

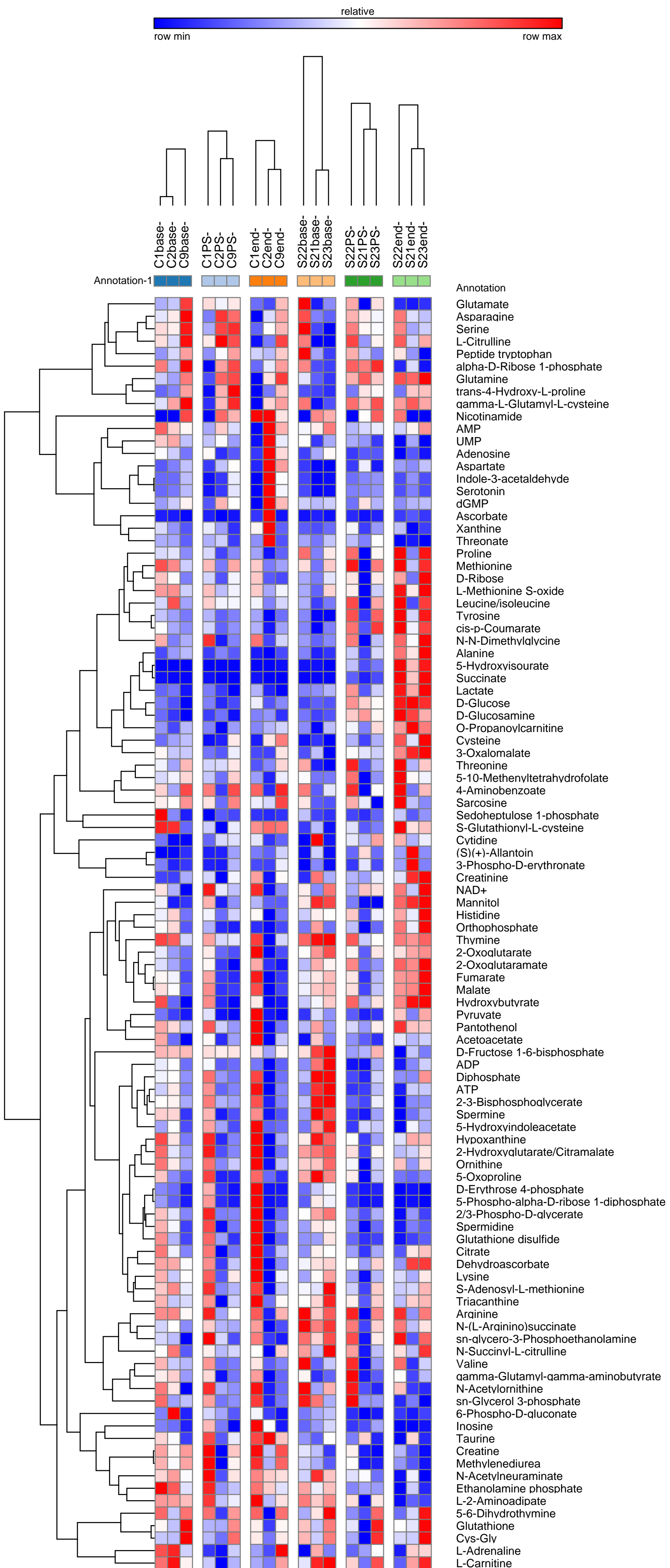
SUPPLEMENTARY MATERIAL

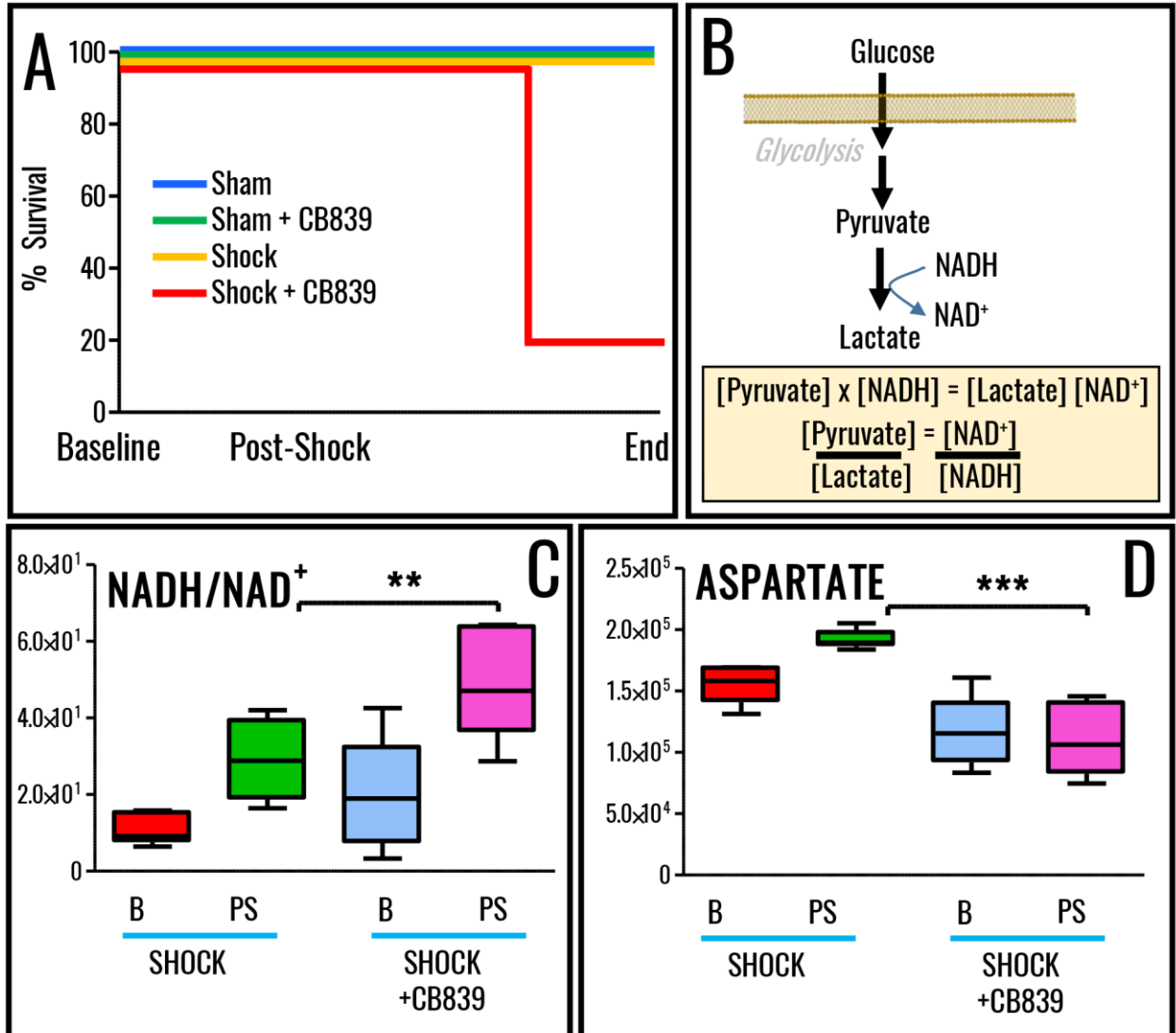
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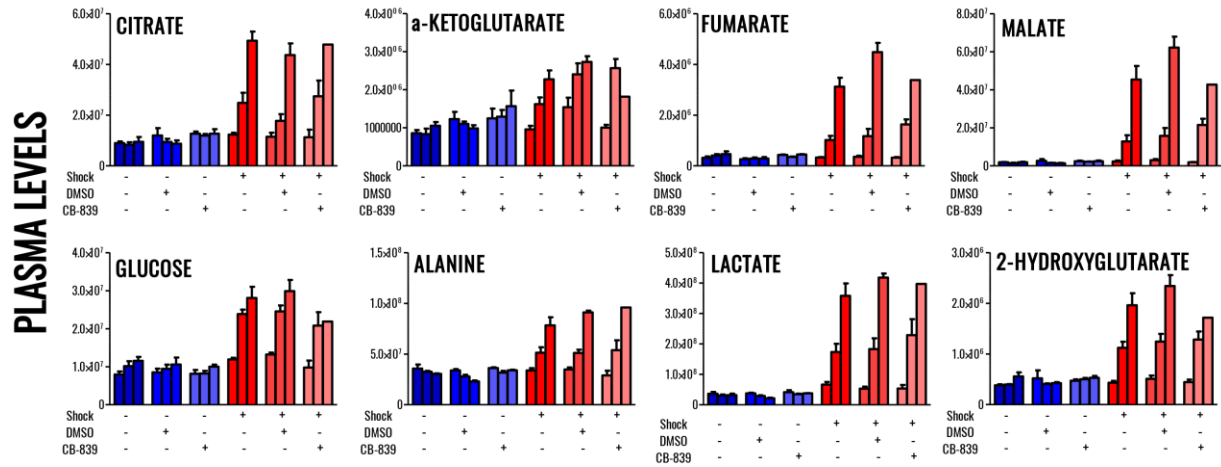
- Annotation-1
- Sham Baseline
- Sham End
- Sham Post-Shock
- Shock Baseline
- Shock End
- Shock Post-Shock

Supplementary Figure 1 – Hierarchical clustering analysis of metabolic measurements in RBCs from sham or hemorrhagic shock rats





Supplementary Figure 2 – Kaplan Meier curves indicate 80% mortality at 60 min from administration of the glutaminase inhibitor CB-839 in rats undergoing hemorrhagic shock (A). In these rats, RBC lactate/pyruvate ratios are proportional to NADH/NAD⁺ ratios (according to the formula in B). NADH/NAD⁺ ratios increase in response to the treatment with CB-839 in hemorrhagic shock rats, suggesting a central role for glutaminolysis in the preservation of reducing equivalent homeostasis in hemorrhagic shock erythrocytes. Inhibition of glutaminolysis in hemorrhagic shock rats increased aspartate consumption (D).



Supplementary Figure 3 – Plasma levels of metabolites from the TCA cycle and glycolysis in sham rats or rats undergoing hemorrhagic shock in presence or absence of the glutaminase inhibitor CB-839.

- Annotation-1
- Sham +CB839 Baseline
- Sham +CB839 End
- Sham +CB839 Post-Shock
- Sham Baseline
- Sham End
- Sham Post-Shock
- Shock +CB839 Baseline
- Shock +CB839 End
- Shock +CB839 Post-Shock
- Shock Baseline
- Shock End
- Shock Post-Shock

Supplementary Figure 4 – Hierarchical clustering analysis of metabolic measurements in RBCs from sham or hemorrhagic shock rats, in presence and absence of the treatment with the glutaminase inhibitor CB-839.

