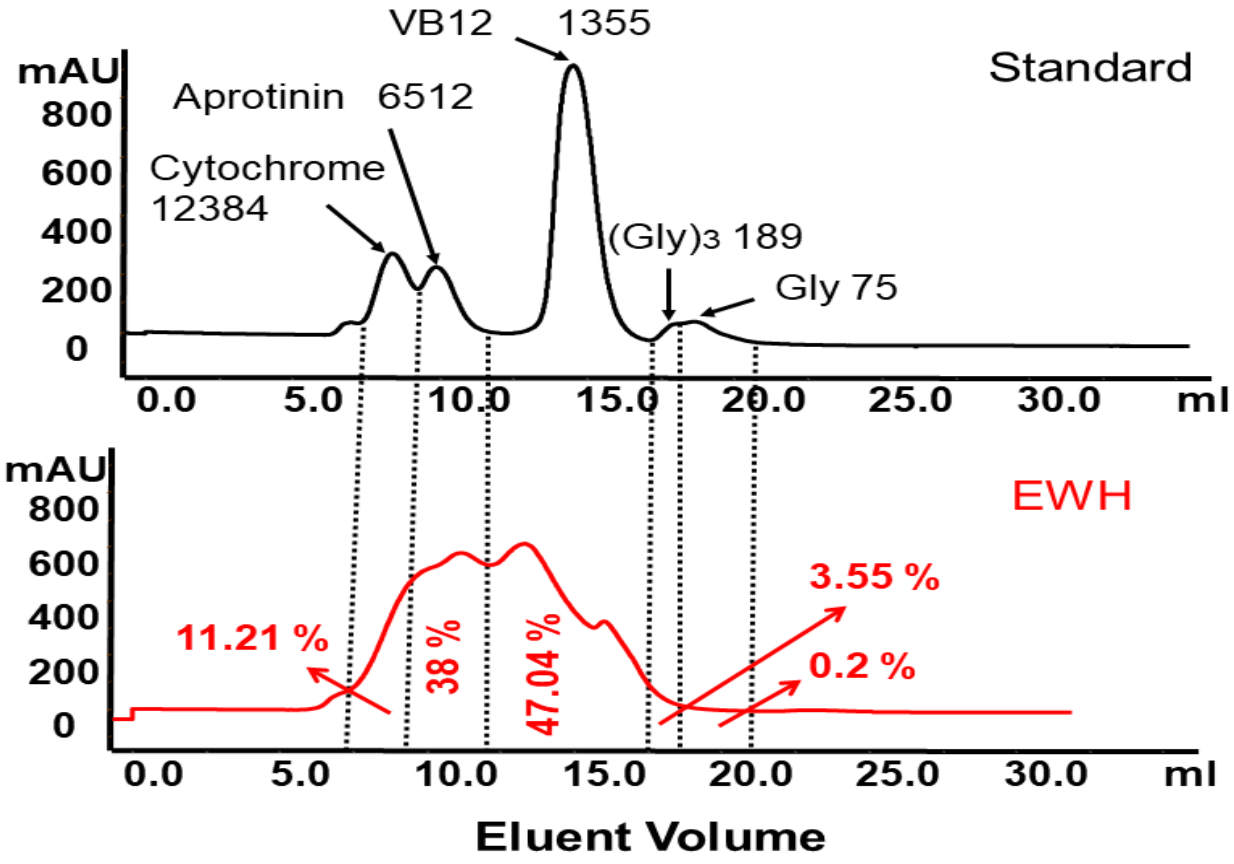
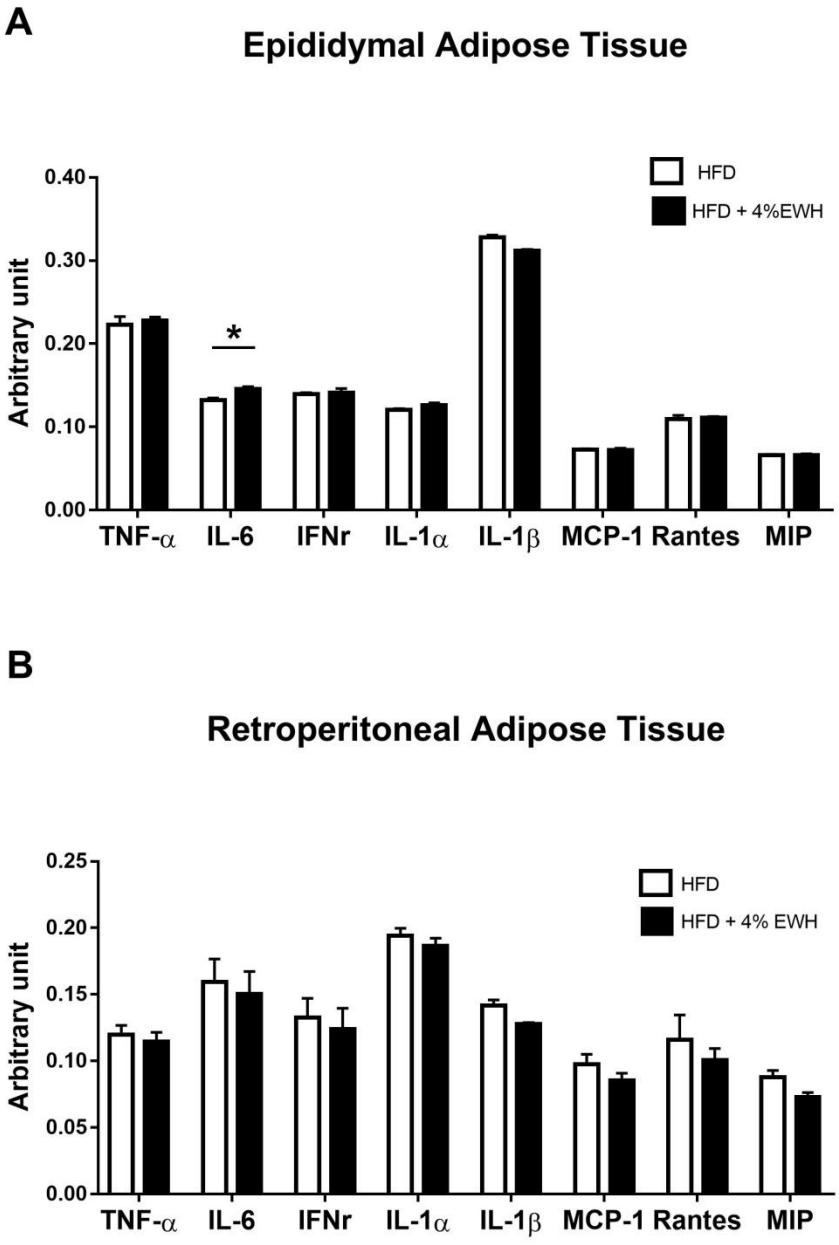
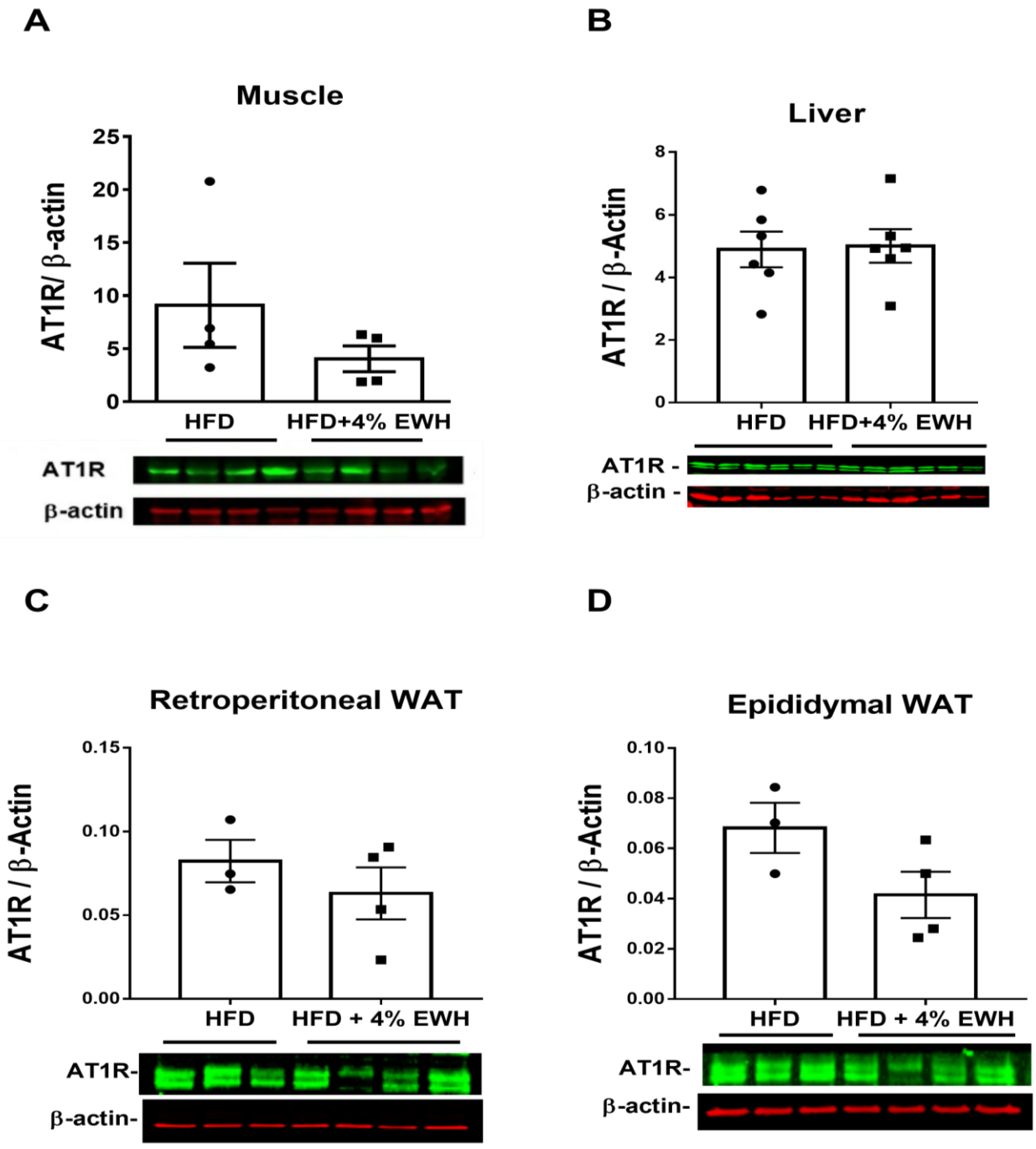
Supplementary material



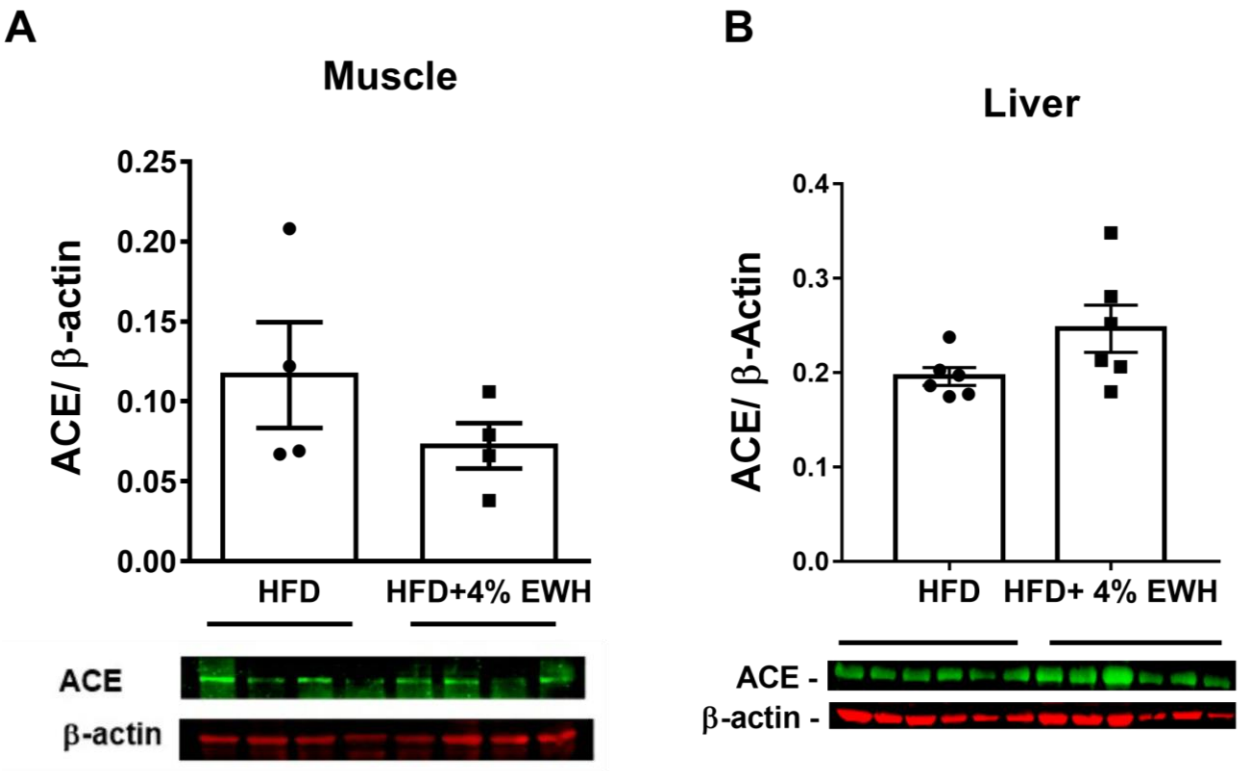
S 1- Size profile of EWH peptides. The majority of EWH-derived peptides (more than 85%) are in the range of 6.51 to 1.36 KDa. A small fraction of peptides (11.21%) with higher molecular weight between 12.38 to 6.51 KDa also present in the EWH. The molecular weight distribution of EWH peptides was determined using size-exclusion chromatography on an AKTA liquid chromatography system (GE Healthcare, Uppsala, Sweden) coupled with a Superdex Peptide 10/300GL column at room temperature. EWH was dissolved in 30% aqueous acetonitrile (ACN) containing 0.1% trifluoroacetic acid (TFA) and filtered through 0.22 µm filters. 100 µL of the sample was injected into the column and separated using an isocratic elution at a flow rate of 0.6 mL/min with 30% ACN containing 0.1% TFA. The absorbance of the eluent was monitored at 215 nm. Molecular weight markers (cytochrome c, 12384 Da; aprotinin, 6512 Da; vitamin B12, 1355 Da; (glycine)3, 189 Da; and glycine, 75 Da) were run under identical conditions to obtain the standard curve.



