10.1038/ncomms3183

Liu, R., Xing, L., Kong, D., Jiang, J., Shang, L., & Hao, W. (2013). Bisphenol A inhibits proliferation and induces apoptosis in micromass cultures of rat embryonic midbrain cells through the JNK, CREB and p53 signaling pathways. *Food and Chemical Toxicology*, *52*, 76–82. https://doi.org/10.1016/j.fct.2012.10.033

Liu, X., & Shi, H. (2015). Regulation of Estrogen Receptor Expression in the Hypothalamus by Sex Steroids: Implication in the Regulation of Energy Homeostasis. In *International Journal of Endocrinology* (Vol. 2015). Hindawi Publishing Corporation. https://doi.org/10.1155/2015/949085 Liu, Y., Zhang, S., Song, N., Guo, R., Chen, M., Mai, D., Yan, Z., Han, Z., & Chen, J. (2017). Occurrence, distribution and sources of bisphenol analogues in a shallow Chinese freshwater lake (Taihu Lake): Implications for ecological and human health risk. *Science of the Total Environment*, *599–600*, 1090–1098. https://doi.org/10.1016/j.scitotenv.2017.05.069

Logroscino, G., Traynor, B. J., Hardiman, O., Chió, A., Mitchell, D., Swingler, R. J., Millul, A., Benn, E., & Beghi, E. (2010). Incidence of amyotrophic lateral sclerosis in Europe. *Journal of Neurology, Neurosurgery and Psychiatry*, *81*(4), 385–390. https://doi.org/10.1136/jnnp.2009.183525

Lu, L., Zhan, T., Ma, M., Xu, C., Wang, J., Zhang, C., Liu, W., & Zhuang, S. (2018). Thyroid Disruption by Bisphenol S Analogues via Thyroid Hormone Receptor β: in Vitro, in Vivo, and Molecular Dynamics Simulation Study. *Environmental Science and Technology*, *52*(11), 6617–6625. https://doi.org/10.1021/acs.est.8b00776

Lukasiewicz, M., Czerniecki, J., Ponikwicka-Tyszko, D., Sztachelska, M., Hryniewicka, M., Nalewajko-Sieliwoniuk, E., Wiczkowski, W., Banaszewska, B., Milewski, R., Toppari, J., Huhtaniemi, I., Rahman, N. A., & Wolczynski, S. (2020). Placenta is Capable of Protecting the Male Fetus from Exposure to Environmental Bisphenol A. *Exposure and Health*, *13*(1), 1–14. https://doi.org/10.1007/s12403-020-00358-5

Malloy, M. A., Kochmanski, J. J., Jones, T. R., Colacino, J. A., Goodrich, J. M., Dolinoy, D. C., & Svoboda, L. K. (2019). Perinatal Bisphenol A Exposure and Reprogramming of Imprinted Gene Expression in the Adult Mouse Brain. *Frontiers in Genetics, 10*. https://doi.org/10.3389/fgene.2019.00951

Mandel, N. D., Gamboa-Loira, B., Cebrián, M. E., Mérida-Ortega, Á., & López-Carrillo, L. (2019). Challenges to regulate products containing bisphenol A: Implications for policy. *Salud Publica de Mexico*, *61*(5), 692–697. https://doi.org/10.21149/10411

Manjourides, J., Zimmerman, E., Watkins, D. J., Carpenito, T., Vélez-Vega, C. M., Huerta-Montañez, G., Rosario, Z., Ayala, I., Vergara, C., Feric, Z., Ondras, M., Suh, H. H., Gu, A. Z., Brown, P., Cordero, J. F., Meeker, J. D., & Alshawabkeh, A. (2020). Cohort profile: Center for Research on Early Childhood Exposure and Development in Puerto Rico. *BMJ Open*, *10*, 36389. https://doi.org/10.1136/bmjopen-2019-036389

Maragou, N. C., Thomaidis, N. S., Theodoridis, G. A., Lampi, E. N., & Koupparis, M. A. (2020). Determination of bisphenol A in canned food by microwave assisted extraction, molecularly imprinted polymer-solid phase extraction and liquid chromatography-mass spectrometry. *Journal of Chromatography B: Analytical Technologies in the Biomedical and Life Sciences*, *1137*, 121938. https://doi.org/10.1016/j.jchromb.2019.121938

Marek, R., Strobel, C., Bredy, T. W., & Sah, P. (2013). The amygdala and medial prefrontal cortex: Partners in the fear circuit. In *Journal of Physiology* (Vol. 591, Issue 10, pp. 2381–2391). Wiley-Blackwell. https://doi.org/10.1113/jphysiol.2012.248575

Marty, S., Beekhuijzen, M., Charlton, A., Hallmark, N., Hannas, B. R., Jacobi, S., Melching-Kollmuss, S., Sauer, U. G., Sheets, L. P., Strauss, V., Urbisch, D., Botham, P. A., & van Ravenzwaay, B. (2021).

Towards a science-based testing strategy to identify maternal thyroid hormone imbalance and neurodevelopmental effects in the progeny–part II: how can key events of relevant adverse outcome pathways be addressed in toxicological assessments? In *Critical Reviews in Toxicology* (Vol. 51, Issue 4, pp. 328–358). Taylor and Francis Ltd. https://doi.org/10.1080/10408444.2021.1910625

Mathisen, G. H., Yazdani, M., Rakkestad, K. E., Aden, P. K., Bodin, J., Samuelsen, M., Nygaard, U. C., Goverud, I. L., Gaarder, M., Løberg, E. M., Bølling, A. K., Becher, R., & Paulsen, R. E. (2013). Prenatal exposure to bisphenol A interferes with the development of cerebellar granule neurons in mice and chicken. *International Journal of Developmental Neuroscience*, *31*(8), 762–769. https://doi.org/10.1016/j.ijdevneu.2013.09.009

Matta, K., Ploteau, S., Coumoul, X., Koual, M., Le Bizec, B., Antignac, J. P., & Cano-Sancho, G. (2019). Associations between exposure to organochlorine chemicals and endometriosis in experimental studies: A systematic review protocol. In *Environment International* (Vol. 124, pp. 400–407). Elsevier Ltd. https://doi.org/10.1016/j.envint.2018.12.063

Matuszczak, E., Komarowska, M. D., Debek, W., & Hermanowicz, A. (2019). The Impact of Bisphenol A on Fertility, Reproductive System, and Development: A Review of the Literature. In *International Journal of Endocrinology* (Vol. 2019). Hindawi Limited. https://doi.org/10.1155/2019/4068717

McCaffrey, K. A., Jones, B., Mabrey, N., Weiss, B., Swan, S. H., & Patisaul, H. B. (2013). Sex specific impact of perinatal bisphenol A (BPA) exposure over a range of orally administered doses on rat hypothalamic sexual differentiation. *NeuroToxicology*, *36*, 55–62. https://doi.org/10.1016/j.neuro.2013.03.001

McCarthy, M. M. (2008). Estradiol and the developing brain. In *Physiological Reviews* (Vol. 88, Issue 1, pp. 91–124). NIH Public Access. https://doi.org/10.1152/physrev.00010.2007

McKlveen, J. M., Myers, B., Flak, J. N., Bundzikova, J., Solomon, M. B., Seroogy, K. B., & Herman, J. P. (2013). Role of prefrontal cortex glucocorticoid receptors in stress and emotion. *Biological Psychiatry*, 74(9), 672–679. https://doi.org/10.1016/j.biopsych.2013.03.024

Melmed, S. (2017). Hypothalamic-Pituitary Regulation. In *Conn's Translational Neuroscience* (pp. 317–331). Elsevier Inc. https://doi.org/10.1016/B978-0-12-802381-5.00025-7

Mileva, G., Baker, S. L., Konkle, A. T. M., & Bielajew, C. (2014). Bisphenol-A: Epigenetic reprogramming and effects on reproduction and behavior. In *International Journal of Environmental Research and Public Health* (Vol. 11, Issue 7, pp. 7537–7561). MDPI AG. https://doi.org/10.3390/ijerph110707537

Mohan, V., Sinha, R. A., Pathak, A., Rastogi, L., Kumar, P., Pal, A., & Godbole, M. M. (2012). Maternal thyroid hormone deficiency affects the fetal neocorticogenesis by reducing the proliferating pool, rate of neurogenesis and indirect neurogenesis. *Experimental Neurology*, *237*(2), 477–488. https://doi.org/10.1016/j.expneurol.2012.07.019

Molina-Molina, J. M., Jiménez-Díaz, I., Fernández, M. F., Rodriguez-Carrillo, A., Peinado, F. M., Mustieles, V., Barouki, R., Piccoli, C., Olea, N., & Freire, C. (2019). Determination of bisphenol A and bisphenol S concentrations and assessment of estrogen- and anti-androgen-like activities in thermal paper receipts from Brazil, France, and Spain. *Environmental Research*, *170*, 406–415. https://doi.org/10.1016/j.envres.2018.12.046

Molina-Molina, José Manuel, Amaya, E., Grimaldi, M., Sáenz, J. M., Real, M., Fernández, M. F., Balaguer, P., & Olea, N. (2013). In vitro study on the agonistic and antagonistic activities of bisphenol-S and other bisphenol-A congeners and derivatives via nuclear receptors. *Toxicology and Applied Pharmacology*, *272*(1), 127–136. https://doi.org/10.1016/j.taap.2013.05.015

Morgan, M. K., Nash, M., Barr, D. B., Starr, J. M., Scott Clifton, M., & Sobus, J. R. (2018). Distribution, variability, and predictors of urinary bisphenol A levels in 50 North Carolina adults over a six-week monitoring period. *Environment International*, *112*, 85–99. https://doi.org/10.1016/j.envint.2017.12.014

Morgan, R. L., Whaley, P., Thayer, K. A., & Schünemann, H. J. (2018). Identifying the PECO: A framework for formulating good questions to explore the association of environmental and other exposures with health outcomes. In *Environment International* (Vol. 121, pp. 1027–1031). Elsevier Ltd. https://doi.org/10.1016/j.envint.2018.07.015

Moriyama, K., Tagami, T., Akamizu, T., Usui, T., Saijo, M., Kanamoto, N., Hataya, Y., Shimatsu, A., Kuzuya, H., & Nakao, K. (2002). Thyroid Hormone Action Is Disrupted by Bisphenol A as an Antagonist. *The Journal of Clinical Endocrinology & Metabolism*, *87*(11), 5185–5190. https://doi.org/10.1210/jc.2002-020209

Morrice, J. R., Gregory-Evans, C. Y., & Shaw, C. A. (2018). Modeling Environmentally-Induced Motor Neuron Degeneration in Zebrafish. *Scientific Reports, 8*(1). https://doi.org/10.1038/s41598-018-23018-w

Mouriec, K., Gueguen, M. M., Manuel, C., Percevault, F., Thieulant, M. L., Pakdel, F., & Kah, O. (2009). Androgens upregulate cyp19a1b (aromatase B) gene expression in the brain of zebrafish (Danio rerio) through estrogen receptors. *Biology of Reproduction*, *80*(5), 889–896. https://doi.org/10.1095/biolreprod.108.073643

Mueller, J. K., & Heger, S. (2014). Endocrine disrupting chemicals affect the Gonadotropin releasing hormone neuronal network. *Reproductive Toxicology*, *44*, 73–84. https://doi.org/10.1016/j.reprotox.2013.10.011

