

## INTERNATIONAL SUMMIT ON DIABETES, ENDOCRINOLOGY, AND METABOLIC DISORDERS



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### Expression of the IL-1 $\beta$ and CX3CR1 pancreatic $\beta$ -cells of mice fed with a high-fat/high-sucrose diet

#### Abstract

#### Background/Objectives

Increased circulatory glucose and free fatty acids lead to glucolipotoxicity, which triggers inflammation, affecting the endocrine pancreatic  $\beta$ -cells. We investigated the comparative effects of a fish-oil-based-high-fat, high-sucrose-diet (Fish-HFHSD) versus a cocoa-butter-based-high-fat, high-sucrose-diet (Cocoa-HFHSD) on the expression of the inflammatory cytokine interleukin (IL)-1 $\beta$  and fractalkine receptor (CX3CR1) in mouse  $\beta$ -cells.

#### Methods

C57BL/6 male mice (n=18) were randomly assigned to three dietary interventions, including Chow, Fish-HFHSD, and Cocoa-HFHSD, for 22-weeks. Pancreatic tissues were collected for immunohistochemistry to quantify insulin-, IL-1 $\beta$ -, and CX3CR1-positive-areas.

#### Results

The immunostaining intensity of insulin in  $\beta$ -cells was significantly higher (P=0.0008) in mice fed with either Fish-HFHSD (180 $\pm$ 10px GV) or Chow (173 $\pm$ 6px GV) compared to those fed with Cocoa-HFHSD (118 $\pm$ 12px GV). Conversely, the immunostaining intensity of IL-1 $\beta$  was significantly higher (P=0.0004) in mice fed with Cocoa-HFHSD (237 $\pm$ 12px GV) compared to those fed with Fish-HFHSD (145 $\pm$ 13px GV) or Chow (180 $\pm$ 10px GV).

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As expected, the IL-1 $\beta$ :insulin ratio was significantly lower ( $P<0.0001$ ) in Fish-HFHSD ( $0.80\pm0.04$ ) and Chow ( $0.82\pm0.06$ ) fed mice compared to that ( $2.11\pm0.22$ ) in Cocoa-HFHSD fed mice. Unlike IL-1 $\beta$ , CX3CR1 immunostaining intensity was significantly higher ( $P<0.0001$ ) in mice fed with either Fish-HFHSD ( $214\pm10$ px GV) or Chow ( $209\pm9$ px GV) compared to those fed with Cocoa-HFHSD ( $111\pm12$ px GV). In addition, the CX3CR1:insulin ratio was significantly higher ( $P<0.0001$ ) in Fish-HFHSD ( $1.17\pm0.02$ ) and Chow ( $1.2\pm0.02$ ) fed mice compared to that ( $0.98\pm0.01$ ) in Cocoa-HFHSD fed mice.

### Conclusions

In contrast to Fish-HFHSD feeding, Cocoa-HFHSD feeding induces a metabolic insult in mice that results in increased IL-1 $\beta$  expression and reduced insulin and CX3CR1 expression in pancreatic  $\beta$ -cells.