

US EPA ARCHIVE DOCUMENT



Endocrine Disruptors and Growth in the Tilapia, a Euryhaline Fish

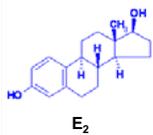
Background

Endocrine disruptors (ED) are compounds that affect normal reproduction, development, and behavior, but little is known about their effects on growth.

Vertebrate growth is regulated by the growth hormone (GH) and insulin-like growth factor (IGF-I) axis.



The primary objective of my study is to examine *in vitro* effects of heptachlor, an estrogenic pesticide, on the GH-IGF-I axis, using estradiol-17 β (E₂) as a positive control.



Methods

Hepatocytes of the male tilapia (*Oreochromis mossambicus*) were isolated by collagenase digestion.

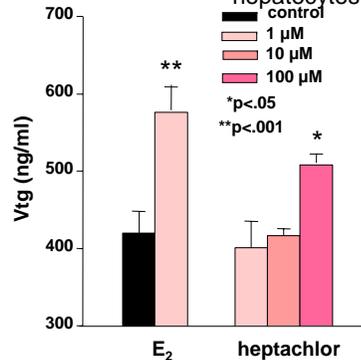
Hepatocytes were dispersed, plated, and treated with 1 nM E₂ or 1-100 μ M heptachlor for 48 h.

Culture media was analyzed for secretion of the yolk precursor protein, vitellogenin, using ELISA.

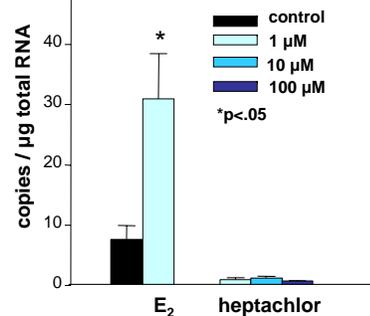
Total RNA was isolated for analyses of gene expression of IGF-I and GH-receptor (GH-R) via real-time PCR.

Results

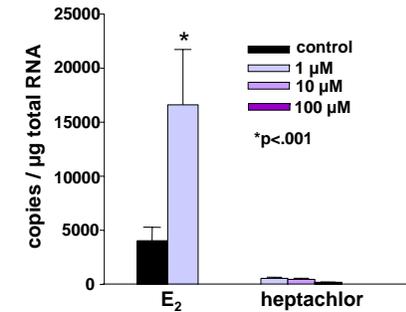
Heptachlor induces a dose-dependent increase in vitellogenin (Vtg) secretion in hepatocytes.



Heptachlor reduces IGF-I gene expression in hepatocytes.



Heptachlor reduces GH-R gene expression in hepatocytes



Conclusions & Future Plans

- Heptachlor stimulated Vtg secretion dose dependently in tilapia hepatocytes.
- E₂ stimulated both IGF-I and GH-R mRNA expression at 1 nM.
- Unexpectedly, however, heptachlor suppressed expression of IGF-I and GH-R genes at all concentrations tested, suggesting that E₂ and heptachlor use different mechanisms in interacting with the GH-IGF-I axis.
- Further studies will be done to clarify these differences, in part by using other EDs such as DDE and PCB #138 and analyzing expression of the three different forms of Vtg.
- In vitro* culture of pituitary glands with EDs will also be done to see how EDs alter normal pituitary activity and responsiveness. In addition, long-term *in vivo* studies will be carried out to examine how EDs affect growth regulation at whole animal level.

Many thanks to Drs. Riley, Hirano, and Grau for their guidance.