Mughal, B. B., Fini, J. B., & Demeneix, B. A. (2018). Thyroid-disrupting chemicals and brain development: An update. In *Endocrine Connections* (Vol. 7, Issue 4, pp. R160–R186). Oxford Academic. https://doi.org/10.1530/EC-18-0029

Mustieles, V., Williams, P. L., Fernandez, M., Mnguez-Alarcn, L., Ford, J. B., Calafat, A. M., Hauser, R., & Messerlian, C. (2018). Maternal and paternal preconception exposure to bisphenols and size at birth. *Human Reproduction*, *33*(8), 1528–1537. https://doi.org/10.1093/humrep/dey234

Naderi, M., & Kwong, R. W. M. (2020). A comprehensive review of the neurobehavioral effects of bisphenol S and the mechanisms of action: New insights from in vitro and in vivo models. In *Environmental International* (Vol. 145, p. 106078). Elsevier Ltd. https://doi.org/10.1016/j.envint.2020.106078

Naderi, M., Wong, M. Y. L., & Gholami, F. (2014). Developmental exposure of zebrafish (Danio rerio) to bisphenol-S impairs subsequent reproduction potential and hormonal balance in adults. *Aquatic Toxicology*, *148*, 195–203. https://doi.org/10.1016/j.aquatox.2014.01.009

Nakamura, K., Itoh, K., Yaoi, T., Fujiwara, Y., Sugimoto, T., & Fushiki, S. (2006). Murine neocortical histogenesis is perturbed by prenatal exposure to low doses of bisphenol A. *Journal of Neuroscience Research*, *84*(6), 1197–1205. https://doi.org/10.1002/jnr.21020

Nesan, D., Sewell, L. C., & Kurrasch, D. M. (2018). Opening the black box of endocrine disruption of brain development: Lessons from the characterization of Bisphenol A. In *Hormones and Behavior* (Vol. 101, pp. 50–58). Academic Press Inc. https://doi.org/10.1016/j.yhbeh.2017.12.001

Nishikawa, M., Iwano, H., Yanagisawa, R., Koike, N., Inoue, H., & Yokota, H. (2010). Placental transfer of conjugated bisphenol A and subsequent reactivation in the rat fetus. *Environmental Health Perspectives*, *118*(9), 1196–1203. https://doi.org/10.1289/ehp.0901575

Noda, M. (2015). Possible role of glial cells in the relationship between thyroid dysfunction and mental disorders. In *Frontiers in Cellular Neuroscience* (Vol. 9, Issue JUNE). Frontiers Media S.A. https://doi.org/10.3389/fncel.2015.00194

Nowicki, B. A., Hamada, M. A., Robinson, G. Y., & Jones, D. C. (2016). Adverse effects of bisphenol A (BPA) on the dopamine system in two distinct cell models and corpus striatum of the Sprague-Dawley rat. *Journal of Toxicology and Environmental Health - Part A: Current Issues, 79*(20), 912–924. https://doi.org/10.1080/15287394.2016.1204577

Nugroho, B., Pramudya, Y., & Widodo, W. (2019). The Content Analysis of Bisphenol A (BPA) on Water in Plastic Glass with Varying Temperatures and Contact Times using UV-VIS Spectrophotometer. *Indonesian Review of Physics*, 1(2), 27. https://doi.org/10.12928/irip.v1i2.263

Nunez, J., Celi, F. S., Ng, L., & Forrest, D. (2008). Multigenic control of thyroid hormone functions in the nervous system. In *Molecular and Cellular Endocrinology* (Vol. 287, Issues 1–2, pp. 1–12). https://doi.org/10.1016/j.mce.2008.03.006

Oh, J., Choi, J. W., Ahn, Y. A., & Kim, S. (2018). Pharmacokinetics of bisphenol S in humans after single oral administration. *Environment International*, *112*, 127–133. https://doi.org/10.1016/j.envint.2017.11.020

Olsvik, P. A., Whatmore, P., Penglase, S. J., Skjærven, K. H., D'Auriac, M. A., & Ellingsen, S. (2019). Associations between behavioral effects of bisphenol A and DNA methylation in zebrafish embryos. *Frontiers in Genetics*, *10*(MAR). https://doi.org/10.3389/fgene.2019.00184

Page, M. J., McKenzie, J. E., Bossuyt, P. M., Boutron, I., Hoffmann, T. C., Mulrow, C. D., Shamseer, L., Tetzlaff, J. M., Akl, E. A., Brennan, S. E., Chou, R., Glanville, J., Grimshaw, J. M., Hróbjartsson, A., Lalu, M. M., Li, T., Loder, E. W., Mayo-Wilson, E., McDonald, S., ... Moher, D. (2021). The PRISMA 2020 statement: An updated guideline for reporting systematic reviews. In *The BMJ* (Vol. 372). BMJ Publishing Group. https://doi.org/10.1136/bmj.n71

Palanza, P., Howdeshell, K. L., Parmigiani, S., & vom Saal, F. S. (2002). Exposure to a low dose of bisphenol A during fetal life or in adulthood alters maternal behavior in mice. *Environmental Health Perspectives*, *110*(SUPPL. 3), 415–422. https://doi.org/10.1289/ehp.02110s3415

Pan, Y., Deng, M., Li, J., Du, B., Lan, S., Liang, X., & Zeng, L. (2020). Occurrence and Maternal Transfer of Multiple Bisphenols, including an Emerging Derivative with Unexpectedly High Concentrations, in the Human Maternal-Fetal-Placental Unit. *Environmental Science and Technology*, *54*(6), 3476–3486. https://doi.org/10.1021/acs.est.0c00206

Pang, Q., Li, Y., Meng, L., Li, G., Luo, Z., & Fan, R. (2019). Neurotoxicity of BPA, BPS, and BPB for the hippocampal cell line (HT-22): An implication for the replacement of BPA in plastics. *Chemosphere*, *226*, 545–552. https://doi.org/10.1016/j.chemosphere.2019.03.177

Patel, J., Landers, K., Li, H., Mortimer, R. H., & Richard, K. (2011). Thyroid hormones and fetal neurological development. In *Journal of Endocrinology* (Vol. 209, Issue 1, pp. 1–8). BioScientifica. https://doi.org/10.1530/JOE-10-0444

Perera, F., Nolte, E. L. R., Wang, Y., Margolis, A. E., Calafat, A. M., Wang, S., Garcia, W., Hoepner, L. A., Peterson, B. S., Rauh, V., & Herbstman, J. (2016). Bisphenol A exposure and symptoms of anxiety and depression among inner city children at 10–12 years of age. *Environmental Research*, *151*, 195–202. https://doi.org/10.1016/j.envres.2016.07.028

Pfaffl, M. W. (2001). A new mathematical model for relative quantification in real-time RT-PCR. *Nucleic Acids Research*, *29*(9), e45. https://doi.org/10.1093/nar/29.9.e45

Prager, E. M., Bergstrom, H. C., Wynn, G. H., & Braga, M. F. M. (2016). The basolateral amygdala γ-aminobutyric acidergic system in health and disease. In *Journal of Neuroscience Research* (Vol. 94, Issue 6, pp. 548–567). John Wiley and Sons Inc. https://doi.org/10.1002/jnr.23690

Prezioso, G., Giannini, C., & Chiarelli, F. (2018). Effect of thyroid hormones on neurons and neurodevelopment. *Hormone Research in Paediatrics, 90*(2), 73–81. https://doi.org/10.1159/000492129

Qiu, W., Shao, H., Lei, P., Zheng, C., Qiu, C., Yang, M., & Zheng, Y. (2018). Immunotoxicity of bisphenol S and F are similar to that of bisphenol A during zebrafish early development. *Chemosphere*, *194*, 1–8. https://doi.org/10.1016/j.chemosphere.2017.11.125

Qiu, W., Yang, M., Liu, S., Lei, P., Hu, L., Chen, B., Wu, M., & Wang, K. J. (2018). Toxic Effects of Bisphenol S Showing Immunomodulation in Fish Macrophages. *Environmental Science and Technology*, *52*(2), 831–838. https://doi.org/10.1021/acs.est.7b04226

Qiu, W., Zhan, H., Hu, J., Zhang, T., Xu, H., Wong, M., Xu, B., & Zheng, C. (2019). The occurrence, potential toxicity, and toxicity mechanism of bisphenol S, a substitute of bisphenol A: A critical review of recent progress. *Ecotoxicology and Environmental Safety*, *173*, 192–202. https://doi.org/10.1016/j.ecoenv.2019.01.114

Qiu, W., Zhao, Y., Yang, M., Farajzadeh, M., Pan, C., & Wayne, N. L. (2016). Actions of bisphenol A and bisphenol S on the reproductive neuroendocrine system during early development in zebrafish. *Endocrinology*, *157*(2), 636–647. https://doi.org/10.1210/en.2015-1785

Rebuli, M. E., Cao, J., Sluzas, E., Barry Delclos, K., Camacho, L., Lewis, S. M., Vanlandingham, M. M., & Patisaul, H. B. (2014). Investigation of the effects of subchronic low dose oral exposure to bisphenol a (BPA) and ethinyl estradiol (EE) on estrogen receptor expression in the juvenile and adult female rat hypothalamus. *Toxicological Sciences*, *140*(1), 190–203. https://doi.org/10.1093/toxsci/kfu074

Rebuli, M. E., Gibson, P., Rhodes, C. L., Cushing, B. S., & Patisaul, H. B. (2016). Sex differences in microglial colonization and vulnerabilities to endocrine disruption in the social brain. *General and Comparative Endocrinology*, *238*, 39–46. https://doi.org/10.1016/j.ygcen.2016.04.018

Regueiro, J., & Wenzl, T. (2015). Development and validation of a stable-isotope dilution liquid chromatography-tandem mass spectrometry method for the determination of bisphenols in readymade meals. *Journal of Chromatography A, 1414*(1), 110–121. https://doi.org/10.1016/j.chroma.2015.08.037

Research and Markets. (2016). *Bisphenol-A - A Global Market Overview - Research and Markets*. https://www.researchandmarkets.com/reports/3625843/bisphenol-a-a-global-market-overview

Resnik, D. B., & Elliott, K. C. (2015). Bisphenol a and risk management ethics. *Bioethics*, *29*(3), 182–189. https://doi.org/10.1111/bioe.12079

Resnik, D. B., Konecny, B., & Kissling, G. E. (2017). Conflict of Interest and Funding Disclosure Policies of Environmental, Occupational, and Public Health Journals. *Journal of Occupational and Environmental Medicine*, *59*(1), 28–33. https://doi.org/10.1097/JOM.0000000000910

Rivollier, F., Krebs, M. O., & Kebir, O. (2019). Perinatal exposure to environmental endocrine disruptors in the emergence of neurodevelopmental psychiatric diseases: A systematic review. In *International Journal of Environmental Research and Public Health* (Vol. 16, Issue 8). MDPI AG. https://doi.org/10.3390/ijerph16081318

Rochester, J. R., & Bolden, A. L. (2015). Bisphenol S and F: A systematic review and comparison of the hormonal activity of bisphenol a substitutes. In *Environmental Health Perspectives* (Vol. 123, Issue 7, pp. 643–650). Public Health Services, US Dept of Health and Human Services. https://doi.org/10.1289/ehp.1408989

