

# Hepatic Leukemia Factor supports the propagation of leukemia and hematopoietic stem cell function during stress-induced regeneration

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## What is the Manuscript Microscope Sentence Audit?

The Manuscript Microscope Sentence Audit is a research paper introspection system that parses the text of your manuscript into minimal sentence components for faster, more accurate, enhanced proofreading.

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- **Accelerated Proofreading:** Examine long technical texts in a fraction of the usual time.
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**Manuscript Source:** <https://www.biorxiv.org/content/10.1101/2021.03.05.434034v1>

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### Features of the Sentence Audit:

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The Minimal Sentence Components shown are the smallest coherent elements of each sentence of your text as derived from it's conjunctions, prepositions and selected punctuation symbols (i.e. commas, semicolons, round and square brackets).

The combined approaches ensure easier, faster, more effective proofreading.

### Comments and Caveats:

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- Depending on the source of the input text, the Sentence Audit may contain occasional html artefacts that are parsed as sentences (E.g. "Download figure. Open in new tab").
- Always consult the original research paper as the true reference source for the text.

### Contact Information:

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### Research Paper Sections:

The sections of the research paper input text parsed in this audit.

[illegible]

**Title**      **Hepatic Leukemia Factor supports the propagation of leukemia and hematopoietic stem cell function during stress-induced regeneration**

**S1 [001]      Abstract**

**S1 [002]**      The processes regulating hematopoietic stem cells (HSC) during aging are not fully understood<sup>1</sup>, but it is clear that the incidence of hematological malignancies increases with age, highlighting the importance of unravelling the cellular and molecular networks involved.

The processes regulating hematopoietic stem cells ...  
... (HSC) ...  
... during aging are not fully understood<sup>1</sup>, ...  
... but it is clear ...  
... that the incidence ...  
... of hematological malignancies increases ...  
... with age, ...  
... highlighting the importance ...  
... of unravelling the cellular ...  
... and molecular networks involved.

**S1 [003]**      Recently, we identified Hepatic Leukemia Factor (HLF) as an essential transcription factor in maintaining the HSC pool during regeneration<sup>2</sup> and showed that failure to downregulate HLF leads to disrupted differentiation<sup>3</sup>.

Recently, ...  
... we identified Hepatic Leukemia Factor ...  
... (HLF) ...  
... as an essential transcription factor ...  
... in maintaining the HSC pool ...  
... during regeneration<sup>2</sup> ...  
... and showed ...  
... that failure ...  
... to downregulate HLF leads ...  
... to disrupted differentiation<sup>3</sup>.

**S1 [004]**      Here, we found that HLF is dispensable for hematopoiesis during systemic aging, but needed during stress-induced hematopoietic recovery of aged HSC after transplantation.

Here, ...  
... we found ...  
... that HLF is dispensable ...  
... for hematopoiesis ...  
... during systemic aging, ...  
... but needed ...  
... during stress-induced hematopoietic recovery ...  
... of aged HSC ...  
... after transplantation.

**S1 [005]**      Additionally, HLF was dispensable for leukemic initiation but required for disease propagation.

Additionally, ...  
... HLF was dispensable ...  
... for leukemic initiation ...  
... but required ...  
... for disease propagation.

**S1 [006]**      Taken together, our findings demonstrate the existence of a HLF-dependent mechanism that uncouples stress-induced regeneration from hematopoietic homeostasis during aging, that can be used by malignant cells to gain stem cell properties to propagate the disease.

Taken together, ...  
... our findings demonstrate the existence ...  
... of a HLF-dependent mechanism ...  
... that uncouples stress-induced regeneration ...  
... from hematopoietic homeostasis ...  
... during aging, ...  
... that can be used ...  
... by malignant cells ...  
... to gain stem cell properties ...  
... to propagate the disease.

**S1 [007]**      Key points

Key points

**S1 [008]**      HLF is dispensable for HSC function and hematopoietic homeostasis during physiological aging, but crucial during stress induced regeneration.

