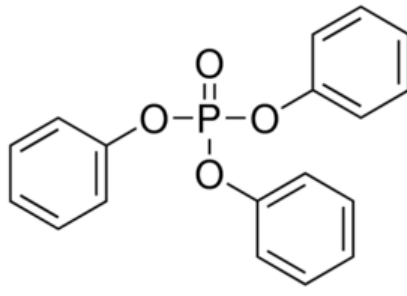


CAS 115-86-6

# Triphenyl phosphate (TPP)

(C<sub>6</sub>H<sub>5</sub>O)<sub>3</sub>PO



## Summary of Health Effects

TPP may harm development or the reproductive or neurological systems, based on animal studies.

## How is TPP used?

TPP is primarily used as a plasticizer in hydraulic fluids, varnishes and lacquers (including nail polish) and as a flame-retardant chemical in polyvinyl chloride (PVC) products, electronics, glues, casting resins, and hydraulic fluids.<sup>1,2</sup> TPP is a component of the commercial flame-retardant chemical Firemaster 550.<sup>2</sup>

## Toxicity: What are its health effects?

Disruption of the endocrine system, developmental, neurological, and reproductive toxicity occurred in animals fed TPP or their offspring.<sup>1,3-8</sup> The U.S. Environmental Protection Agency characterized TPP to have a moderate potential for carcinogenicity and bioaccumulation based on modeling.<sup>9</sup> Mice fed TPP for 35 days showed oxidative liver stress, testicular tissue damage, and decreased testicular testosterone levels, testes weight, and testosterone synthesis related gene

expression.<sup>10</sup> TPP was shown to be a moderate androgen-receptor binder and estrogen receptor agonist in *in vitro* testing.<sup>9</sup> *In vitro* testing has also showed TPP to be a mitochondrial activity inhibitor.<sup>6,11</sup> Metabolic disruption occurred in offspring of rats exposed to a commercial mixture containing TPP.<sup>12</sup>

## Exposure: How can a person come in contact with it?

A person may come in contact with TPP by breathing in, eating, or skin contact with contaminated dust, or from skin contact or from skin contact with consumer products containing TPP. TPP has been detected throughout the environment in air, household dust, surface water, soil and sediment.<sup>13-22</sup>

A TPP metabolite was found through biomonitoring in North American human urine,<sup>17,23-25</sup> and in pregnant women in China.<sup>26</sup> An Indiana study detected TPP in hair and nails of young adults.<sup>27</sup> A TPP metabolite was detected in human urine after application of nail polish.<sup>1</sup> TPP was detected in breast milk in Swedish and Asian studies.<sup>28,29</sup>

