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### ACCEPTED MANUSCRIPT

# Accurate quantification of PGE<sub>2</sub> in the polyposis in rat colon (Pirc) model by surrogate analyte-based UPLC-MS/MS

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#### **Highlights**

- The colon PGE<sub>2</sub> of normal (wild type) rat and Pirc rat (an *Apc*-mutant rat) were firstly accurately determined by specific and sensitive UPLC-MS/MS.
- PGE<sub>2</sub> was confirmed to promote Pirc rat colon polyps growth.
- An unusual mobile phase (0.1% ammonia hydroxide) was found to greatly improve the chromatographic separation of prostaglandins.

#### **Abstract**

An accurate and reliable UPLC-MS/MS method is reported for quantification of endogenous Prostaglandin E2 (PGE<sub>2</sub>) in rat colon mucosa and polyps. This method adopted the "surrogate analyte plus authentic bio-matrix" approach, using two different stable isotopic labeled analogs — PGE<sub>2</sub>-d9 as the surrogate analyte and PGE<sub>2</sub>-d4 as the internal standard. Quantitative standard curve was constructed with the surrogate analyte in colon mucosa homogenate; and the method was also successfully validated with the authentic bio-matrix. Concentrations of endogenous PGE<sub>2</sub> in both normal and

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