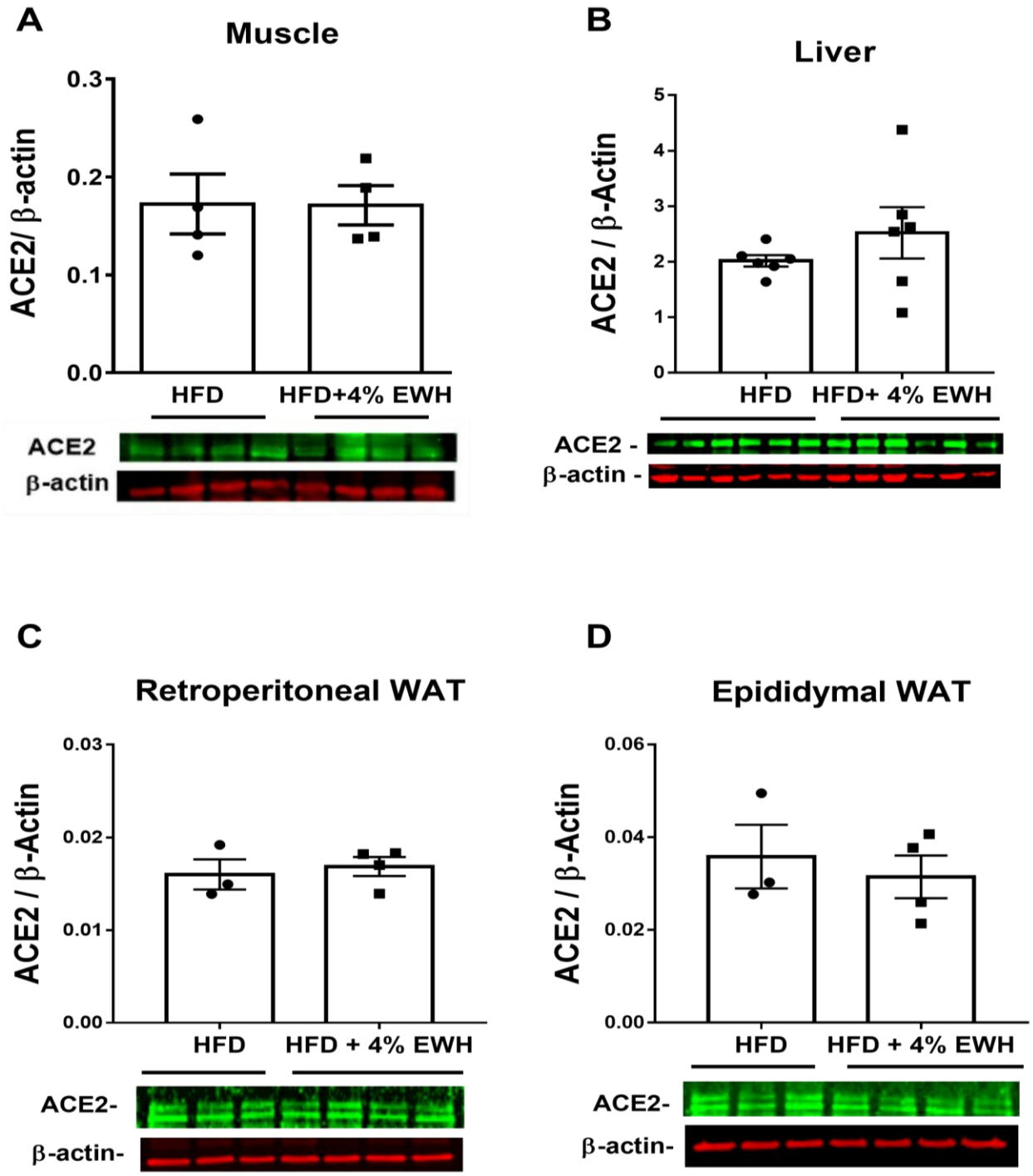
S 2- Epididymal and retroperitoneal adipose tissue inflammatory markers in (A) Epididymal adipose tissue n=3 rats and (B) Retroperitoneal adipose n=6 rats. Data are shown as the Mean ± SEM and were analyzed by two-tailed t-test. * shows significant difference at $p < 0.05$. TNF-α, Tumor necrosis factor; IL-6, Interleukin-6; IFNr, Interferon production regulator; IL-1α, Interleukin-1 alpha; IL-1β, Interleukin- 1 beta; MCP-1, Monocyte chemoattractant protein-1; Rantes, regulated on activation, normal T cell expressed and secreted; MIP, Macrophage inflammatory protein.