Rooney, A. A., Boyles, A. L., Wolfe, M. S., Bucher, J. R., & Thayer, K. A. (2014). Systematic review and evidence integration for literature-based environmental health science assessments. *Environmental Health Perspectives*, *122*(7), 711–718. https://doi.org/10.1289/ehp.1307972

Rycroft-Malone, J., McCormack, B., Hutchinson, A. M., DeCorby, K., Bucknall, T. K., Kent, B., Schultz, A., Snelgrove-Clarke, E., Stetler, C. B., Titler, M., Wallin, L., & Wilson, V. (2012). Realist synthesis: illustrating the method for implementation research. *Implementation Science*, 7(1), 33. https://doi.org/10.1186/1748-5908-7-33

Sadow, P. M., Chassande, O., Koo, E. K., Gauthier, K., Samarut, J., Xu, J., O'Malley, B. W., & Weiss, R. E. (2003). Regulation of expression of thyroid hormone receptor isoforms and coactivators in liver and heart by thyroid hormone. *Molecular and Cellular Endocrinology*, 203(1–2), 65–75. https://doi.org/10.1016/S0303-7207(03)00122-9

Saili, K. S., Corvi, M. M., Weber, D. N., Patel, A. U., Das, S. R., Przybyla, J., Anderson, K. A., & Tanguay, R. L. (2012). Neurodevelopmental low-dose bisphenol A exposure leads to early life-stage hyperactivity and learning deficits in adult zebrafish. *Toxicology*, *291*(1–3), 83–92. https://doi.org/10.1016/j.tox.2011.11.001

Samuels, H. H., Stanley, F., & Shapiro, L. E. (1977). Modulation of thyroid hormone nuclear receptor level by 3,5,3'-triiodo-L-thyronine in GH1 cells. Evidence for two functional components of nuclear-bound receptor and relationship to the induction of growth hormone synthesis. *Journal of Biological Chemistry*, *252*(17), 6052–6060. https://doi.org/10.1016/s0021-9258(17)40028-7

Sánchez-Piñero, J., Bowerbank, S. L., Moreda-Piñeiro, J., López-Mahía, P., & Dean, J. R. (2020). The occurrence and distribution of polycyclic aromatic hydrocarbons, bisphenol A and organophosphate flame retardants in indoor dust and soils from public open spaces: Implications for human exposure. *Environmental Pollution*, *266*. https://doi.org/10.1016/j.envpol.2020.115372

Santos, G. M., Fairall, L., & Schwabe, J. W. R. (2011). Negative regulation by nuclear receptors: A plethora of mechanisms. In *Trends in Endocrinology and Metabolism* (Vol. 22, Issue 3, pp. 87–93). Trends Endocrinol Metab. https://doi.org/10.1016/j.tem.2010.11.004

Semple, B. D., Blomgren, K., Gimlin, K., Ferriero, D. M., & Noble-Haeusslein, L. J. (2013). Brain development in rodents and humans: Identifying benchmarks of maturation and vulnerability to injury across species. In *Progress in Neurobiology* (Vols. 106–107, pp. 1–16). NIH Public Access. https://doi.org/10.1016/j.pneurobio.2013.04.001

Shahid, M. A., & Sharma, S. (2019). Physiology, Thyroid Hormone. In *StatPearls*. StatPearls Publishing. http://www.ncbi.nlm.nih.gov/pubmed/29763182

Shamseer, L., Moher, D., Clarke, M., Ghersi, D., Liberati, A., Petticrew, M., Shekelle, P., Stewart, L. A., Altman, D. G., Booth, A., Chan, A. W., Chang, S., Clifford, T., Dickersin, K., Egger, M., Gøtzsche, P. C., Grimshaw, J. M., Groves, T., Helfand, M., ... Whitlock, E. (2015). Preferred reporting items for systematic review and meta-analysis protocols (prisma-p) 2015: Elaboration and explanation. In *BMJ (Online)* (Vol. 349). BMJ Publishing Group. https://doi.org/10.1136/bmj.g7647

Sheikh, I. A. (2020). Molecular interactions of thyroxine binding globulin and thyroid hormone receptor with estrogenic compounds 4-nonylphenol, 4-tert-octylphenol and bisphenol A metabolite (MBP). *Life Sciences*, *253*. https://doi.org/10.1016/j.lfs.2020.117738

Sheng, Z. G., Tang, Y., Liu, Y. X., Yuan, Y., Zhao, B. Q., Chao, X. J., & Zhu, B. Z. (2012). Low concentrations of bisphenol a suppress thyroid hormone receptor transcription through a nongenomic mechanism. *Toxicology and Applied Pharmacology*, 259(1), 133–142. https://doi.org/10.1016/j.taap.2011.12.018

Skorupskaite, K., George, J. T., & Anderson, R. A. (2014). The kisspeptin-GnRH pathway in human reproductive health and disease. In *Human Reproduction Update* (Vol. 20, Issue 4, pp. 485–500). Oxford University Press. https://doi.org/10.1093/humupd/dmu009

Skranes, J. (2010). The Newborn Brain - neuroscience and clinical applications. *Acta Paediatrica*, *99*(8), 1277–1277. https://doi.org/10.1111/j.1651-2227.2010.01891.x

Somogyi, V., Horváth, T. L., Tóth, I., Bartha, T., Frenyó, L. V., Kiss, D. S., Jócsák, G., Kerti, A., Naftolin, F., & Zsarnovszky, A. (2016). Bisphenol a influences oestrogen- and thyroid hormone-regulated thyroid hormone receptor expression in rat cerebellar cell culture. *Acta Veterinaria Hungarica*, *64*(4), 497–513. https://doi.org/10.1556/004.2016.046

Sotres-Bayon, F., & Quirk, G. J. (2010). Prefrontal control of fear: More than just extinction. In *Current Opinion in Neurobiology* (Vol. 20, Issue 2, pp. 231–235). NIH Public Access. https://doi.org/10.1016/j.conb.2010.02.005

Stenzel, D., Wilsch-Bräuninger, M., Wong, F. K., Heuer, H., & Huttner, W. B. (2014). Integrin $\alpha\nu\beta3$ and thyroid hormones promote expansion of progenitors in embryonic neocortex. *Development (Cambridge)*, 141(4), 795–806. https://doi.org/10.1242/dev.101907

Stepien, B. K., & Huttner, W. B. (2019). Transport, metabolism, and function of thyroid hormones in the developing mammalian brain. *Frontiers in Endocrinology*, *10*(APR), 209. https://doi.org/10.3389/fendo.2019.00209

T.Chow, J. (2007). Environmental Assessment for Bisphenol A and Polycarbonate.

Teng, C., Goodwin, B., Shockley, K., Xia, M., Huang, R., Norris, J., Merrick, B. A., Jetten, A. M., Austin, C. P., & Tice, R. R. (2013). Bisphenol A affects androgen receptor function via multiple mechanisms. *Chemico-Biological Interactions*, 203(3), 556–564. https://doi.org/10.1016/j.cbi.2013.03.013

Thayer, K. A., Taylor, K. W., Garantziotis, S., Schurman, S. H., Kissling, G. E., Hunt, D., Herbert, B., Church, R., Jankowich, R., Churchwell, M. I., Scheri, R. C., Birnbaum, L. S., & Bucher, J. R. (2016). Bisphenol a, bisphenol s, and 4-hydro xyphenyl 4-isopro oxyphenyl sulfone (bpsip) in urine and blood of cashiers. *Environmental Health Perspectives*, 124(4), 437–444. https://doi.org/10.1289/ehp.1409427

Thoene, M., Dzika, E., Gonkowski, S., & Wojtkiewicz, J. (2020). Bisphenol S in food causes hormonal and obesogenic effects comparable to or worse than bisphenol a: A literature review. *Nutrients*, *12*(2). https://doi.org/10.3390/nu12020532

Tiwari, S. K., Agarwal, S., Chauhan, L. K. S., Mishra, V. N., & Chaturvedi, R. K. (2015). Bisphenol-A Impairs Myelination Potential During Development in the Hippocampus of the Rat Brain. *Molecular Neurobiology*, *51*(3), 1395–1416. https://doi.org/10.1007/s12035-014-8817-3

Tiwari, S. K., Agarwal, S., Seth, B., Yadav, A., Ray, R. S., Mishra, V. N., & Chaturvedi, R. K. (2015). Inhibitory Effects of Bisphenol-A on Neural Stem Cells Proliferation and Differentiation in the Rat Brain Are Dependent on Wnt/ β -Catenin Pathway. *Molecular Neurobiology*, *52*(3), 1735–1757. https://doi.org/10.1007/s12035-014-8940-1

Tiwari, S. K., Agarwal, S., Tripathi, A., & Chaturvedi, R. K. (2016). Bisphenol-A Mediated Inhibition of Hippocampal Neurogenesis Attenuated by Curcumin via Canonical Wnt Pathway. *Molecular Neurobiology*, *53*(5), 3010–3029. https://doi.org/10.1007/s12035-015-9197-z

Toda, C., Santoro, A., Kim, J. D., & Diano, S. (2017). POMC Neurons: From Birth to Death. In *Annual Review of Physiology* (Vol. 79, pp. 209–236). Annual Reviews Inc. https://doi.org/10.1146/annurev-physiol-022516-034110

Tovote, P., Fadok, J. P., & Lüthi, A. (2015). Neuronal circuits for fear and anxiety. In *Nature Reviews Neuroscience* (Vol. 16, Issue 6, pp. 317–331). Nature Publishing Group. https://doi.org/10.1038/nrn3945

Tse, W. K. F., Yeung, B. H. Y., Wan, H. T., & Wong, C. K. C. (2013). Early embryogenesis in zebrafish is affected by bisphenol A exposure. *Biology Open*, *2*(5), 466–471. https://doi.org/10.1242/bio.20134283

Ullah, H., Ullah, F., Rehman, O., Jahan, S., Afsar, T., Al-Disi, D., Almajwal, A., & Razak, S. (2021). Chronic exposure of bisphenol S (BPS) affect hypothalamic-pituitary-testicular activities in adult male rats: possible in estrogenic mode of action. *Environmental Health and Preventive Medicine*, *26*(1). https://doi.org/10.1186/s12199-021-00954-0

Valbonesi, P., Profita, M., Vasumini, I., & Fabbri, E. (2021). Contaminants of emerging concern in drinking water: Quality assessment by combining chemical and biological analysis. *Science of the Total Environment*, *758*, 143624. https://doi.org/10.1016/j.scitotenv.2020.143624

Vandenberg, L. N. (2014). Non-monotonic dose responses in studies of endocrine disrupting chemicals: Bisphenol a as a case study. *Dose-Response*, *12*(2), 259–276. https://doi.org/10.2203/dose-response.13-020.Vandenberg

Vandenberg, L. N., Ågerstrand, M., Beronius, A., Beausoleil, C., Bergman, Å., Bero, L. A., Bornehag, C. G., Boyer, C. S., Cooper, G. S., Cotgreave, I., Gee, D., Grandjean, P., Guyton, K. Z., Hass, U., Heindel, J. J., Jobling, S., Kidd, K. A., Kortenkamp, A., Macleod, M. R., ... Rudén, C. (2016). A proposed framework for the systematic review and integrated assessment (SYRINA) of endocrine disrupting chemicals. *Environmental Health: A Global Access Science Source*, *15*(1), 74. https://doi.org/10.1186/s12940-016-0156-6

