Triphenyl phosphate (CAS no. 115-86-6)

Tier 1 in vivo Test

(1) Results

Fish were exposed to concentrations of 2.13, 7.19, 17.1 and $44.9\mu g/L$ (measured). No significant differences were observed in fertility rate, male and female mortality, gonadosomatic index, secondary sex characteristics, male total length, body weight, hepatic vitellogenin and female hepatosomatic Index.

At 7.19 μ g/L and higher, a significant increase was observed in male hepatosomatic index, while a significant decrease was observed in female hepatic vitellogenin level.

A significant decrease was observed in number of eggs, number of fertile eggs, female total length and body weight at $44.9\mu g/L$.

(2) Summary

From the significant decrease observed in number of eggs, number of fertile eggs, female total length and body weight at $44.9 \mu g/L$ considered adverse reproductive effects on Medaka.

Estrogenic and/or antiandrogenic activity of triphenyl phosphate have been indicated from literature. In this study, an increase in male hepatic vitellogenin level was not observed at sublethal concentrations to suggest estrogenic effect. It was not concluded that triphenyl phosphate is an estrogenic compound. Regarding antiandrogenic activity, this study is not designed to detect it.

Since a significant decrease was observed in female hepatic vitellogenin level in this study, antiestrogenic activity should be tested in Tier 1 *in vitro* Test.

The adverse exposure level of $44.9\mu g/L$ was ca. 180 times as high as the highest environmental water concentration of $0.25\mu g/L$ that was measured in MOE's Environmental Survey and Monitoring of Chemicals in FY1975 (also detected in air in FY2007).

Table 1-A Results

Concentration (µg/L)		Number of fish		Mortality (%)		Total length (mm)		Body weight (mg)	
nominal	measure d	male	femal e	male	female	male	female	male	female
Control		12	12	0	0	38.0±0.51	36.5±0.50	543±45	512±37
20.0	2.13	12	12	0	0	37.9±1.1	35.3±0.65	532±44	449±19
64.0	7.19	12	12	0	0	38.0±1.1	35.5±0.28	544±49	464±27
200	17.1	12	12	0	0	37.7±1.0	35.2±1.0	539±31	468±34
640	44.9	12	12	0	8.3	38.3±1.1	34.4±0.35**	583±68	432±32*

Table 1-BResults (continued)

Measured	Measured Number of eggs		Fertility rate	Gonadosomatic Index (%)		
concentration	(eggs/female/day)	eggs	(%)	male	female	
(µg/L)		(eggs/female/day)				
Control	23.3±3.7	21.7±4.0	93.0±5.3	0.86±0.083	8.0±0.72	
2.13	20.4±2.7	18.3±3.1	89.3±4.9	0.87±0.17	7.9±0.47	
7.19	18.7±3.1	17.1±3.4	90.8±3.8	0.95 ± 0.092	8.1±0.62	
17.1	20.1±2.0	18.6±1.8	92.6±2.2	0.94 ± 0.098	7.7±0.72	
44.9	16.4±3.2*	14.9±3.5*	91.0±3.9	0.80 ± 0.067	9.4±1.3	

Table 1-C Results (continued)

Measured Hepatoson concentration		ic Index (%)	Vitellogenir	n (ng/mg liver)	Secondary sex characteristics	
(µg/L)	male	female	male	female	male	female
Control	1.2±0.19	2.7±0.52	9.1±5.4	764±220	88±7.5	0
2.13	1.4 ± 0.12	3.5±0.39	6.9 ± 4.0	542±115	83±7.4	0
7.19	1.7±0.29*	3.1±0.21	8.4±9.0	360±59.6**	84±6.9	0
17.1	1.7±0.24*	3.6±0.46	11.2±9.4	420±76.7*	83±7.2	0
44.9	1.7±0.13*	3.6±0.60	8.8 ± 8.1	417±70.3*	90 ± 8.0	0

Table 1-D Results (continued)

		Tuble I D	results (continued)	
Measured concentration			Other observations	
(µg/L)				
Control	Not found			
2.13	Not found			
7.19	Not found			
17.1	Not found			
44.9	Not found			

Data show mean \pm SD (standard deviation)

Statistically significant differences from control group (**p<0.01, *p<0.05) nd: not detected (below detection limit of vitellogenin: 1ng/mg liver) (-): not measured

Secondary sex characteristics: number of papillary processes