HLF is dispensable ...  
... for HSC function ...  
... and hematopoietic homeostasis ...  
... during physiological aging, ...  
... but crucial ...  
... during stress induced regeneration.

**S1 [009]**      HLF supports the propagation of leukemia-initiating cells

HLF supports the propagation ...  
... of leukemia-initiating cells

## **S2 [010]      Methods**

**S2 [011]**      Mice

Mice

**S2 [012]** The generation of KO mice was previously described<sup>4</sup>, and mice were backcrossed to achieve pure C57BL/6 background.

The generation ...  
... of KO mice was previously described<sup>4</sup>, ...  
... and mice were backcrossed ...  
... to achieve pure C57BL/6 background.

**S2 [013]** Animals were housed in ventilated racks, given autoclaved food and water, and maintained in accordance with Swedish Animal Welfare organisation guidelines, at the Biomedical Center animal facilities in Lund.

Animals were housed ...  
... in ventilated racks, ...  
... given autoclaved food ...  
... and water, ...  
... and maintained ...  
... in accordance ...  
... with Swedish Animal Welfare organisation guidelines, ...  
... at the Biomedical Center animal facilities ...  
... in Lund.

**S2 [014]** All animal experiments were approved by local ethical committees (permit M94-15).

All animal experiments were approved ...  
... by local ethical committees ...  
... (permit M94-15).

**S2 [015]** Competitive transplantation assay

Competitive transplantation assay

**S2 [016]** For transplantation, 2 x10<sup>5</sup> unfractionated cells from BM from 18-month-old mice (CD45.2) were mixed in a 1:1 ratio with 2x10<sup>5</sup> unfractionated BM competitor cells (CD45.1).

For transplantation, ...  
... 2 x10<sup>5</sup> unfractionated cells ...  
... from BM ...  
... from 18-month-old mice ...  
... (CD45.2) ...  
... were mixed ...  
... in a 1:1 ratio ...  
... with 2x10<sup>5</sup> unfractionated BM competitor cells ...  
... (CD45.1).

**S2 [017]** For secondary transplant, a femur per donor was split into 2 recipients.

For secondary transplant, ...  
... a femur ...  
... per donor was split ...  
... into 2 recipients.

- S2 [018]**      Grafts were intravenously injected into lethally irradiated (900 cGy) recipients (CD45.1/CD45.2).
- Grafts were intravenously injected ...  
... into lethally irradiated ...  
... (900 cGy) ...  
... recipients ...  
... (CD45.1/CD45.2).
- S2 [019]**      Generation of MLL/AF9 leukemia
- Generation ...  
... of MLL/AF9 leukemia
- S2 [020]**      MLL/AF9 leukemia was generated as described in5 from either WT or KO cKit+ cells.
- MLL/AF9 leukemia was generated ...  
... as described in5 ...  
... from either WT ...  
... or KO cKit+ cells.
- S2 [021]**      HLF expression was established by semi-quantitative PCR as described in2.
- HLF expression was established ...  
... by semi-quantitative PCR ...  
... as described in2.
- S2 [022]**      Peripheral blood and bone marrow preparation
- Peripheral blood ...  
... and bone marrow preparation
- S2 [023]**      Peripheral blood (PB) was collected from the tail vein.
- Peripheral blood ...  
... (PB) ...  
... was collected ...  
... from the tail vein.
- S2 [024]**      Blood parameters were analyzed using SysmexXE-5000 (Sysmex Europe GmbH).
- Blood parameters were analyzed ...  
... using SysmexXE-5000 ...  
... (Sysmex Europe GmbH).
- S2 [025]**      Before staining, erythrocytes were lysed with NH4Cl (StemCell Technologies).
- Before staining, ...  
... erythrocytes were lysed ...  
... with NH4Cl ...  
... (StemCell Technologies).

## **End of Sample Audit**

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This is a truncated Manuscript Microscope Sample Audit.

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