Vandenberg, L. N., Colborn, T., Hayes, T. B., Heindel, J. J., Jacobs, D. R., Lee, D. H., Shioda, T., Soto, A. M., vom Saal, F. S., Welshons, W. V., Zoeller, R. T., & Myers, J. P. (2012). Hormones and endocrine-

disrupting chemicals: Low-dose effects and nonmonotonic dose responses. In *Endocrine Reviews* (Vol. 33, Issue 3, pp. 378–455). Oxford Academic. https://doi.org/10.1210/er.2011-1050

Viñas, P., Campillo, N., Martínez-Castillo, N., & Hernández-Córdoba, M. (2010). Comparison of two derivatization-based methods for solid-phase microextraction-gas chromatography-mass spectrometric determination of bisphenol A, bisphenol S and biphenol migrated from food cans. *Analytical and Bioanalytical Chemistry*, *397*(1), 115–125. https://doi.org/10.1007/s00216-010-3464-7

Visser, T. J. (2018). *Regulation of Thyroid Function, Synthesis, and Function of Thyroid Hormones* (pp. 3–32). https://doi.org/10.1007/978-3-319-45013-1_1

Visser, T. J., & Peeters, R. P. (2017). Metabolism of thyroid hormone. *New Comprehensive Biochemistry*, *18*, 81–103. https://doi.org/10.1016/S0167-7306(08)60641-9

Vogel, S. A. (2009). The politics of plastics: the making and unmaking of bisphenol a "safety". *American Journal of Public Health, 99 Suppl 3*. https://doi.org/10.2105/ajph.2008.159228

Wan, Y., Huo, W., Xu, S., Zheng, T., Zhang, B., Li, Y., Zhou, A., Zhang, Y., Hu, J., Zhu, Y., Chen, Z., Lu, S., Wu, C., Jiang, M., Jiang, Y., Liu, H., Yang, X., & Xia, W. (2018). Relationship between maternal exposure to bisphenol S and pregnancy duration. *Environmental Pollution*, *238*, 717–724. https://doi.org/10.1016/j.envpol.2018.03.057

Wan, Y., Xia, W., Yang, S., Pan, X., He, Z., & Kannan, K. (2018). Spatial distribution of bisphenol Sin surface water and human serum from Yangtze River watershed, China: Implications for exposurethroughdrinkingwater.Chemosphere,199,595–602.https://doi.org/10.1016/j.chemosphere.2018.02.040

Wang, W., Abualnaja, K. O., Asimakopoulos, A. G., Covaci, A., Gevao, B., Johnson-Restrepo, B., Kumosani, T. A., Malarvannan, G., Minh, T. B., Moon, H. B., Nakata, H., Sinha, R. K., & Kannan, K. (2015). A comparative assessment of human exposure to tetrabromobisphenol A and eight bisphenols including bisphenol A via indoor dust ingestion in twelve countries. *Environment International, 83*, 183–191. https://doi.org/10.1016/j.envint.2015.06.015

Wang, X., Dong, Q., Chen, Y., Jiang, H., Xiao, Q., Wang, Y., Li, W., Bai, C., Huang, C., & Yang, D. (2013). Bisphenol A affects axonal growth, musculature and motor behavior in developing zebrafish. *Aquatic Toxicology*, *142–143*, 104–113. https://doi.org/10.1016/j.aquatox.2013.07.011

Warita, K., Mitsuhashi, T., Ohta, K. ichi, Suzuki, S., Hoshi, N., Miki, T., & Takeuchi, Y. (2013). In vitro evaluation of gene expression changes for gonadotropin-releasing hormone 1, brain-derived neurotrophic factor and neurotrophic tyrosine kinase, receptor, type 2, in response to bisphenol A treatment. *Congenital Anomalies*, *53*(1), 42–45. https://doi.org/10.1111/j.1741-4520.2012.00381.x

Wei, P., Zhao, F., Zhang, X., Liu, W., Jiang, G., Wang, H., & Ru, S. (2018). Transgenerational thyroid endocrine disruption induced by bisphenol S affects the early development of zebrafish offspring. *Environmental Pollution*, *243*, 800–808. https://doi.org/10.1016/j.envpol.2018.09.042

Wen, L., He, C., Sifuentes, C. J., & Denver, R. J. (2019). Thyroid hormone receptor alpha is required for thyroid hormone-dependent neural cell proliferation during tadpole metamorphosis. *Frontiers in Endocrinology*, *10*(JUN). https://doi.org/10.3389/fendo.2019.00396

Wen, L., & Shi, Y. B. (2015). Unliganded thyroid hormone receptor α Controls developmental timing in Xenopus tropicalis. *Endocrinology*, *156*(2), 721–734. https://doi.org/10.1210/en.2014-1439

Wilkinson, M., & Imran, S. A. (2019). Hypothalamic Regulation of Thyroid Function. In *Clinical Neuroendocrinology* (pp. 97–117). Cambridge University Press. https://doi.org/10.1017/9781108149938.007

Wirth, E. K., Schweizer, U., & Köhrle, J. (2014). Transport of thyroid hormone in brain. In *Frontiers in Endocrinology* (Vol. 5, Issue JUN). Frontiers Research Foundation. https://doi.org/10.3389/fendo.2014.00098

Wolstenholme, J. T., Rissman, E. F., & Connelly, J. J. (2011). The role of Bisphenol A in shaping the brain, epigenome and behavior. In *Hormones and Behavior* (Vol. 59, Issue 3, pp. 296–305). https://doi.org/10.1016/j.yhbeh.2010.10.001

Wu, C. C., Shields, J. N., Akemann, C., Meyer, D. N., Connell, M., Baker, B. B., Pitts, D. K., & Baker, T. R. (2020). The phenotypic and transcriptomic effects of developmental exposure to nanomolar levels of estrone and bisphenol A in zebrafish. *Science of the Total Environment*, *757*, 143736. https://doi.org/10.1016/j.scitotenv.2020.143736

Wu, L. H., Zhang, X. M., Wang, F., Gao, C. J., Chen, D., Palumbo, J. R., Guo, Y., & Zeng, E. Y. (2018). Occurrence of bisphenol S in the environment and implications for human exposure: A short review. In *Science of the Total Environment* (Vol. 615, pp. 87–98). Elsevier B.V. https://doi.org/10.1016/j.scitotenv.2017.09.194

Wu, X., Majumder, A., Webb, R., & Stice, S. L. (2016). High content imaging quantification of multiple in vitro human neurogenesis events after neurotoxin exposure. *BMC Pharmacology and Toxicology*, *17*(1), 1–15. https://doi.org/10.1186/s40360-016-0107-4

Xiao, X., Zhang, X., Zhang, C., Li, J., Zhao, Y., Zhu, Y., Zhang, J., & Zhou, X. (2019). Toxicity and multigenerational effects of bisphenol S exposure to: Caenorhabditis elegans on developmental, biochemical, reproductive and oxidative stress. *Toxicology Research*, *8*(5), 630–640. https://doi.org/10.1039/c9tx00055k

Xiao, Y., Liu, R., Xing, L., Xu, Y., Shang, L., & Hao, W. (2011). Combined developmental toxicity of bisphenol A and genistein in micromass cultures of rat embryonic limb bud and midbrain cells. *Toxicology in Vitro*, *25*(1), 153–159. https://doi.org/10.1016/j.tiv.2010.10.010

Xing, L., Esau, C., & Trudeau, V. L. (2016). Direct regulation of aromatase B expression by 17βestradiol and dopamine D1 receptor agonist in adult radial glial cells. *Frontiers in Neuroscience*, *9*(JAN), 504. https://doi.org/10.3389/fnins.2015.00504 Xu, Xiao Bin, He, Y., Song, C., Ke, X., Fan, S. J., Peng, W. J., Tan, R., Kawata, M., Matsuda, K. I., Pan, B. X., & Kato, N. (2014). Bisphenol a regulates the estrogen receptor alpha signaling in developing hippocampus of male rats through estrogen receptor. *Hippocampus*, *24*(12), 1570–1580. https://doi.org/10.1002/hipo.22336

Xu, Xiaohong, Lu, Y., Zhang, G., Chen, L., Tian, D., Shen, X., Yang, Y., & Dong, F. (2014). BisphenolA promotes dendritic morphogenesis of hippocampal neurons through estrogen receptor-mediatedERK1/2signalpathway.Chemosphere,96,129–137.https://doi.org/10.1016/j.chemosphere.2013.09.063

Xue, J., Wan, Y., & Kannan, K. (2016). Occurrence of bisphenols, bisphenol A diglycidyl ethers (BADGEs), and novolac glycidyl ethers (NOGEs) in indoor air from Albany, New York, USA, and its implications for inhalation exposure. *Chemosphere*, 151, 1–8. https://doi.org/10.1016/j.chemosphere.2016.02.038

Yamazaki, E., Yamashita, N., Taniyasu, S., Lam, J., Lam, P. K. S., Moon, H. B., Jeong, Y., Kannan, P., Achyuthan, H., Munuswamy, N., & Kannan, K. (2015). Bisphenol A and other bisphenol analogues including BPS and BPF in surface water samples from Japan, China, Korea and India. *Ecotoxicology and Environmental Safety*, *122*, 565–572. https://doi.org/10.1016/j.ecoenv.2015.09.029

Yang, Yang, Fang, Z., Dai, Y., Wang, Y., Liang, Y., Zhong, X., Wang, Q., Hu, Y., Zhang, Z., Wu, D., & Xu, X. (2018). Bisphenol-A antagonizes the rapidly modulating effect of DHT on spinogenesis and long-term potentiation of hippocampal neurons. *Chemosphere*, *195*, 567–575. https://doi.org/10.1016/j.chemosphere.2017.12.086

Yang, Ying, & Wang, J. Z. (2017). From structure to behavior in basolateral amygdalahippocampus circuits. In *Frontiers in Neural Circuits* (Vol. 11, p. 86). Frontiers Media S.A. https://doi.org/10.3389/fncir.2017.00086

Yang, Yunjia, Lu, L., Zhang, J., Yang, Y., Wu, Y., & Shao, B. (2014). Simultaneous determination of seven bisphenols in environmental water and solid samples by liquid chromatography-electrospray tandem mass spectrometry. *Journal of Chromatography A*, *1328*, 26–34. https://doi.org/10.1016/j.chroma.2013.12.074

Yaoi, T., Itoh, K., Nakamura, K., Ogi, H., Fujiwara, Y., & Fushiki, S. (2008). Genome-wide analysis of epigenomic alterations in fetal mouse forebrain after exposure to low doses of bisphenol A. *Biochemical and Biophysical Research Communications*, *376*(3), 563–567. https://doi.org/10.1016/j.bbrc.2008.09.028

Yen, P. M., & Koibuchi, N. (2016). Thyroid Hormone Disruption and Neurodevelopment. In *Thyroid Hormone Disruption and Neurodevelopment*. https://doi.org/10.1007/978-1-4939-3737-0

