

# **Characterization of epigenetic and transcriptional landscape in heat stressed-rats** using ATAC-seq and RNA-seq

#### Introduction

- Understanding animal physiology and identifying reliable biomarkers may help to establish effective management strategies for prevention of heat stress (HS)
- Little is known about the molecular mechanism of mammal tolerance to high temperature
- In a previous study with Sprague-Dawley rats, we performed RNA-seq assays on liver of rats in control check (CK; 22 °C, n = 5) and heat stress (H120; 42 °C for 120 min, n=5) groups. A total of 3,909 differential expression genes (DEGs, Q <0.05) were observed

#### **Materials and Methods**

- Animals: Female Sprague Dawley rats weighing  $205 \pm 7.16$  g (8 weeks old)
- **Methods:** Three liver tissues were selected from the RNA-seq samples in the CK and H120 groups and had an Assay for Transpose Accessible Chromatin sequencing (ATAC-seq) performed
- **Bioinformatics analysis:**

Quality control of raw data (Trimm-galore, FastQC)

Align to the reference genome (Bowtie2, MarkDuplicates)

Peak calling & annotation (MACS2, ChIPseeker)

Irreproducible Discovery Rate analysis (IDR)

Differential peak (DP) (P < 0.05& fold change >1.5) analysis (DiffBind)

Transcript factors (TFs) motif analysis (MEME)

**DPEGs\*** DPGs\* DEGs

Functional annotation (GO, KEGG)

\* DPGs: Differential peaks related genes DPEGs: Differential peaks related genes that were differentially expressed in RNA-seq data

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- All ATAC-seq libraries generated an expected distribution of the insert fragment lengths, with the majority of fragments being small and progressively fewer fragments of larger size (**Fig 1A**)
- The accessibility of transcriptional start sites (TSS) was significantly enriched (**Fig 1B**)
- which mapped to 571 DEGs (**Fig 2A**)
- oxidative stress were proposed as candidate TFs (Fig 2C-D)
- Thirty six and 22 TF-motifs were predicted by the up- and down-regulated DPs in CK vs. H120 (Fig 2B) • Based on the literature and integration of ATAC-seq and RNA-seq analyses, *Cebpa*, *Foxo4* and *Sp3* involving in the process of



• The nuclear chromatin in the liver of heat stressed-rats was less open than that of control rats • It is suggested that TFs, including *Cebpa*, *Foxo4* and *Sp3*, and their corresponding DEGs may be involved in the physiological mechanism of HS regulation

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### **Results and Discussion**



• A total of 2,586 DPs were identified comparing of CK vs. H120, including 230 up regulated DPs and 2,356 down regulated DPs,

### Conclusions

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