Supplemental Information

## *Title*: Isothiocyanates induce UGT1A1 in humanized *UGT1* mice in a CAR dependent fashion that is highly dependent upon oxidative stress

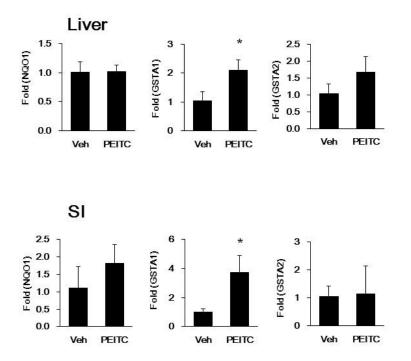
Short title: Cruciferous vegetables and activation of nuclear receptors

Emiko Yoda <sup>1,2</sup>, Miles Paszek<sup>1</sup>, Camille Konopnicki<sup>1</sup>, Ryoichi Fujiwara<sup>3</sup>, Shujuan Chen<sup>1</sup> and Robert H. Tukey<sup>1,\*</sup>

<sup>1</sup>Laboratory of Environmental Toxicology, Department of Pharmacology, University of California, San Diego, La Jolla, CA 92093.

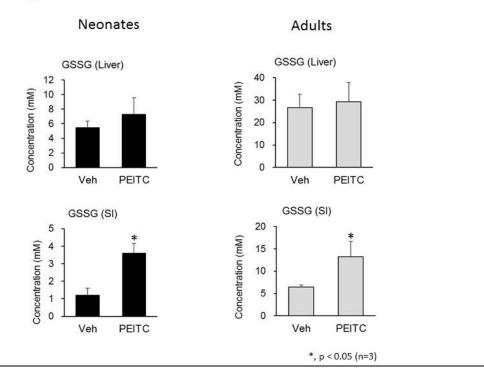
<sup>2</sup>Division of Health Chemistry, Department of Healthcare and Regulatory Sciences, School of Pharmacy, Showa University, 1-5-8 Hatanodai, Shinagawa-ku, Tokyo 142-8555, Japan. <sup>3</sup>Department of Pharmaceutics, School of Pharmacy, Kitasato University, 6-9-1 Shirokane, Minato-ku, Tokyo 108-8641, Japan.

## Supplemental Figure 1



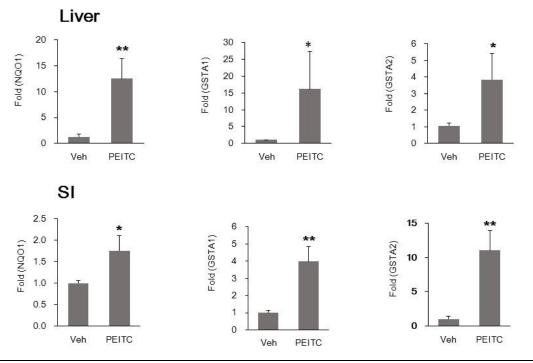
**Supplemental Figure 1**. Analysis of Nrf2 target genes. Using RNA isolated in Figure 1 (neonatal hUGT1 mice), gene expression patterns for Nqo1, Gsta1 and Gsta2 were quantitated by RT-qPCR expressed as Fold induction. Values are the means of ± SD (n=3). Statistically significant differences between vehicle (Veh) and PEITC are indicated by asterisks (Student t test: \*, P<0.05).

## Supplemental Figure 2



**Supplemental Figure 2.** Oxidative stress following PEITC treatment. **A**. Neonatal hUGT1 mice were given an oral dose of PEITC (200 mg/kg) and GSSG levels measured in liver and small intestine (SI) after 24 hours. **B**. The same experiment was conducted on adult (6 week old) hUGT1 mice. Values are the means of  $\pm$  SD (n=3). Statistically significant differences between vehicle (Veh) and PEITC are indicated by asterisks (Student *t* test: \*, *P*<0.05).

## Supplemental Figure 3



**Supplemental Figure 3**. Impact of PEITC on Nrf2 target genes in adult mice. RNA prepared from PEITC treated  $hUGT1/Car^{+/-}$  and  $hUGT1/Car^{-/-}$  mice from liver and small intestine (Figure 4) was used to quantitate Nqo1, Gsta1 and Gsta2 gene expression by real time PCR. Values are the means of ± SD (n>3). Statistically significant differences between vehicle (Veh) and PEITC are indicated by asterisks (Student t test: \*, P<0.05; \*\*, P<0.01).

