Reviewer's report

**Title:** The Potential Interaction Between Clopidogrel and Proton Pump Inhibitors: A Systematic Review

**Version:** 1 **Date:** 16 October 2010

**Reviewer:** Deepak Bhatt

**Reviewer's report:**

Minor Essential Revisions

This is an excellent systematic review of whether there is or is not an interaction between clopidogrel and PPIs. The authors wisely refrain from pooling these very heterogeneous data. They correctly realize that residual confounding is the most likely explanation for the very large effects seen in observational studies of whether there is a clopidogrel-PPI interaction. They also wisely note that if such an interaction were present, it should be most obvious early after stenting or PCI, when clopidogrel provides the majority of its absolute benefits (though I might argue that it is more than just 3 months as the authors contend). This paper is very well-written and quite thoughtful and injects some common sense (as well as basic epidemiological principles) into the debate about this putative interaction.

1. Reference 25 should be updated, as COGENT has now been published in NEJM. Also, since COGENT has now been peer-reviewed, this review should upgrade its status to high quality.

2. Would consider adding the recent, large analysis by Charlot in Annals of Internal Medicine. The present authors wrote a nice editorial for that article so they should be quite familiar with it.

**Quality of written English:** Acceptable

**Statistical review:** No, the manuscript does not need to be seen by a statistician.

**Declaration of competing interests:**

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