UTSouthwestern

Medical Center

Department of Surgery

Mitochondrial fission with Function Impairment in Burn Serum Treated C2C12 Cells

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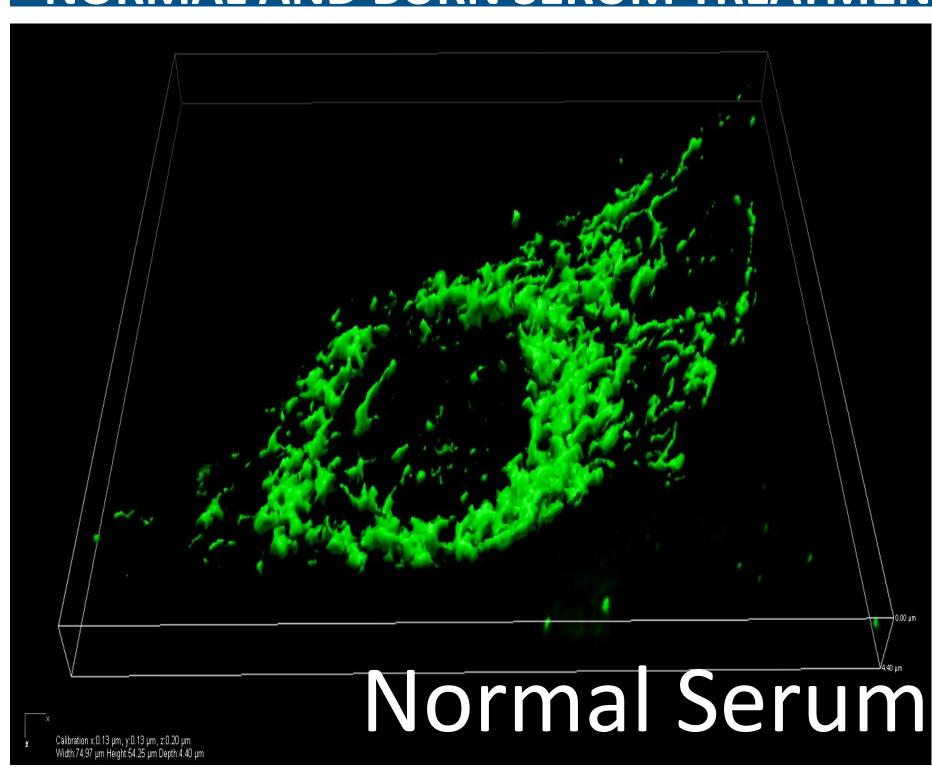
BACKGROUND

- Normal mitochondria have a fission and fusion dynamic cycle to maintain its function.
- Mitochondria fission is associated with cell death by releasing cytochrome C activated caspase cascade.
- Burn patients suffer muscle mass loss associated with increased muscle cell death.
- Mitochondria function impairment associated with muscle mass loss has been observed in severe burn patients.
- We hypothesize that severe burn impaired muscle atrophy is associated with increased mitochondria fission with function impairment.