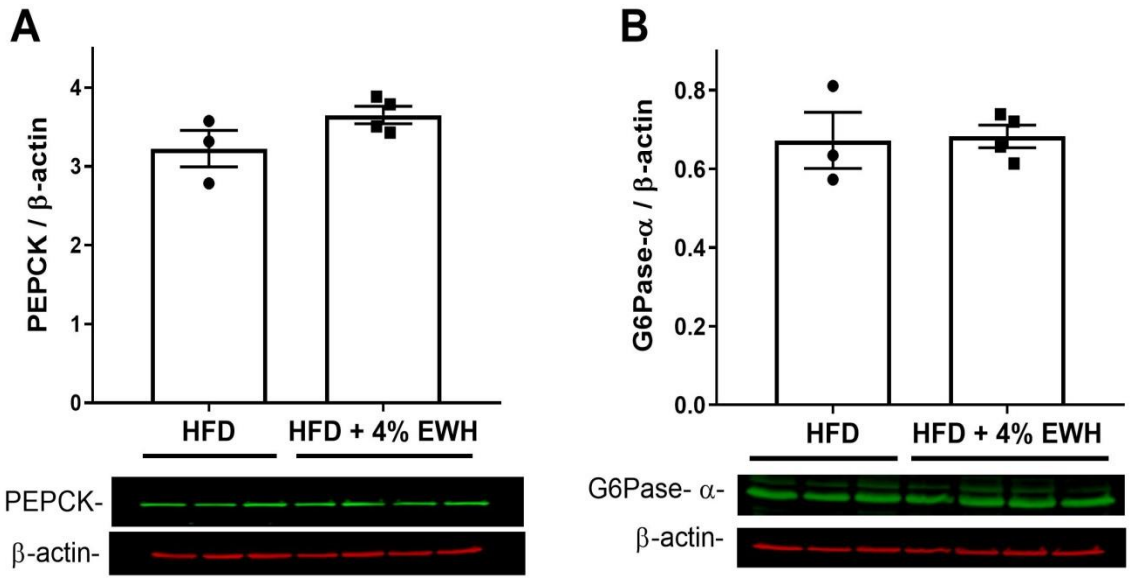
S 3- AT1R protein abundance in skeletal muscle (A), liver (B), and adipose tissue (C and D). The protein band of AT1R was normalized to β-actin as the loading control in HFD and HFD + 4% EWH treated groups. Data are shown as the Mean ± SEM for n= 4 rats and were analyzed by two-tailed t-test. AT1R, Angiotensin II type 1 receptor; WAT, white adipose tissue.



S 4- ACE protein abundance in skeletal muscle (A) and liver (B). The protein band of ACE was normalized to β-actin as the loading control in HFD and HFD+4%EWH treated groups. Data are shown as the Mean ± SEM for n= 4-6 rats and were analyzed by two-tailed t-test. ACE, angiotensin converting enzyme; WAT, white adipose tissue.



S 5- ACE2 protein abundance in skeletal muscle (A), liver (B), and adipose tissue (C and D). The protein band of ACE2 was normalized to β-actin as the loading control in HFD and HFD+4%EWH treated groups. Data are shown as the Mean ± SEM for n= 3-6 rats and were analyzed by two-tailed t-test. ACE2, angiotensin converting enzyme 2; WAT, white adipose tissue.



S 6- Liver PEPCK and G6Pase-α abundance. PEPCK and G6Pase protein bands were normalized to β-actin as the loading control. **(A)** PEPCK and **(B)** G6Pase in HFD and HFD+4%EWH treated groups. Data are shown as the Mean ± SEM for n= 3-4 rats and were analyzed by two-tailed t-test. PEPCK, Phosphoenolpyruvate carboxykinase; G6Pase-α, Glucose 6 phosphatase-α