

Guidelines/Criteria	
Reference:	<p>NTP. 1993a. NTP Technical Report on toxicity studies of a chemical mixture of 25 groundwater contaminants. NTP toxicity report series no. 35 - NIH Publication 93-3384. National Toxicology Program, National Institute of Environmental Health Sciences (NIEHS), Research Triangle Park, NC, USA</p> <p>Hong HL, Yang RSH, Boorman GA. 1991. Residual damage to hematopoietic system in mice exposed to a mixture of groundwater contaminants. <i>Toxicol Lett</i> 57:101-111.</p> <p>Hong HL, Yang RSH, Boorman GA. 1992. Alterations in hematopoietic responses in B6C3F1 mice caused by drinking a mixture of 25 groundwater contaminants. <i>J Environ Pathol Toxicol Oncol</i> 11(2):1-10.</p>
In vivo study type Route of Administration Species & age of animals	<p>drinking water, water control group</p> <p>B6C3F1 mice, 10 weeks</p>
Study Duration	2.5, 15.5, 31.5 weeks and 108 days with 2-days and 10 weeks of recovery
Type of Mixture Binary >2 components Similar acting or dissimilar What Mode of Action was investigated?	<p>no</p> <p>yes</p> <p>dissimilar action assumed (chemicals mix to simulate groundwater supplies near hazardous dumps)</p> <p>haematotoxicity</p>
Parameters/End points Measured Target organs/Critical effects Pharmacological changes or adverse effects	<p>Haematopoietic system (whole body irradiation, haematology, bone marrow cellularity, progenitor (granulocyte macrophage) cell formation (CFU-GM), erythroid precursor formation (CFU-E)); water consumption, body, liver, spleen, kidney, thymus weights, histopathology of lung, heart, liver, kidney, adrenal glands, spleen thymus, stomach, uterus, bone marrow (sternum), urinary bladder, small and large intestines)</p> <p>adverse and non-adverse effects</p>
Individual Components Characterisation of individual compounds Name, exact chemical name, CAS no. Were dose responses established for individual components? Were no effect levels established? Were doses below the NO(A)ELs investigated?	<p>yes</p> <p>One mixture was investigated: Acetone, Aroclor 1260, Arsenic, Benzene, Cadmium, Carbon tetrachloride, Chloroform, Chlorobenzene, Chromium, 1,1-Dichloroethane, 1,1-Dichloroethylene, 1,2-Dichloroethane, 1,2-t-Dichloroethylene, Di(2-ethylhexyl)phthalate, Ethylbenzene, Lead, Mercury, Methylene chloride, Nickel acetate tetrahydrate, Phenol, Tetrachloroethylene, Toluene, 1,1,1-Trichloroethane, Trichloroethylen, Xylenes</p> <p>No, only mixtures at three dose levels were administered: Target dose (see below and 2 and 10 fold dilutions thereof)</p> <p>Yes</p> <p>Yes presumably</p>
Mixtures Investigated Number of dose levels How does the mixture make-up compare to individual components? (e.g. low dose) equivalents used? No. of technical replicates per exposure condition (<i>in vitro</i>) No. of animals per dose group (<i>in vivo</i>)	<p>Target concentrations: Acetone: 53 ppm, Aroclor 1260: 0.01 ppm, Arsenic: 9.0 ppm, Benzene: 12.5 ppm, Cadmium: 51.0 ppm, Carbon tetrachloride: 0.40 ppm, Chloroform: 7.0 ppm, Chlorobenzene: 0.10 ppm, Chromium: 36.0 ppm 1,1-Dichloroethane: 1.4 ppm, 1,1-Dichloroethylene: 0.5 ppm, 1,2-Dichloroethane: 40.0 ppm, 1,2-t-Dichloroethylene: 2.5 ppm, Di-2-ethylhexyl-phthalate: 0.015 ppm, Ethylbenzene: 0.3 ppm, Lead: 70 ppm, Mercury: 0.50 ppm, Methylene chloride: 37.5 ppm, Nickel: 6.80 ppm, Phenol: 29.0 ppm, Tetrachloroethylene: 3.40 ppm, Toluene: 7.0 ppm, 1,1,1-Trichloroethane: 2.0 ppm, Trichloroethylen: 6.50 ppm, Xylenes: 1.60 ppm</p> <p>Target concentrations: 375 ppm</p> <p>Would have to be evaluated, NOAELs of individual compounds are not given.</p> <p>not applicable</p> <p>8 mice/group</p>
Observations/Findings	<p>108 days</p> <p>Decreased water consumption in mid and high dose</p> <p>High dose: Increased rel. Kidney weights, decreased rel. Thymus weights no other changes observed</p> <p>Haematology: High dose: decreased MCV values</p> <p>Haematopoiesis: mid and high dose: decreased number of progenitor cells capable of forming granulocyte-macrophage colonies (CFU-GM) (reversible after 10 weeks); significantly slower recovery of bone marrow progenitor (even after 10 weeks of recovery)</p> <p>2.5, 15.5, 31.5 weeks exposures</p> <p>High dose: 15 days: rel. increased kidney and decreased thymus weight; 31 days: decreased rel. liver and thymus weights, thymus atrophy;</p> <p>mid and high dose: 15 and 31 days decreased MCV levels (at 15 days only in high dose), dose-related decrease in the numbers of CFU-GM</p>
Overall opinion (e.g. sufficient numbers of groups investigated, group sizes adequate, observations reproducible, low dose levels used investigated)	Questionable relevance of some parameters (CFU-GM...).