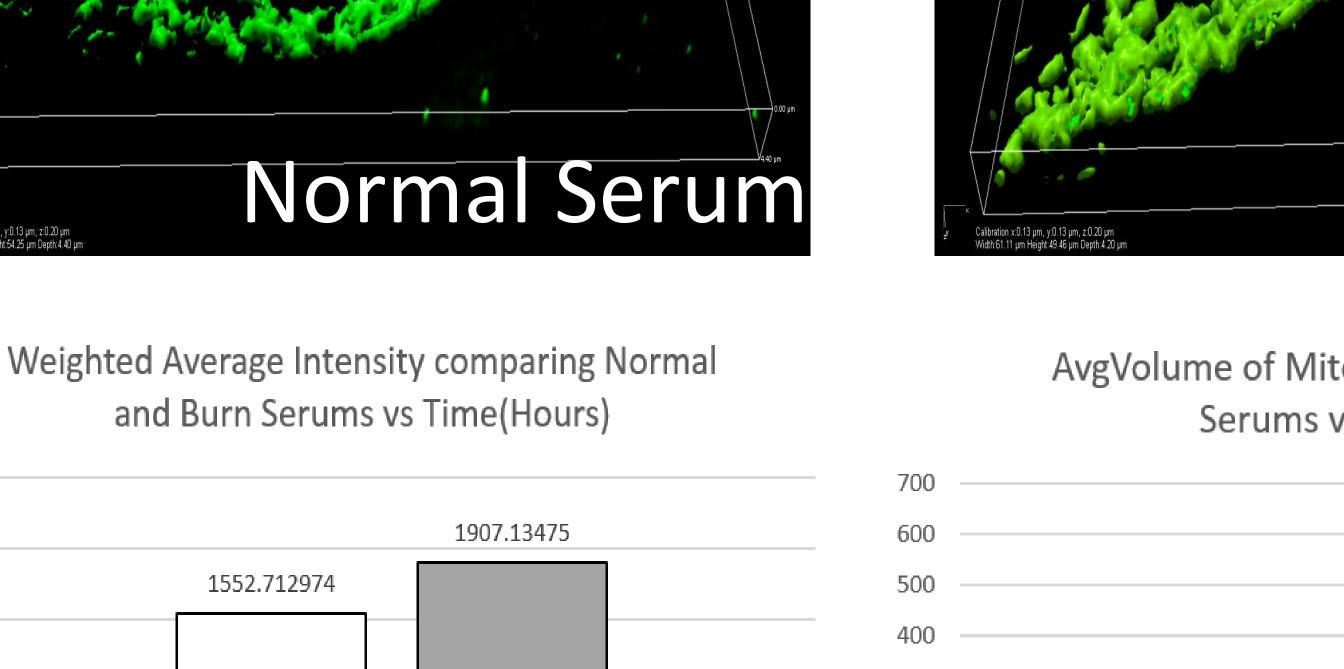
METHODS

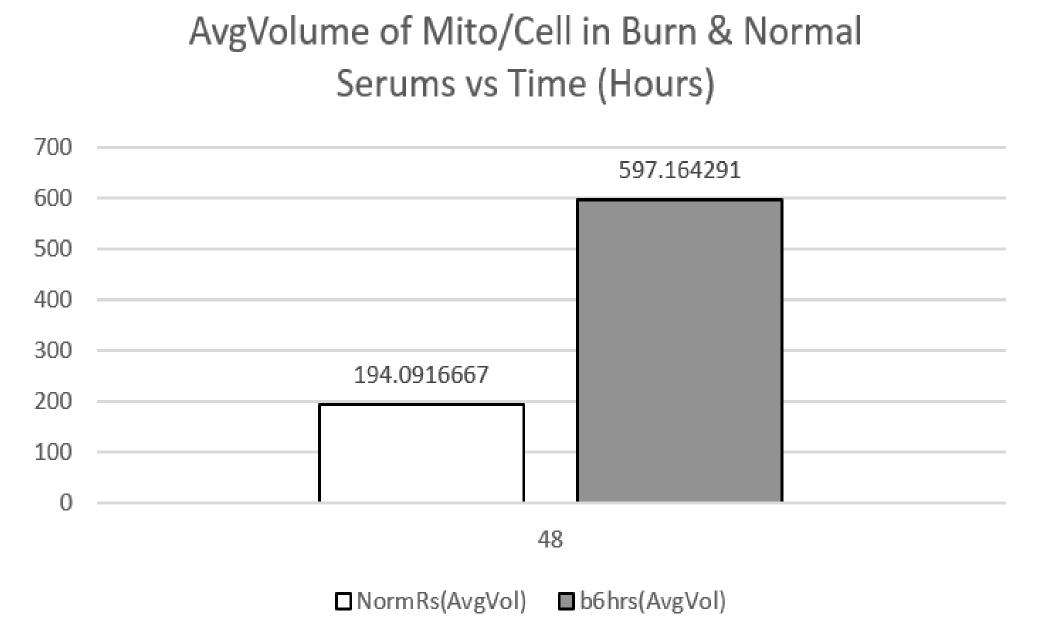
- Murine myoblast C2C12 cells were treated with DMEM media containing 10% rat serum isolated either from 40% TBSA scald burn rats, or control rats for 3 days.
- The cells were stained with MitoTracker Green FM dye and live cell images were taken under a Nikon Ti Eclipse Confocal microscope. Mitochondrial function was evaluated with Enzo Mito-ID membrane potential cytotoxicity kit.
- Cell lysate was collected, and western blotting for fission/fusion proteins, and caspase 3 activity was measured.

• RESULTS #1: THE INTENSITY AND VOLUME OF STAINED CELL MITOCHONDRIA WITH NORMAL AND BURN SERUM TREATMENT



■ Normrs_Intensity
■ B6hrs_Intensity





Burn Serum

CONCLUSION

- Mitochondria fission/fusion ratio increased in C2C12 cells stimulated with burn serum isolated 6 hours after burn.
- Mfn1, a fusion protein, decreased and confirmed the observation of mitochondrial fission in response to burn serum.
- Mitochondrial function impaired with dropped MMP level in muscle cells in response to burn serum stimulation.
- Muscle cell death increased with elevated caspase 3 activity in burn serum stimulated C2C12 cells.
- Continued work is needed to specify the effect of the burn serum and uncover target signals to the decreased mitochondrial function.