Yin, N., Yao, X., Qin, Z., Wang, Y. L., & Faiola, F. (2015). Assessment of Bisphenol A (BPA) neurotoxicity in vitro with mouse embryonic stem cells. *Journal of Environmental Sciences (China)*, *36*, 181–187. https://doi.org/10.1016/j.jes.2015.06.004

Yoo, S. J., Joo, H., Kim, D., Lim, M. H., Kim, E., Ha, M., Kwon, H. J., Paik, K. C., & Kim, K. M. (2020). Associations between Exposure to Bisphenol A and Behavioral and Cognitive Function in Children with Attention-deficit/Hyperactivity Disorder: A Case-control Study. *Clinical Psychopharmacology and Neuroscience*, *18*(2), 261–269. https://doi.org/10.9758/cpn.2020.18.2.261

Yu, X., Xue, J., Yao, H., Wu, Q., Venkatesan, A. K., Halden, R. U., & Kannan, K. (2015). Occurrence and estrogenic potency of eight bisphenol analogs in sewage sludge from the U.S. EPA targeted national sewage sludge survey. *Journal of Hazardous Materials*, *299*, 733–739. https://doi.org/10.1016/j.jhazmat.2015.07.012

Zenata, O., Dvorak, Z., & Vrzal, R. (2017). Profiling of bisphenol S towards nuclear receptors activities in human reporter cell lines. *Toxicology Letters*, *281*, 10–19. https://doi.org/10.1016/j.toxlet.2017.09.006

Zhang, B., He, Y., Zhu, H., Huang, X., Bai, X., Kannan, K., & Zhang, T. (2020). Concentrations of bisphenol A and its alternatives in paired maternal–fetal urine, serum and amniotic fluid from an e-waste dismantling area in China. *Environment International*, *136*, 105407. https://doi.org/10.1016/j.envint.2019.105407

Zhang, D. H., Zhou, E. X., & Yang, Z. L. (2017). Waterborne exposure to BPS causes thyroid endocrine disruption in zebrafish larvae. *PLoS ONE*, *12*(5), e0176927. https://doi.org/10.1371/journal.pone.0176927

Zhang, H., Zhang, Y., Li, J., & Yang, M. (2019). Occurrence and exposure assessment of bisphenol analogues in source water and drinking water in China. *Science of the Total Environment*, *655*, 607–613. https://doi.org/10.1016/j.scitotenv.2018.11.053

Zhang, J., Begum, A., Brännström, K., Grundström, C., Iakovleva, I., Olofsson, A., Sauer-Eriksson, A. E., & Andersson, P. L. (2016). Structure-Based Virtual Screening Protocol for in Silico Identification of Potential Thyroid Disrupting Chemicals Targeting Transthyretin. *Environmental Science and Technology*, *50*(21), 11984–11993. https://doi.org/10.1021/acs.est.6b02771

Zhang, Zhi, Boelen, A., Kalsbeek, A., & Fliers, E. (2018). TRH Neurons and Thyroid Hormone Coordinate the Hypothalamic Response to Cold. In *European Thyroid Journal* (Vol. 7, Issue 6, pp. 279– 288). S. Karger AG. https://doi.org/10.1159/000493976

Zhang, Zifeng, Alomirah, H., Cho, H. S., Li, Y. F., Liao, C., Minh, T. B., Mohd, M. A., Nakata, H., Ren, N., & Kannan, K. (2011). Urinary bisphenol a concentrations and their implications for human exposure in several Asian countries. *Environmental Science and Technology*, *45*(16), 7044–7050. https://doi.org/10.1021/es200976k

Zhou, R., Chen, F., Chang, F., Bai, Y., & Chen, L. (2013). Persistent overexpression of DNA methyltransferase 1 attenuating GABAergic inhibition in basolateral amygdala accounts for anxiety in rat offspring exposed perinatally to low-dose bisphenol A. *Journal of Psychiatric Research*, 47(10), 1535–1544. https://doi.org/10.1016/j.jpsychires.2013.05.013

Zhou, X., Kramer, J. P., Calafat, A. M., & Ye, X. (2014). Automated on-line column-switching high performance liquid chromatography isotope dilution tandem mass spectrometry method for the quantification of bisphenol A, bisphenol F, bisphenol S, and 11 other phenols in urine. *Journal of Chromatography B: Analytical Technologies in the Biomedical and Life Sciences*, 944, 152–156. https://doi.org/10.1016/j.jchromb.2013.11.009

Zoeller, R. T., Bansal, R., & Parris, C. (2005). Bisphenol-A, an environmental contaminant that acts as a thyroid hormone receptor antagonist in vitro, increases serum thyroxine, and alters RC3/neurogranin expression in the developing rat brain. *Endocrinology*, *146*(2), 607–612. https://doi.org/10.1210/en.2004-1018

APPENDICES

Appendix A. Conversion Criteria for Risk of Bias Assessment

Question	Definitely Low Risk of Bias (++)	Probably Low Risk of Bias (+)	Probably High Risk of Bias (-/NR)	Definitely High Risk of Bias ()
Was the administered dose or exposure level adequately randomized?	There is direct evidence that animals were allocated to any study group (including control) using random consistent method AND there is direct concurrent control group is used as indication that randomization covered to all study groups	There is indirect evidence that animals were allocated to any study group (including control) using random consistent method AND there is direct/indirect evidence that concurrent control is used as indication that randomization covered all study groups OR It is deemed that allocation without clear random component would not appreciably bias the result	There is indirect evidence that animals were allocated to any study group (including controls) using the non-random method OR There is indirect evidence that there was a lack of concurrent control group OR There is insufficient information provided about how subjects were allocated to study groups (NR)	There is direct evidence that animals were allocated to any study group (including controls) using the non-random method OR There is direct evidence that there was a lack of concurrent control group
Was allocation to study groups adequately concealed?	There is direct evidence at the time of assigning study groups that the research personnel did not know what group of animals were allocated to AND it is unlikely that they could broke the blinding of allocation until the assignment was completed	There is indirect evidence at the time of assigning study groups that the research personnel did not know what group of animals were allocated to AND it is unlikely that they could broke the blinding of allocation until the assignment was completed OR It is deemed that allocation that lack of adequate allocation concealment would not appreciably bias the result	There is indirect evidence at the time of assigning study groups which the research personnel did not know what group of animals were allocated to AND It is likely that they could break the blinding of allocation until the assignment was completed OR There is insufficient information provided about allocation to study groups (NR)	There is direct evidence at the time of assigning study groups which the research personnel did not know what group of animals were allocated to AND It is likely that they could have broken the blinding of allocation until the assignment was completed
Were experimental conditions identical across study groups?	There is direct evidence that same vehicle was used in control and experimental animals	There is indirect evidence that same vehicle was used in control and experimental animals	There is indirect evidence that difference vehicle was used in control and experimental animals	There is direct evidence that difference vehicle was used in control and experimental animals

	AND evidence that non-treatment-related experimental conditions were identical across study groups (Study reports in detail explicitly)	AND Identical non- treatment related experimental conditions are assumed if authors did not report differences in housing or husbandry OR It is deemed that vehicle used would not appreciably bias the result AND Identical non-treatment related experimental conditions are assumed	OR Author did not report the vehicle used (NR) OR There is indirect evidence that non- treatment-related experimental conditions were not comparable between study group	or the control is untreated OR There is direct evidence that non- treatment-related experimental conditions were not comparable between study groups
		if authors did not report differences in housing or husbandry		
Were the research personnel and human subjects blinded to the study group during the study?	There is direct evidence that the research personnel were adequately blinded to study group and it is unlikely that they broke the blinding during the study	There is direct evidence that the research personnel were adequately blinded to the study groups AND it is unlikely they broke the blinding during the study OR It is deemed that lack of adequate blinding during the study would not appreciably bias the results	There is indirect evidence that the research personnel were not adequately blinded to study group OR There is insufficient information provided about blinding to study group (NR)	There is direct evidence that the research personnel were not adequately blinded to study group
Were outcome data complete without attrition or exclusion from analysis?	There is direct evidence that loss of animals was adequately addressed AND reasons were documented when animal was removed from the study OR Missing data have been imputed using appropriate methods (Ensure that characteristics of animals are not significantly different	There is indirect evidence that loss of animals was adequately addressed and reasons were documented when animal were removed from the study OR It is deemed that the proportion lost would not appreciably bias the results	There is indirect evidence that loss of animal was not adequately addressed and loss of animals was unacceptably large OR There is insufficient information provided about loss of animals (NR)	There is direct evidence that loss of animals was not adequately addressed AND loss of animal was unacceptably large

from animals retained from the analysis)

Can we be confident in the exposure characterization? There is direct evidence that the exposure was independently characterized by the researcher and purity confirmed as ≥99% for single substance or non-mixture evaluations AND The exposure was consistently administered across treatment groups (The same method and time-frame)

There is indirect evidence that the exposure was independently characterized by the researcher and purity confirmed as ≥99% for single substance or non-mixture evaluations AND there is indirect evidence that exposure was consistently administered across treatment groups (The same method and time-frame) OR

There is a direct evidence that purity was independently confirmed as ≥98% AND it is deemed that impurity up to 2% would not appreciably bias the results **AND** there is indirect evidence that exposure was consistently administered across treatment groups (The same method and time-frame) There is indirect evidence that the exposure was assessed using poorly validated methods **OR**

There is insufficient information provided about the validity of the exposure assessment method but no evidence for concern (NR) There is direct evidence that the exposure was assessed using poorly validated methods

Can we be confident in the outcome assessment? There is direct evidence that the outcome was assessed using well-established or gold standards method **AND** the assessment conducted at the same length of time in all study groups **AND** there is direct evidence that the outcome assessors were adequately There is direct evidence that the outcome was assessed using well-established or gold standards method **AND** the assessment conducted at the same length of time in all study groups **AND** there is indirect evidence that the outcome assessors were adequately There is indirect evidence that the outcome assessment method is an insensitive instrument **OR**

The length of time after initial exposure differed by study group **OR**

There is indirect evidence that it was possible for outcome There is direct evidence that the outcome assessment method is an insensitive instrument **OR** The length of time after initial exposure

differed by study group OR There is direct evidence that it was possible for outcome

	blinded to the study group and it is unlikely that they broke the blinding prior to reporting outcomes	blinded to the study group and it is unlikely that they broke the blinding prior to reporting outcomes OR It is deemed that the outcome assessment method used would not appreciably bias the results AND there is indirect evidence that the outcome assessors were adequately blinded to the study group and it is unlikely that they broke the blinding prior to reporting outcomes OR It is deemed that lack of adequate blinding of outcome assessors would not appreciably bias the results	assessors to infer the study group prior to reporting outcomes without sufficient quality control measures OR There is insufficient information provided about blinding of outcome assessors (NR)	assessors to infer the study group prior to reporting outcomes without sufficient quality control measures
Were all measured outcomes reported?	There is direct evidence that all of the study measured outcomes outline in the protocol, methods, abstract, and/or introduction (data that relevant for evaluation) have been reported	There is indirect evidence that all of the study measured outcomes outline in the protocol, methods, abstract, and/or introduction (data that relevant for evaluation) have been reported OR Analyses that had not been planned in advance are clearly indicated as such AND it is deemed that the additional analyses were appropriate and selective reporting would not appreciably bias the results	There is indirect evidence that all of the study measured outcomes outline in the protocol, methods, abstract, and/or introduction have been reported OR There is indirect evidence that unplanned analyses were included that may appreciably bias results OR There is insufficient information provided about selective outcome reporting (NR)	There is direct evidence that all of the study measured outcomes outline in the protocol, methods, abstract, and/or introduction have been reported
Were there no other potential threats to internal validity (statistical	There is direct evidence that the statistical method selected is appropriate	There is indirect evidence that the statistical method selected is appropriate	There is insufficient information provided about selective outcome reporting	There is direct evidence that the statistical method selected is not

methods were appropriate and researchers adhered to study protocol)? (rationale on the selection of statistical method) **AND** there is indirect evidence that the researcher adhere to the study protocol **AND** there is direct evidence on limitation of the study or potential confounding and modifying variables (rationale on the selection of statistical method) **AND** there is indirect evidence that the researcher adhere to study protocol **AND** there is indirect evidence on limitation of the study or potential confounding and modifying variables (NR)

appropriate (rationale on the selection of statistical method) **AND/OR** There is direct evidence that the researcher adhere to study protocol **AND/OR** there is direct evidence that the confounding and modifying variables cause bias in the study

