## 2,2',4,4'-Tetrahydroxybenzophenone (Benzophenone-2) (CAS no. 131-55-5)

## Tier 1 in vivo Test

## (1) Results

Fish were exposed to concentrations of 0.0943, 0.939, 9.53 mg/L (measured). No significant differences were observed in mortality, total length, body weight, total number of eggs, number of fertile eggs, gonadosomatic index, hepatosomatic index, secondary sex characteristics and female hepatic vitellogenin level.

At 9.53 mg/L, a significant increase in male hepatic vitellogenin level and a significant decrease in fertility rate was observed.

## (2) Summary

Estrogenic activity of 2,2',4,4'-tetrahydroxybenzophenone (benzophenone-2) has been indicated from literature. In this study, an increase in male hepatic vitellogenin level was observed at sublethal concentrations indicating its estrogenic effect. It was concluded that 2,2',4,4'-tetrahydroxybenzophenone (benzophenone-2) is an estrogenic compound.

The adverse exposure level indicating of 9.53mg/L was ca. 733,000 times as high as the highest environmental water concentration of  $0.013\mu$ g/L that was measured in MOE's Environmental Survey and Monitoring of Chemicals in FY2014.

The no observed adverse effect level of 0.939 mg/L was ca. 72,200 times as high as the detection limit of environmental water concentration of 0.013  $\mu$ g/L for MOE's Environmental Survey and Monitoring of Chemicals in FY2014.

|                                     |                             |                  | Table I-A         | Results  |                  |              |
|-------------------------------------|-----------------------------|------------------|-------------------|----------|------------------|--------------|
| Measured<br>concentration<br>(mg/L) | Number of survived fish     | Mortality<br>(%) | Total length (mm) |          | Body weight (mg) |              |
|                                     | male and male<br>female fem | male and female  | male              | female   | male             | female       |
| Control (<0.0199)                   | 24                          | 0                | 30.9±7.0          | 30.1±1.0 | 288±38           | 317±39       |
| 0.0943                              | 24                          | 0                | 30.8±1.2          | 30.2±0.4 | 291±27           | 296±16       |
| 0.939                               | 23                          | 4.2              | 30.6±1.1          | 30.3±1.0 | 290±32           | $304 \pm 48$ |
| 9.53                                | 24                          | 0                | 31.9±1.1          | 30.4±0.5 | 327±34           | 295±8        |

Table 1-A Results

Table 1-B Results (continued)

| Measured                | Number of eggs    | Number of fertile         | Fertility rate<br>(%) | Gonadosomat       | Gonadosomatic Index (%) |  |
|-------------------------|-------------------|---------------------------|-----------------------|-------------------|-------------------------|--|
| concentration<br>(mg/L) | (eggs/female/day) | eggs<br>(eggs/female/day) |                       | male              | female                  |  |
| Control (<0.0199)       | 14.4±1.8          | 13.6±1.6                  | 94.6±1.4              | 0.856±0.107       | 7.50±1.38               |  |
| 0.0943                  | 15.4±2.3          | 14.6±2.2                  | 94.5±2.3              | $1.07 \pm 0.14$   | 6.93±0.34               |  |
| 0.939                   | 14.5±2.7          | 13.6±2.3                  | 94.3±2.5              | 0.692±0.154       | 7.56±1.20               |  |
| 9.53                    | 13.8±1.7          | 12.5±1.7                  | 89.7±3.5*             | $0.790 \pm 0.226$ | 8.56±0.41               |  |

Table 1-C Results (continued)

| Measured concentration | Hepatosoma | tic Index (%)   | Vitellogenin (ng/mg liver) |         | Secondary sex characteristics |         |
|------------------------|------------|-----------------|----------------------------|---------|-------------------------------|---------|
| (mg/L)                 | male       | female          | male                       | female  | male                          | female  |
| Control (<0.0199)      | 1.93±0.28  | 3.36±0.42       | 0.567±0.133                | 345±152 | 79.8±4.8                      | 0±0     |
| 0.0943                 | 1.79±0.37  | $2.83 \pm 0.87$ | nd                         | 354±105 | 72.6±7.0                      | $0\pm0$ |
| 0.939                  | 1.89±0.24  | 3.08±0.68       | $0.888 \pm 0.460$          | 398±230 | 77.0±19.8                     | $0\pm0$ |
| 9.53                   | 1.95±0.55  | $3.48 \pm 0.58$ | 393±306*                   | 449±114 | 78.8±8.4                      | 0±0     |

Table 1-D Results (continued)

| Measured          |                    |  |
|-------------------|--------------------|--|
| concentration     | Other observations |  |
| (mg/L)            |                    |  |
| Control (<0.0199) | Not found          |  |
| 0.0943            | Not found          |  |
| 0.939             | Not found          |  |
| 9.53              | Not found          |  |

Note. The test compound was a pale yellow-green to red-yellow crystalline power.

Data show mean ± 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 joint plates with papillary processes