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Supplemental Information

Developmental Stage-Specific Changes in Protein Synthesis Differentially Sensitize Hematopoietic Stem Cells and Erythroid Progenitors to Impaired Ribosome Biogenesis

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SUPPLEMENTAL INFORMATION

Developmental stage-specific changes in protein synthesis differentially sensitize hematopoietic stem cells and erythroid progenitors to impaired ribosome biogenesis

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Supplemental Figures and Legends

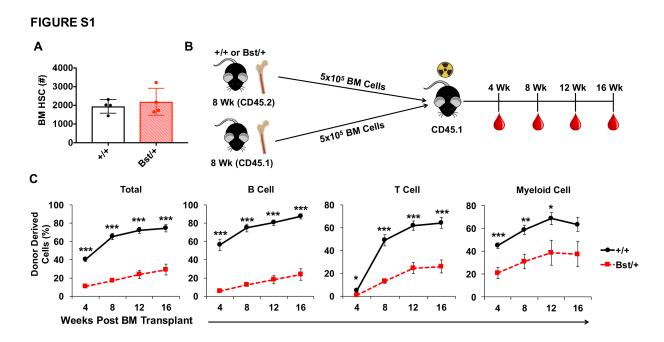


Figure S1. Young adult $Rpl24^{\text{Bst/+}}$ HSCs have impaired long-term multilineage reconstituting activity. Related to Figure 3. (A) Absolute number of HSCs in young adult $Rpl24^{\text{Bst/+}}$ and control mice (1 femur + 1 tibia/mouse; N=4 mice/genotype). (B) Diagram of experimental strategy to test long-term multilineage reconstituting activity of young adult $Rpl24^{\text{Bst/+}}$ HSCs. (C) Donor cell engraftment when $5x10^5 Rpl24^{\text{Bst/+}}$ (Bst/+) or littermate control (+/+) BM cells were transplanted with $5x10^5$ recipient-type young adult BM cells into irradiated mice. Total hematopoietic, B-, T-and myeloid cell engraftment is shown 4, 8, 12 and 16 weeks after transplantation (N=5 recipients/genotype). Data represent mean \pm SD (A) or SEM (C). Statistical significance was assessed using a two-tailed Student's t-test (*P<0.05, **P<0.01, ***P<0.001).

FIGURE S2

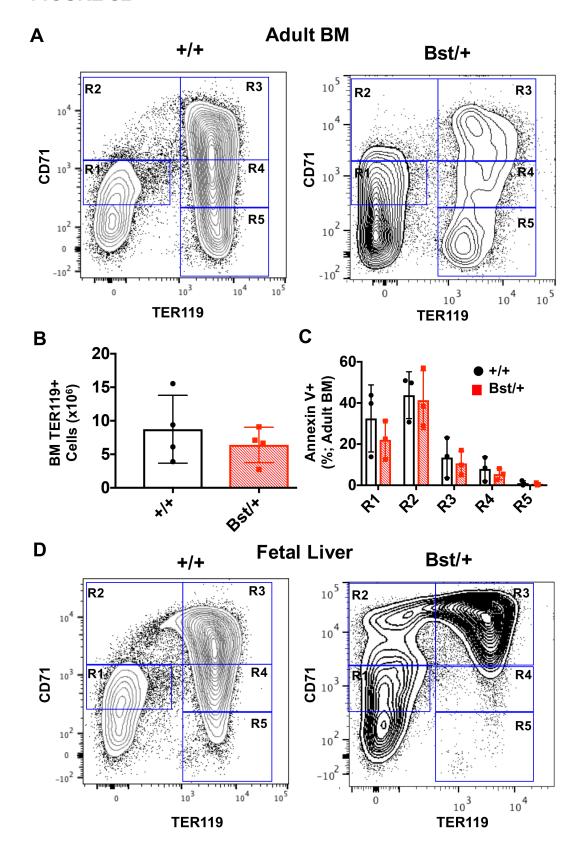


Figure S2. The $Rpl24^{\text{Bst/+}}$ mutation impairs fetal but not adult erythroid differentiation. Related to Figure 4. (A) Representative flow cytometry plots of R1-R5 erythroid lineage cells in $Rpl24^{\text{Bst/+}}$ (Bst/+) or control (+/+) young adult BM. (B) Number of TER119⁺ cells in $Rpl24^{\text{Bst/+}}$ (Bst/+) or control (+/+) young adult $Rpl24^{\text{Bst/+}}$ (Bst/+) or control (+/+) BM (1 femur + 1 tibia/mouse; n=4 mice/genotype). (C) Frequency of erythroid progenitors that are Annexin V⁺ in young adult $Rpl24^{\text{Bst/+}}$ (Bst/+) or control (+/+) BM (N=3 mice/genotype). (D) Representative flow cytometry plots of R1-R5 erythroid lineage cells in $Rpl24^{\text{Bst/+}}$ (Bst/+) or control (+/+) E15.5 fetal liver. Data represent mean \pm SD. Statistical significance was assessed using a two-tailed Student's t-test.