REFERENCE

- Gao AW, Canto C, Houtkooper RH. Mitochondrial response to nutrient availability and its role in metabolic disease. EMBO Mol Med. 2014;6(5):580–9.
- Porter C, Herndon DN, Sidossis LS, Børsheim E. (2013) The impact of severe burns on skeletal muscle mitochondrial function. Burns, 39: 1039–47.

ACKNOWLEDGEMENT

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• RESULTS #4: CASPASE 3 ACTIVITY IN C2C12 CELLS

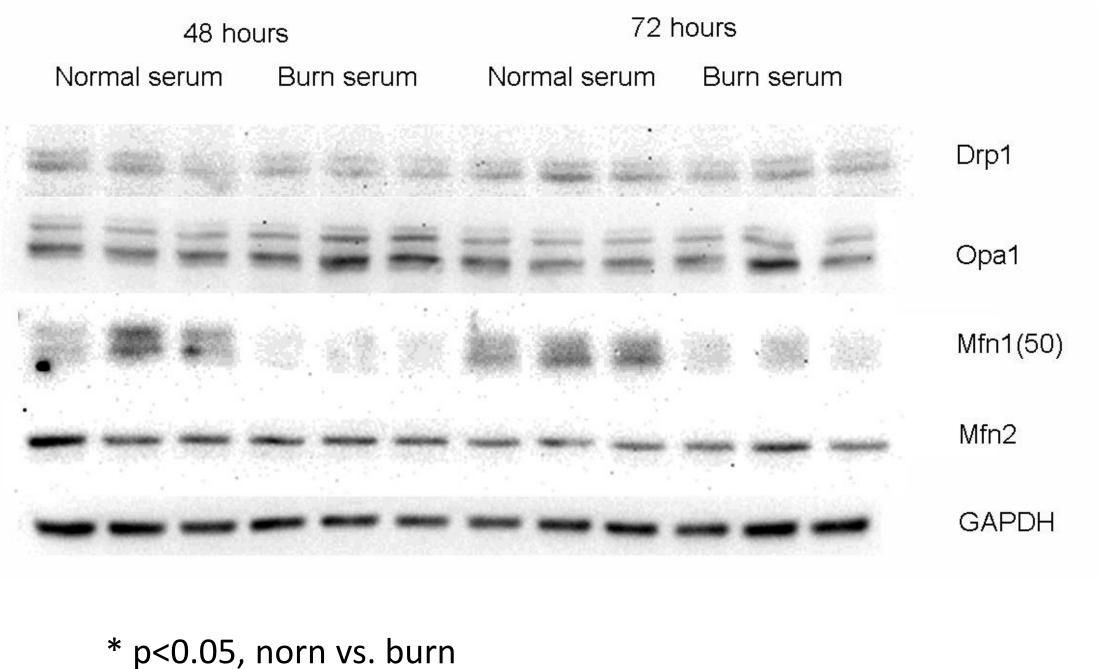
• RESULTS #2: WESTERN BLOT DETECTION OF MITOCHONDRIAL FISSION/FUSION PROTEINS EXPRESSION IN C2C12 CELLS