## References

1. Mendelsohn, E., Hagopian, A., Hoffman, K., Butt, C.M., Lorenzo, A., Congleton, J., Webster, T.F., Stapleton, H.M. (2015). Nail polish as a source of exposure to triphenyl phosphite. *Environment International*, 86, 45–51.
2. Toxicology Excellence for Risk Assessment (TERA) (2015). *Environmental concentrations and consumer exposure data for selected flame retardants (TDCPP, TCPP, TEP, TPP)*. June 1, 2015: Consumer Product Safety Commission contract Number CPSC-D-12-0001. Retrieved from [www.cpsc.gov/s3fs-public/pdfs/CPSC%2520Staff%2520Statement%2520on%2520Toxicology%2520ExcellenceRiskAssessmentsReportExposureDataSelectedFlameRetardants.pdf](http://www.cpsc.gov/s3fs-public/pdfs/CPSC%2520Staff%2520Statement%2520on%2520Toxicology%2520ExcellenceRiskAssessmentsReportExposureDataSelectedFlameRetardants.pdf)
3. Boris, V., Krivoshev, F.D., Covaci, A., Blust, R., Husson, S.J., (2016). Assessing in-vitro estrogenic effects of currently used flame retardants. *Toxicology in Vitro*, 33, 153-162.
4. Du, Z., Zhang, Y., Wang, G., Peng, J., Wang, Z., & Gao, S. (2016). TPhP exposure disturbs carbohydrate metabolism, lipid metabolism, and the DNA damage repair system in zebrafish liver. *Scientific Reports*, 6, 21827. doi: 10.1038/srep21827.
5. Green, A.J., Graham, J.L., Gonzalez, E.A., La Frano, M.R., Petropoulou, S.E., Park, J.S., Newman, J.W., Stanhope, K.L., Havel, P.J., La Merrill, M.A. (2016). Perinatal triphenyl phosphite exposure accelerates type 2 diabetes onset and increases adipose accumulation in UCD-type 2 diabetes mellitus rats. *Reproductive Toxicology*, 68, 119-129. doi: 10.1016/j.reprotox.2016.07.009.
6. Behl, M., Hsieh, J.H., Shafer, T.J., Mundy, W.R., Rice, J.R., Boyd, W.A., Freedman, J.H., Hunter, E.S., Jarema, K.A., Padilla, S., Tice, R.R. (2015). Use of alternative assays to identify and prioritize organophosphorus flame retardants for potential developmental and neurotoxicity. *Neurotoxicology and Teratology*, 52, 181–193.
7. Liu, X., Ji, K., Choi, K. (2012). Endocrine disruption potentials of organophosphate flame retardants and related mechanisms in *H295R* and *MVLN* cell lines and in zebrafish. *Aquatic Toxicology*, 114-115, 173-181.
8. Liu, X., Ji, K., Jo, A., Moon, H., & Choi, K. (2013). Effects of TDCPP or TPP on gene transcriptions and hormones of HPG axis, and their consequences on reproduction in adult zebrafish (*Danio rerio*). *Aquatic Toxicology*, 134-135, 104-111.
9. EPA (2015). *Flame retardants used in flexible polyurethane foam: An alternatives assessment update*. U.S. Environmental Protection Agency. Retrieved from [www.epa.gov/sites/production/files/2015-08/documents/ffr\\_final.pdf](http://www.epa.gov/sites/production/files/2015-08/documents/ffr_final.pdf)
10. Chen, G., Jin, Y., Wu, Y., Liu, L., Fu, Z. (2015). Exposure of male mice to two kinds of organophosphate flame retardants (OPFRs) induced oxidative stress and endocrine disruption. *Environmental Toxicology and Pharmacology*, 40(1), 310-8.
11. Behl, M. & Smith, M.V. (2016). Comparative toxicity of organophosphate flame retardants and polybrominated diphenyl ethers to *Caenorhabditis elegans*. *Toxicol Sciences*, 154(2), 241-252.
12. Patisaul, H.B., Roberts, S.C., Mabrey, N., McCaffrey, K.A., Gear, R.B., Braun, J., Belcher, S.M., Stapleton, H.M. (2013). Accumulation and endocrine disrupting effects of the flame retardant mixture Firemaster(R) 550 in rats: an exploratory assessment. *Journal of Biochemical and Molecular Toxicology*, 27(2), 124-36.
13. Meeker, J.D. & Stapleton, H.M. (2010). House dust concentrations of organophosphate flame retardants in relation to hormone levels and semen quality parameters. *Environmental Health Perspectives*, 118(3), 318-23.
14. van der Veen, I. & de Boer, J. (2012). Phosphorus flame retardants: properties, production, environmental occurrence, toxicity and analysis. *Chemosphere*, 88(10), 1119-53.
15. He, R., Li, Y., Xiang, P., Li, C., Zhou, C., Zhang, S., Cui, X., Ma, L.Q. (2015). Organophosphorus flame retardants and phthalate esters in indoor dust from different microenvironments: Bioaccessibility and risk assessment. *Chemosphere*, 150, 528-535. doi: 10.1016/j.chemosphere.2015.10.087.
16. Abdallah, M.A., & Covaci, A. (2014). Organophosphate flame retardants in indoor dust from Egypt: Implications for human exposure. *Environmental Science & Technology*, 48(9), 4782-4789.