Appendix B. Criteria for Determining Initial Confidence in the Body of Evidence Quality Assessment

Study Design Features	Description
Controlled Exposure	The exposure of the substance should be experimentally controlled
Exposure Conducted before Outcome	The exposure assessment showed that the exposure occurred before the development of the outcome or concurrent with aggravation/amplification of an existing condition
Individual Outcome Data	The outcomes should be assessed on individual level
Comparison Group Used	Appropriate comparison group should be used in the study

Appendix C. Criteria for Determining Final Confidence in the Body of Evidence Quality Assessment

Domains	Description
Risk of Bias Across Studies	Confidence was downgraded if most of the information was derived from tier 3 risk of bias

	study.
Unexplained Consistency	Confidence was downgraded if there is an unexplained consistency that not linked to the variation in the characteristic of the animal model used, exposure or treatment settings, and timing of outcome measurement
Indirectness	Confidence was downgraded if the study using non-vertebrate mammalian model or genetically modified rodents, bird, reptile, amphibian, and fish
Imprecision	Confidence was downgraded if the study has a large standard deviations and improper statistical analysis was used
Publication Bias	Confidence was downgraded if the studies were published in abstracts or type of grey literature or the conflict of interests present, or preliminary study
Large Magnitude Effect	Confidence was increased if statistically significant association or causation relationship observed
Dose Response	Confidence was increased if non-monotonic dose-response observed within studies
All Plausible Confounding	Confidence was increased if the study address the possible confounding that might affect the interpretation
Consistency Across Animal Study	Confidence was increased if there is a consistency across different strain of animals or species

Appendix D. Risk of Bias Across Studies Tiering Criteria

Tier 1 : A Study must be rated as "definitely low" or "probably low" risk of bias for key criteria AND have most other risk of bias criteria answered "definitely low" or "probably low" risk of bias Tier 2 : A Study meets neither criteria for tier 1 or tier 3 Tier 3 : A study must be rated as "definitely high" or "probably high" risk of bias for key criteria AND have most other risk of bias criteria answered "definitely high" or "probably high" risk of bias criteria	Risk of Bias Domains and Ratings								
	Key Criteria			Other Risk of Bias Criteria					
	Can we be confident in the exposure characterization?	Can we be confident in the outcome assessment ?	Were there no other potential threats to internal validity (statistical methods were appropriate and researchers adhered to study protocol)?	Was the administered dose or exposure level adequately randomized?	Was allocation to study groups adequately concealed?	Were experimenta l conditions identical across study groups?	Were the research personnel and human subjects blinded to the study group during the study?	Were outcome data complete without attrition or exclusion from analysis?	Were all measured outcomes reported?

Appendix F. Prisma-P 2020 Checklist

Section and Topic	ltem #	Checklist item	Location where item is reported
TITLE			
Title	1	Identify the report as a systematic review.	Cover Page
ABSTRACT			
Abstract	2	See the PRISMA 2020 for Abstracts checklist.	Page IV
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of existing knowledge.	Page 1-5
Objectives	4	Provide an explicit statement of the objective(s) or question(s) the review addresses.	Page 5
METHODS			
Eligibility criteria	5	Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses.	Page 41-42
Information sources	6	Specify all databases, registers, websites, organisations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted.	Page 42-43

Search strategy	7	Present the full search strategies for all databases, registers and websites, including any filters and limits used.	Page 42-43
Selection process	8	Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process.	Page 44
Data collection process	9	Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the process.	Page 44
Data items	10a	List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g. for all measures, time points, analyses), and if not, the methods used to decide which results to collect.	Page 45-47
	10b	List and define all other variables for which data were sought (e.g. participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information.	-
Study risk of bias assessment	11	Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process.	Page 47
Effect measures	12	Specify for each outcome the effect measure(s) (e.g. risk ratio, mean difference) used in the synthesis or presentation of results.	-
Synthesis methods	13a	Describe the processes used to decide which studies were eligible for each synthesis (e.g. tabulating the study intervention characteristics and comparing against the planned groups for each synthesis (item #5)).	Page 45-47
	13b	Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions.	-

13c Describe any methods used to tabulate or visually display results of individual studies and syntheses.

	13d	Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta-analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used.	-
	13e	Describe any methods used to explore possible causes of heterogeneity among study results (e.g. subgroup analysis, meta- regression).	-
	13f	Describe any sensitivity analyses conducted to assess robustness of the synthesized results.	-
Reporting bias assessment	14	Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases).	Page 47
Certainty assessment	15	Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome.	Page 48
RESULTS			
Study selection	16a	Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram.	Page 57
	16b	Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded.	-
Study characteristics	17	Cite each included study and present its characteristics.	Page 58

-

Risk of bias in studies	18	Present assessments of risk of bias for each included study.	Page 59
Results of individual studies	19	For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g. confidence/credible interval), ideally using structured tables or plots.	Page 69, Page 77, Page 81
Results of syntheses	20a	For each synthesis, briefly summarise the characteristics and risk of bias among contributing studies.	Page 57
	20b	Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g. confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect.	-
	20c	Present results of all investigations of possible causes of heterogeneity among study results.	-
	20d	Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results.	-
Reporting biases	21	Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed.	Page 58
Certainty of evidence	22	Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed.	Page 58
DISCUSSION			
Discussion	23a	Provide a general interpretation of the results in the context of other evidence.	Page 86- Page 97

OTHER INFORMATION			
23d	Discuss implications of the results for practice, policy, and future research.	Page 109	
23c	Discuss any limitations of the review processes used.	Page 105	
23b	Discuss any limitations of the evidence included in the review.	Page 98	

Registration and protocol	24a	Provide registration information for the review, including register name and registration number, or state that the review was not registered.	-
	24b	Indicate where the review protocol can be accessed, or state that a protocol was not prepared.	-
	24c	Describe and explain any amendments to information provided at registration or in the protocol.	-
Support	25	Describe sources of financial or non-financial support for the review, and the role of the funders or sponsors in the review.	-
Competing interests	26	Declare any competing interests of review authors.	-
Availability of data, code and other materials	27	Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review.	-

Appendix F. GENEzol[™] Reagent (Geneaid) RNA extraction Protocol

RNA Extraction Protocol Procedure

Please read the entire instruction manual prior to starting the Protocol Procedure.

1. Sample Homogenization

Sample preparation should be performed at room temperature. Please follow the table below for specific sample preparation. To avoid DNA contamination of extracted RNA, be sure and use the indicated volume of GENEzol™ Reagent.

Sample	Procedure			
	1. Remove the culture medium from culture dish.			
	 Directly add 100 µl of GENEzol[™] Reagent per cm² of culture dish surface area. 			
Adherent Cultured Cells	Lyse the cells directly in the culture dish by pipetting several times.			
	Incubate the sample mixture for 5 minutes at room temperature.			
	Transfer the sample to a new 1.5 ml of microcentrifuge tube (RNase-free).			
	 Transfer cells (up to 5 x 10⁸) to a 1.5 ml microcentrifuge tube (RNase-free). 			
	Harvest cells by centrifugation at 300 x g for 5 minutes to form a cell pellet.			
Suspension Cultured Cells	Remove the culture medium completely.			
	 Add 1 ml of GENEzol[™] Reagent to the cell pellet and lyse the cells by pipetting several times. 			
	Incubate the sample mixture for 5 minutes at room temperature.			
	 Add 1 ml of GENEzol™ Reagent to 50–100 mg of tissue sample. 			
Tiesue	Homogenize tissue samples using a glass-Teflon or Polytron homogenizer.			
TISSUE	Incubate the homogenized sample for 5 minutes at room temperature.			
	Transfer the sample to a new 1.5 ml of microcentrifuge tube (RNase-free).			
	 Transfer up to 300 µl of liquid sample to a 1.5 ml of microcentrifuge tube (RNase-free). 			
Body Fluids (blood, buffy	 Add 3 volumes of GENEzol[™] Reagent to each volume of liquid sample (3:1) 			
coat, plasma, serum)	Mix well by vortex.			
	Incubate the sample mixture for 5 minutes at room temperature.			
NOTE: For samples which contain high levels of fat, proteins, polysaccharides, or extracellular material, perform this optional step following				
sample homogenization. However, if DNA extraction is required, DO NOT perform this additional step.				
 Centrifuge the sample at 12-16,000 x g for 10 minutes to remove insoluble particles. 				
NOTE: Following centrifugation of high fat content samples, a layer a fat will float on the supernatant. Remove and discard the fatty layer.				
Transfer the clear supernatant to a new 1.5 ml microcentrifuge tube (RNase-free).				

3. Proceed to Step 2 Phase Separation.

2. Phase Separation

1. Add 200 µl of chloroform to the sample per 1 ml of GENEzol™ Reagent used in sample homogenization.

2. Shake the microcentrifuge tube vigorously for 10 seconds.

3. Centrifuge the sample at 12–16,000 x g for 15 minutes at 4°C to separate the phases.

NOTE: RNA is in the colorless upper aqueous phase which is approximately 50% of the total volume.

4. Transfer the upper aqueous phase to a new 1.5 ml microcentrifuge tube (RNase-free).

NOTE: Be careful not to draw any of the interphase layer (white) or organic phase layer (red) when transferring the aqueous layer. If DNA isolation is required, save the interphase and organic phase then proceed with the DNA Extraction protocol on page 3.

3. RNA Precipitation

1. Add 1 volume of isopropanol to the aqueous phase then mix by inverting the tube several times.

- 2. Incubate the sample mixture for 10 minutes at room temperature.
- 3. Centrifuge the sample at 12-16,000 x g for 10 minutes at 4°C to form a tight RNA pellet.
- 4. Carefully remove and discard the supernatant.

4. RNA Wash

- 1. Add 1 ml of 70% ethanol to wash the RNA pellet then vortex briefly.
- 2. Centrifuge the sample at 12-16,000 x g for 5 minutes at 4°C.
- 3. Being careful not to contact the RNA pellet, remove the supernatant with a pipette.
- 4. Air-dry the RNA pellet for 5-10 minutes at room temperature.
- NOTE: DO NOT dry the RNA pellet by vacuum centrifuge and avoid over drying the RNA pellet.