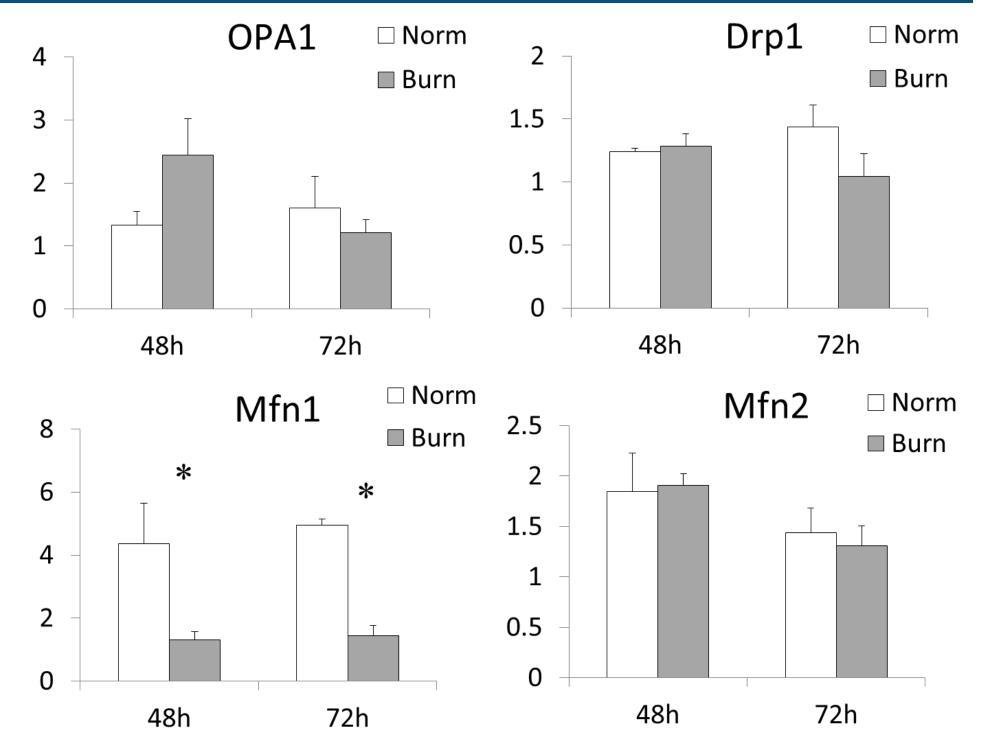
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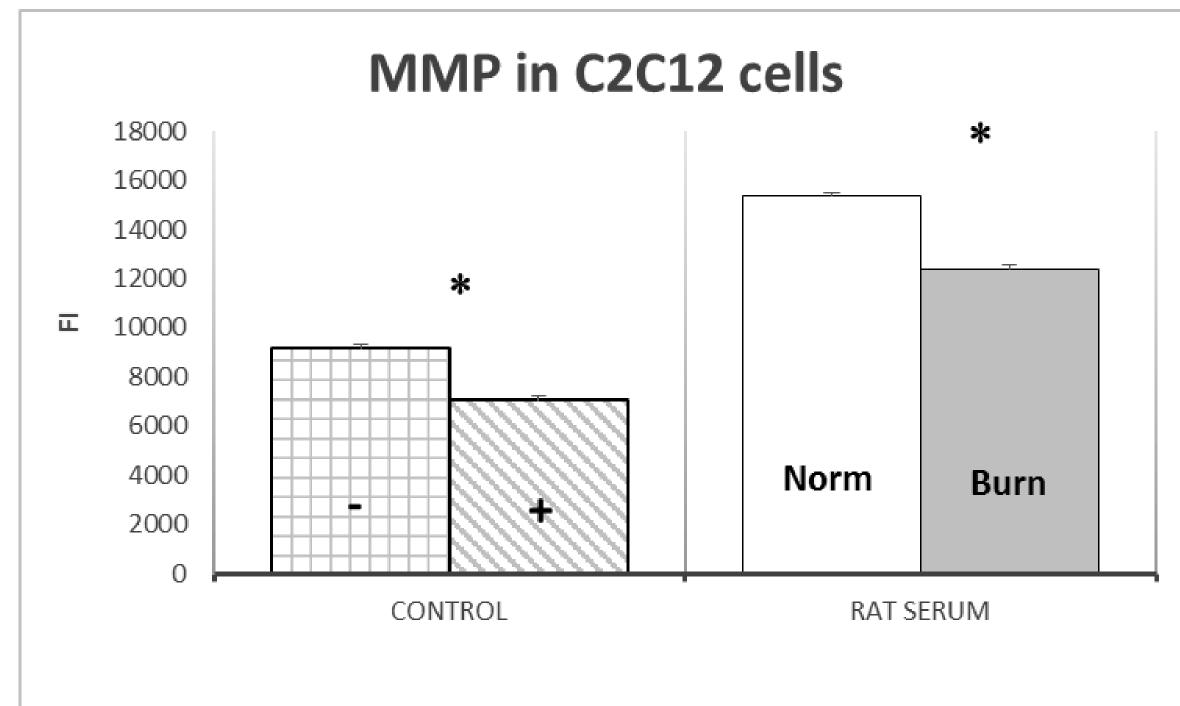
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• RESULTS #3: MITOCHONDRIAL MEMBRANE POTENTIAL IN RESPONSE TO RAT SERUM



• RESULTS #4: CASPASE 3 ACTIVITY IN C2C12 CELLS WITH RAT SERUM STIMULATION FOR 48 HOURS

