**Professor Barbara P. Rattner**

**Executive Editor**

***Epigenetics***

30th July, 2013

Dear Professor Rattner,

I would like to submit a presubmission inquiry regarding suitability of our manuscript, “Diagnostic Role of APC Promoter Methylation in Non-Small Cell Lung Cancer (NSCLC): An Integrate-Analysis of Published Articles and Microarray Data” by Shicheng Guo, Lixing Tan, Weilin Pu, Junjie Wu, Kuan Xu, Jinhui Wu, Qiang Li, Yanyun Ma, Jibin Xu, Li Jin\*, Jiucun Wang\* for publication in ***Epigenetics***.

APC is one of most important human tumor suppressor gene and has been considered as an important biomarker for early diagnosis of cancer. However, there was still not yet any quantitative assessment for the sensitivity and specificity of the diagnostic performance in NSCLC.

In this report, we made a comprehensively assessment on the diagnosis ability of APC promoter hypermethylation in NSLCL with a combination of Meta-analysis and independent Microarray dataset from TCGA project. Three credible heterogeneity sources to the association between APC methylation and NSCLC were identified, including primer set, heterogeneous or autogenous controls, proportion of adenocarcinoma in NSCLC samples. The present study shows that the methylation status of APC would be an excellent diagnosis biomarker in remote non-invasive media. In addition, APC methylation test has better diagnosis ability in adenocarcinoma than that in squamous cell carcinoma. The study made a professional analysis for the diagnostic ability of APC methylation with summary receiver operating characteristic curve (SROC) and independently validation with substantial methylation microarray dataset.

*Epigenetics* would be an outstanding forum for this paper due to its intention of featuring DNA methylation research publications. We believe that this paper will be of interest to epigenetic epidemiologists, due to the double validation analysis method, and to medical and clinical oncologists , due to its interesting findings, such as heterogeneity discovery and APC methylation diagnostic role in adenocarcinoma but squamous cell carcinoma.

I have attached the full-length manuscript. Thank you in advance for considering this pre-submission inquiry.

Sincerely,

Shicheng Guo