5. RNA Resuspension

- 1. Add 20-50 µl of RNase-free Water to resuspend the RNA pellet.
- 2. Incubate at 55-60°C for 10-15 minutes to dissolve the RNA pellet.

NOTE: Occasionaly tapping the bottom of the tube during incubation will promote RNA rehydration.

The RNA is ready for downstream applications or storage at -70°C.

GENEzol[™] Reagent Page 2 of 4

www.geneaid.com

Ver. 06.18.15

Appendix G. GENEzol[™] TriRNA Pure Kit (Geneaid) Protocol

RNA Purification Protocol Procedure

Please read the entire instruction manual prior to starting the Protocol Procedure.

Additional Requirements

absolute ethanol, lysozyme and bacteria lysis buffer (bacteria only), 1.5 ml microcentrifuge tubes (RNase-free)

Optional Requirements

1 µL of 20 mM EGTA (pH=8.0) for Optional Step 2: DNA Digestion in Solution

1. Sample Homogenization and Lysis

Sample preparation should be performed at room temperature. Please follow the table below for specific sample preparation. To avoid DNA contamination of extracted RNA, be sure and use the indicated volume of GENEzol™ Reagent. Lysozyme (LY420) and Bacteria Lysis Buffer (BLB00030) can be purchased directly from Geneaid.

Sample	Procedure
	 Remove the culture medium from the culture dish.
	 Directly add 100 µl of GENEzol[™] Reagent per cm² of culture dish surface area.
Adherent Cultured Cells	Lyse the cells directly in the culture dish by pipetting several times.
	Incubate the sample mixture for 5 minutes at room temperature.
	Transfer the sample to a 1.5 ml microcentrifuge tube (RNase-free).
	 Transfer cells (up to 5 x 10⁶) to a 1.5 ml microcentrifuge tube (RNase-free).
Suppopulation Cultured Colla	Harvest by centrifugation at 300 x g for 5 minutes then remove the culture medium completely.
Suspension Guitarea Celis	3. 700 µl of GENEzol™ Reagent should be added to the cell pellet then mixed several times by pipette.
	Incubate the sample mixture for 5 minutes at room temperature.
	1. Excise 10-50 mg of tissue directly from the animal or remove the tissue sample from storage. Do not use
	more than 50 mg of tissue per reaction.
	2. Homogenize tissue samples using one of the following methods: A. Transfer the tissue and 700 μl
	of GENEzol™ Reagent to a 2 ml centrifuge tube containing ceramic beads or stainless steel beads then
Tierua	homogenize the sample with a TissueLyser, Disruptor Genie or similar. B. Transfer the tissue and 700 µl
113505	of GENEzol™ Reagent to a 1.5 ml centrifuge tube and grind the tissue with a micropestle a few times
	then shear the tissue by passing the lysate through a 20-G needle syringe 10 times. C. Transfer the tissue
	and 700 µl of GENEzol™ Reagent to a glass-Teflon or Polytron homogenizer. Transfer the homogenized
	sample to a 1.5 ml microcentrifuge tube (RNase-free).
	 Incubate the homogenized sample for 5 minutes at room temperature.
Body Fluids (blood, buffy	 Transfer up to 200 µl of liquid sample to a 1.5 ml of microcentrifuge tube (RNase-free).
coat plasma serum)	2. Add 3 volumes of GENEzol™ Reagent per 1 volume of sample (3:1) then mix well by vortex.
coat, plasma, scramy	3. Incubate the sample mixture for 5 minutes at room temperature.
	 Transfer bacteria cells (up to 1 x 10⁸) to a 1.5 ml microcentrifuge tube (RNase-free).
	Centrifuge at 12-16,000 x g for 2 minutes then remove the supernatant completely.
	3. Weigh and transfer 10 mg of lysozyme powder to a new 1.5 ml microcentrifuge tube (RNase-free).
	Add 1 ml of bacteria lysis buffer to the microcentrifuge tube containing 10 mg of lysozyme.
Bacteria	Vortex the tube until the lysozyme powder is completely dissolved.
Dactoria	Add 100 µl of bacteria lysis buffer containing lysozyme to the bacteria cell pellet.
	Resuspend the cell pellet by vortex or pipetting.
	NOTE: Residual bacteria lysis buffer containing lysozyme should be stored at 4°C for 2 weeks.
	 Incubate the sample for 5 minutes at room temperature.
	 Add 700 µl of GENEzol[™] Reagent, mix well by pipette then incubate at room temperature for 5 minutes.
	 Cut off 20-50 mg of fresh or frozen plant tissue. Do not use more than 50 mg of plant tissue per rxn.
	2. Homogenize plant tissue samples using one of the following methods: A. Transfer the plant tissue
	and 700 µI GENEzol™ Reagent to a 2 ml centrifuge tube containing ceramic beads or stainless steel
Plant	beads then homogenize the sample with a TissueLyser, Disruptor Genie or similar. B. Add liquid nitrogen
	to a mortar (RNase-free) and grind the plant tissue thoroughly using a pestle (RNase-free). Transfer the
	plant tissue powder and 700 µl of GENEzol™ Reagent to a 1.5 ml centrifuge tube then vortex briefly.
	Incubate the homogenized sample for 5 minutes at room temperature.

2. RNA Binding

1. Centrifuge the sample at 12-16,000 x g for 1 minute to remove cell debris then transfer the clear supernatant to a new 1.5 ml microcentrifuge tube (RNase-free).

NOTE: When extracting RNA from cultured cell samples, cell debris will not commonly collect on the bottom of the microcentrifuge tube. In this case, proceed without transferring the supernatant.

2. Add 1 volume of absolute ethanol directly to 1 volume of sample mixture (1:1) in GENEzol™ Reagent.

3. Mix well by vortex then place a RB Column in a 2 ml Collection Tube.

4. Transfer 700 µl of the sample mixture to the RB Column. Centrifuge at 14-16,000 x g for 1 minute then discard the flow-through.

5. Repeat the RNA Binding Step by transferring the remaining sample mixture to the RB Column.

6. Centrifuge at 14-16,000 x g for 1 minute then discard the flow-through. Place the RB Column in a new 2 ml Collection Tube.

www.geneaid.com

Ver. 03.28.17

Optional Step 1: In Column DNase I Digestion IMPORTANT

DNA contamination is significantly reduced following In Column DNase I Digestion. However, traces of residual DNA may be detected in very sensitive applications. In this situation, please perform Optional Step 2: DNA Digestion In Solution instead to efficiently remove trace amounts of DNA. Standard DNase buffers are incompatible with In Column DNase I Digestion and may affect RNA integrity and reduce yield.

1. Add 400 µl of Wash Buffer (make sure ethanol was added) to the RB Column then centrifuge at 14-16,000 x g for 30 seconds.

2. Discard the flow-through and place the RB Column back in the 2 ml Collection Tube.

3. Prepare DNase I solution in a 1.5 ml microcentrifuge tube (RNase-free) as follows:

DNase I	5 μl (2 U/μl)
DNase I Reaction Buffer	45 µl
Total volume	50 ul

4. Gently pipette the DNase I solution to mix (DO NOT vortex) then add DNase I solution (50 µl) into the CENTER of the RB column matrix.

5. Incubate the column for 15 minutes at room temperature (20-30°C) then proceed with RNA Wash.

3. RNA Wash

1. Add 400 µl of Pre-Wash Buffer (make sure ethanol was added) to the RB Column then centrifuge at 14-16,000 x g for 30 seconds.

2. Discard the flow-through then place the RB Column back in the 2 ml Collection Tube.

3. Add 600 µl of Wash Buffer (make sure ethanol was added) to the RB Column.

- 4. Centrifuge at 14-16,000 x g for 30 seconds then discard the flow-through. Place the RB Column back in the 2 ml Collection Tube.
- 5. Add 600 µl of Wash Buffer (make sure ethanol was added) to the RB Column.

6. Centrifuge at 14-16,000 x g for 30 seconds then discard the flow-through.

7. Place the RB Column back in the 2 ml Collection Tube.

NOTE: For blood samples only, wash the RB Column again with 600 µl of Wash Buffer.

8. Centrifuge at 14-16,000 x g for 3 minutes to dry the column matrix.

4. RNA Elution

1. Place the dry RB Column in a clean 1.5 ml microcentrifuge tube (RNase-free).

2. Add 25-50 µl of RNase-free Water into the CENTER of the column matrix.

- 3. Let stand for at least 3 minutes to ensure the RNase-free Water is completely absorbed by the matrix.
- 4. Centrifuge at 14-16,000 x g for 1 minute to elute the purified RNA.

Optional Step 2: DNA Digestion In Solution

1. Prepare DNase I reaction in a 1.5 ml microcentrifuge tube (RNase-free) as follows:

RNA in RNase-free water	1-40 µl
DNase I	0.5 μl/μg RNA
DNase I Reaction Buffer	5 µl
RNase-free water	add to final volume = 50 µl
Total volume	50 µl

2. Gently pipette the DNase I reaction solution to mix (DO NOT vortex) then incubate the microcentrifuge tube at 37°C for 15-30 minutes.

3. Stop the reaction by adding 1 µl of 20 mM EGTA (pH=8.0) then incubate the microcentrifuge tube at 65°C for 10 minutes.

NOTE: DNase I Reaction Buffer may cause aberrant migration or smearing of RNA on gels. If analyzing RNA by gel electrophoresis, repurify the RNA sample by using the Geneaid ™ RNA Cleanup Kit instead of stopping the reaction with EGTA.

Appendix H. RevertAid First Strand cDNA Synthesis Kit (THermo Scientific[™]) Protocol

PROTOCOLS

I. First Strand cDNA Synthesis

After thawing, mix and briefly centrifuge the components of the kit. Store on ice.

 Add the following reagents into a sterile, nucleasefree tube on ice in the indicated order:

Template RNA	total RNA or poly(A) mRNA or specific RNA	0.1 ng - 5 µg 10 pg - 0.5 µg 0.01 pg - 0.5 µg
Primer	Oligo (dT)18 primer or Random Hexamer primer or gene-specific primer	1 μL 1 μL 15-20 pmol
Water, nucl	to 12 µL	
	12 µL	

- Optional. If the RNA template is GC-rich or contains secondary structures, mix gently, centrifuge briefly and incubate at 65°C for 5 min. Chill on ice, spin down and place the vial back on ice.
- 3. Add the following components in the indicated order:

5X Reaction Buffer	4 µL
RiboLock RNase Inhibitor (20 U/µL)	1 µL
10 mM dNTP Mix	2 µL
RevertAid M-MuLV RT (200 U/µL)	1 µL
Total volume	20 µL

- 4. Mix gently and centrifuge briefly.
- For oligo(dT)₁₈ or gene-specific primed cDNA synthesis, incubate for 60 min at 42°C. For random hexamer primed synthesis, incubate for 5 min at 25°C followed by 60 min at 42°C. Note. For GC-rich RNA templates the reaction temperature can be increased up to 45°C.
- Terminate the reaction by heating at 70°C for 5 min.