17. Cequier, E., Skhi, A.K., Marce, R.M., Becher, G., Thomsen, C. (2015). Human exposure pathways to organophosphate triesters - a biomonitoring study of mother-child pairs. *Environment International*, 75, 159-65.
18. Fan, X., Kubwabo, C., Rasmussen, P.E., Wu, F. (2014). Simultaneous determination of thirteen organophosphate esters in settled indoor house dust and a comparison between two sampling techniques. *Science of the Total Environment*, 491-492, 80-6.
19. Salamova, A., Ma, Y., Venier, M., Hites, R.A. (2014). High levels of organophosphate flame retardants in the Great Lakes atmosphere. *Environmental Science & Technology Letters*, 1(1), 8-14.
20. Cao, S., Zeng, X., Song, H., Li, H., Yu, Z., Sheng, G., Fu, J. (2012). Levels and distributions of organophosphate flame retardants and plasticizers in sediment from Taihu Lake, China. *Environmental Toxicology and Chemistry*, 31(7), 1478-84.
21. Dodson, R.E., Perovich, L.J., Covaci, A., Van den Eede, N., Ionas, A.C., Dirtu, A.C., Brody, J.G., Rudel, R.A. (2012). After the PBDE phase-out: A broad suite of flame retardants in repeat house dust samples from California. *Environmental Science & Technology*, 46, 13056–13066.
22. EPA. (2015). *TSCA Work plan chemical problem formulation and initial assessment - chlorinated phosphate ester cluster flame retardants*. Environmental Protection Agency. Retrieved from [www.epa.gov/sites/production/files/2015-09/documents/cpe\\_fr\\_cluster\\_problem\\_formulation.pdf](http://www.epa.gov/sites/production/files/2015-09/documents/cpe_fr_cluster_problem_formulation.pdf)
23. Meeker, J.D., Cooper, E.M., Stapleton, H.M., Hauser, R. (2013). Urinary metabolites of organophosphate flame retardants: temporal variability and correlations with house dust concentrations. *Environmental Health Perspectives*, 121(5), 580-5.
24. Butt, C.M., Congleton, J., Hoffman, K., Fang, M., Stapleton, H.M. (2014). Metabolites of organophosphate flame retardants and 2-ethylhexyl tetrabromobenzoate in urine from paired mothers and toddlers. *Environmental Science & Technology*, 48(17), 10432-8.
25. Centers for Disease Control and Prevention (CDC) (2015). *Fourth national report on human exposure to environmental chemicals: Updated tables February 2015*. U.S. Department of Health & Human Services. Atlanta, GA. Retrieved from [www.cdc.gov/exposurereport](http://www.cdc.gov/exposurereport)
26. Liping F., Fengxiu O., Liangpo L., Xu, W., Xia, W., Yi-Ju, L., Amy, M., Heqing, S., Jungfeng, Z., Jun Jim, Z. (2016). Levels of urinary metabolites of organophosphate flame retardants, TDCIPP, and TPHP, in pregnant women in Shanghai. *Journal of Environmental and Public Health*, Article ID 9416054. doi:10.1155/2016/9416054.
27. Liang-Ying Liu, K.H., Hites, R.A., & Salamova, A. (2016). Hair and nails as noninvasive biomarkers of human exposure to brominated and organophosphate flame retardants. *Environmental Science & Technology*, 50, 3065–3073.
28. Kim, J.W., Isobe, T., Muto, M., Tue, N.M., Katsura, K., Malarvannan, G., Sudaryanto, A., Chang, K.H., Prudente, M., Viet, P.H., Takahashi, S., Tanabe, S. (2014). Organophosphorus flame retardants (PFRs) in human breast milk from several Asian countries. *Chemosphere*, 116, 91-7.
29. Sundkvist, A.M., Olofsson, U., and Haglund, P. (2010). Organophosphorus flame retardants and plasticizers in marine and fresh water biota and in human milk. *J Journal of Environmental Monitoring.*, 12(4), 943-51.