Author's response to reviews

Title: Genetic Background Determines Response to Hemostasis and Thrombosis

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Re: Submission of Revised Manuscript #6056319367688982

Dear Dr. Pemberton,

We are submitting to the journal *BMC Blood Disorders* the revised manuscript entitled: *Genetic Background Determines Response to Haemostasis and Thrombosis*. Jane Hoover-Plow, Aleksey Shchurin, Erika Hart, and Jingfeng Sha, Annie E. Hill, Jonathan B. Singer, Joseph H. Nadeau are now co-authors of this manuscript.

Additional experiments have been performed and we have addressed the comments of the reviewer and made changes in the manuscript. The manuscript is substantially improved since the first submission. We look forward to a favorable decision.

All authors concur with the submission of this manuscript. This is an original work (research article), has not been reported previously, and is not under consideration elsewhere. None of the authors have any financial interest in relation to the submission. The library at this institution is a member of BioMed Central.
Thank you for your consideration of this manuscript for publication in *BMC Blood Disorders*

Sincerely yours,

Jane Hoover-Plow, PhD

**Response to Reviewers**

**Reviewer 1** had no comments.

**Reviewer 2**

The authors thank the reviewer for the constructive comments. We have made the suggested changes.

As suggested, Plg was measured in the CSS-5 x 17 strain and was not different from the B6 mice. In the CSS-17 F1 (heterosomic for Chr 17), Plg was also not different than for the B6 mice, suggesting the increased Plg was a recessive trait. In addition, as the reviewer points out, results also suggest that other genes are interacting in the CSS-5 x 17 cross for the expression of the occlusion time and PAI-1 activity. This is clarified in the text, p. 14, and p. 19. α2-antiplasmin values for CSS-11 initially had a high SD and additional mice were tested, and values for CSS-11 are significantly higher than for B6. In future studies additional crosses with the CSS will be investigated.

Statistical comparisons of CSSS were made only to B6 values and indicated in the text (Statistics), and figure legends.

On p. 19, the reference was to a congenic Chr 17 strain (V19H) that has only a fragment of the A/J Chr 17 that includes the Plg gene. The rest of the Chr 17 is from B6 mice and the background is from B6. In this congenic strain with the Chr 17 A/J fragment, no difference in bleeding time or rebleeding time was found compared to B6 mice. This is clarified in the text. p. 19.

The possibility that DNA sequences in the regulatory region may be important is added to the text, p. 20