The reverse transcription reaction product can be directly used in PCR applications or stored at -20°C for less than one week. For longer storage, -70°C is recommended

Appendix I. QuantiNova SYBR Green RT-PCR Kit (Qiagen) Protocol

- Thaw QuantiNova SYBR Green RT-PCR Master Mix, QuantiNova Yellow Template Dilution Buffer, template RNA, QuantiNova Internal Control RNA (optional), primers, QN ROX Reference Dye (if required) and RNase-free water. Mix the individual solutions.
- Prepare a reaction mix according to Table 1. Due to the 2-phase hot start of both the RT and the PCR reactions, it is not necessary to keep samples on ice during reaction setup or while programming the real-time cycler.

Table 1. Reaction mix setup

Component	96-well block, Rotor-Gene	384-well block	Final concentration
2x SYBR Green RT-PCR Master Mix	10 µl	5 µl	lx
QN ROX Reference Dye (AB instruments only)	1 µl/0.1 µl*	0.5 µl/0.05 µl*	lx
QN SYBR Green RT-Mix	0.2 µl	0.1 µl	1x
20x primer mix (or Ctrl_QNIC_1_SG QuantiTect Primer Assay!)	1 µlt	0.5 µl†	0.5 µM forward primer 0.5 µM reverse primer
QN IC RNA (optional)	1 µl	1 µl	lx
RNase-free water	Variable	Variable	-
Template RNA (added at step 4)	Variable	Variable	≤200 ng/reaction
Total reaction volume	20 µl	10 µl	-

*Results in a 1:20 dilution for high ROX dye cyclers (i.e., ABI PRISM 7000, Applied Biosystems 7300, 7900 and StepOne Real-Time PCR Systems) and a 1:200 dilution for low-ROX dye cyclers (i.e., Applied Biosystems 7500 and ViiA7 Real-Time PCR Systems) in the final 1x reaction.

t If using the QN IC RNA to monitor RT-PCR amplification, add 2 µl (for 96-well) or 1 µl (for 384-well) of the 10x Ctrl_QNIC_1_SG QuantiTect Primer Assay.

- Mix the reaction thoroughly and dispense appropriate volumes into PCR tubes, PCR capillaries or wells of a PCR plate.
- Add template RNA (200 ng 100 fg per reaction, depending on target transcript abundance) to the individual PCR tubes, capillaries or wells containing the reaction mix.
- 5. Program the real-time cycler according to Table 2.

Note: Data acquisition should be performed during the combined annealing/extension step.

6. Place the PCR tubes or plates in the real-time cycler and start the cycling program.

Table 2. Cycling condit	Cycling conditions
-------------------------	--------------------

Step	Time	Temperature	Ramp rate
RT-step	10 min	50°C	Maximal/fast mode
PCR initial heat activation	2 min	95°C	Maximal/fast mode
2-step cycling			
Denaturation	5 s	95°C	Maximal/fast mode
Combined annealing/extension	10 s*	60°C	Maximal/fast mode
Number of cycles	40 [†]		

*If your cycler does not accept this short time for data acquisition, choose the shortest acceptable time.

* The number of cycles depends on the amount of template RNA.



Appendix J. Normalized RT-PCR Amplification Curve for Primer Efficiency Determination

Figure J.1. Normalized RT-PCR Amplification Curve for GAPDH. Each dilution of the cDNA was analyzed for GAPDH expression in duplicates. Each line represents a single reaction and the Ct-value obtained from the replicates were averaged and plotted against the log concentration of the reaction



Figure J.2. Normalized RT-PCR Amplification Curve for THR α . Each dilution of the cDNA was analyzed for THR α expression in duplicates. Each line represents a single reaction and the Ct-value obtained from the replicates were averaged and plotted against the log concentration of the reaction.

Appendix K. Ct-value Data Obtained from RT-PCR Analysis for Primer Efficiency Determination

Sample Name	Line Color	Ct-Value	Average Ct-value
GAPDH Non-diluted A		13.86	13.95
GAPDH Non-diluted B		14.04	
GAPDH 1:10 A		16.89	16.925
GAPDH 1:10 B		16.96	
GAPDH 1:100 A		20.38	20.365
GAPDH 1:100 B		20.39	
GAPDH 1:500 A		22.62	22.72
GAPDH 1:500 B		22.82	
GAPDH 1:1000 A		22.94	22.96
GAPDH 1:1000 B		22.98	
NTC A		24.4	24,35
NTC B		24.3	

Table K.1. Ct-values from RT-PCR Analysis of GAPDH for Primer Efficiency Determination

Table K.2. Ct-values from RT-PCR Analysis of $THR\alpha$ for Primer Efficiency Determination

Sample Name	Line Colour	Ct-Value	Average Ct-value
THR α Non-diluted A		23.13	23.16
THR α Non-diluted B		23.19	
THRα 1:10 A		25.92	25.92
THRα 1:10 B		25.92	
THRα 1:100 A		29.98	29.95
THRα 1:100 B		29.92	
THRα 1:500 A		30.76	31.24



Appendix L. Normalized RT-PCR Amplification Curve for Gene Expression Analysis



Figure L.1. Normalized RT-PCR Amplification Curve for THR α expression analysis in embryonic day 16 mice brain (1st technical replication). Each line represents a single reaction of either BPS or control samples. There are 4 samples analyzed from both treatment groups. The Ct-value obtained from this



technical replication were averaged with the Ct-value obtained from other technical replicates.

Figure L.2. Normalized RT-PCR Amplification Curve for THRα expression analysis in embryonic day 16 mice brain (2nd technical replication). Each line represents a single reaction of either BPS or control samples. There are 4 samples analyzed from both treatment groups. The Ct-value obtained from this technical replication were averaged with the Ct-value obtained from other technical replicates.



Figure L.3. Normalized RT-PCR Amplification Curve for THR α expression analysis in postnatal day 1 mice brain (1st technical replication). Each line represents a single reaction of either BPS or control



samples. There are 4 samples analyzed from both treatment groups. The Ct-value obtained from this technical replication were averaged with the Ct-value obtained from other technical replicates.

Figure L.3. Normalized RT-PCR Amplification Curve for THR α expression analysis in postnatal day 1 mice brain (2nd technical replication). Each line represents a single reaction of either BPS or control samples. There are 4 samples analyzed from both treatment groups. The Ct-value obtained from this technical replication were averaged with the Ct-value obtained from other technical replicates.

Appendix M. Ct-value Data Obtained from RT-PCR for Gene Expression Analysis

Time point	Gene	Sample Name	Line Color	Ct-v	alue	Average Ct-value
Embryonic day 16 Brain	GAPDH	Control 1		11.76	12.78	12.27
		Control 2		12.09	13.09	12.59
		Control 3		11.65	13.07	12.36
		Control 4		12.08	13.33	12.705
		BPS 1		12.51	13.97	13.24
		BPS 2		12.41	13.5	12.955
		BPS 3		11.62	13.44	12.53
		BPS 4		11.89	13.22	12.555
		NTC		-	26.8	-
	THRα	Control 1		17.41	18.73	18.07
		Control 2		17.57	19.05	18.31
		Control 3		17.09	18.27	17.68
		Control 4		17.83	18.94	18.385
		BPS 1		19.1	20.49	19.795
		BPS 2		19.38	20.91	20.145
		BPS 3		17.64	18.88	18.26
		BPS 4		17.62	18.74	18.18
		NTC		-	-	-
Postnatal day 1 brain	GAPDH	Control 1		16.79	14.85	15.82
		Control 2		14.51	15.64	15.075
		Control 3		16.34	15.04	15.69
		Control 4		14.1	14.79	14.445
		BPS 1		13.94	14.5	14.22
		BPS 2		18	15.41	16.705

Table K.2. Ct-values from RT-PCR Analysis of $THR\alpha$ for Primer Efficiency Determination

	BPS 3	15.52	15.11	15.315
	BPS 4	14.4	16.24	15.32
	NTC	-	30.19	15.82
THRα	Control 1	20.6	21.08	20.84
	Control 2	19.84	20	19.92
	Control 3	19.76	19.98	19.87
	Control 4	20.15	19.77	19.96
	BPS 1	20.2	20.07	20.135
	BPS 2	20.81	20.94	20.875
	BPS 3	20.13	20.47	20.3
	BPS 4	20.36	20.27	20.315
	NTC	-	32.31	-

Appendix N. Melting Curve of RT-PCR for Gene Expression Analysis



Figure N.1. Melting Curve Analysis of 1^{st} technical replication of THR α expression analysis in embryonic day 16 mice brain. Each line represents a single reaction of either BPS or control samples.

There is only a single peak observed in each sample indicating no contamination or primer dimer



Figure N.2. Melting Curve Analysis of 2^{nt} technical replication of THR α expression analysis in embryonic day 16 mice brain. Each line represents a single reaction of either BPS or control samples. There is only a single peak observed in each sample indicating no contamination or primer dimer



formation.

Figure N.3. Melting Curve Analysis of 1^{nt} technical replication of THR α expression analysis in postnatal day 1 mice brain. Each line represents a single reaction of either BPS or control samples. There is only a single peak observed in each sample indicating no contamination or primer dimer formation.



Figure N.4. Melting Curve Analysis of 2^{nd} technical replication of THR α expression analysis in postnatal day 1 mice brain. Each line represents a single reaction of either BPS or control samples. There is only a single peak observed in each sample indicating no contamination or primer dimer formation.

Appendix O. Details on Technical Issues due to Nanodrop Imprecision

Inconsistencies were found on the nanodrop measurement as demonstrated by the measurement for sample 3 & 4 from BPS treated group on embryonic day 16. As seen on figure O.1. below, the measurement of RNA purity and concentration for sample E16 BPS 3 & 4 showed a good result with A260/A280 ratio of 1.95 and concentration of 581.0 ng/µl for sample 3 and A260/A280 ratio of 1.95 and concentration of 581.0 ng/µl for sample 3 and A260/A280 ratio of 1.95 and concentration of 578.7 ng/µl for sample 4. This measurement was taken on 29th April 2021. The same samples then measured on 11th May 2021, following the claims that the nanodrop cannot be used and showed a negative result. Surprisingly, the measurement of the same samples showed a negative result with no A260/A280 ratio (See figure 0.2). Furthermore, the measurement of nucleus free water (blank) showed a positive result with A260/A280 ratio of 48.67 and concentration of 0.5 ng/µl. Another measurement showed no A260/A280 ratio value and concentration of -35.9 ng/ µl, indicating imprecision results generated by Nanodrop used.

RNA (Factor: 40) RNA (Factor: 40) A260 (10 mm): 14.467 A260 (10)A260/A280 60/A2

Figure O.1. Result of Nanodrop Reading of Sample 3 (left) & 4 (right) BPS E16. This measurement was taken on 29th April 2021 right after the extraction.

RNA (Factor: 40)	RNA (Factor: 40)
#1 A260 (10 mm): -0.015 A260/A280: 0.00	#2 A260 (10 mm): -1.133 A260/A280: 0.00
-0.6 ng/µl	-45.3 ng/µl

Figure O.2. Result of Nanodrop Reading of Sample 3 (left) & 4 (right) BPS E16. This measurement was taken on 11th May 2021.

RNA (Factor: 40)	#1 2
#14 A260 (10 mm): 0.012 A260/A280: 48.67	A260 (10 mm): -1.087 A260/A280: 0.00
0.5 ng/µ	rigiµ[

Figure O.3. Result of Nanodrop Reading of Nucleus free water (blank). Two measurement was made and the results were inconsistent. This measurement was taken on 11th May 2021