

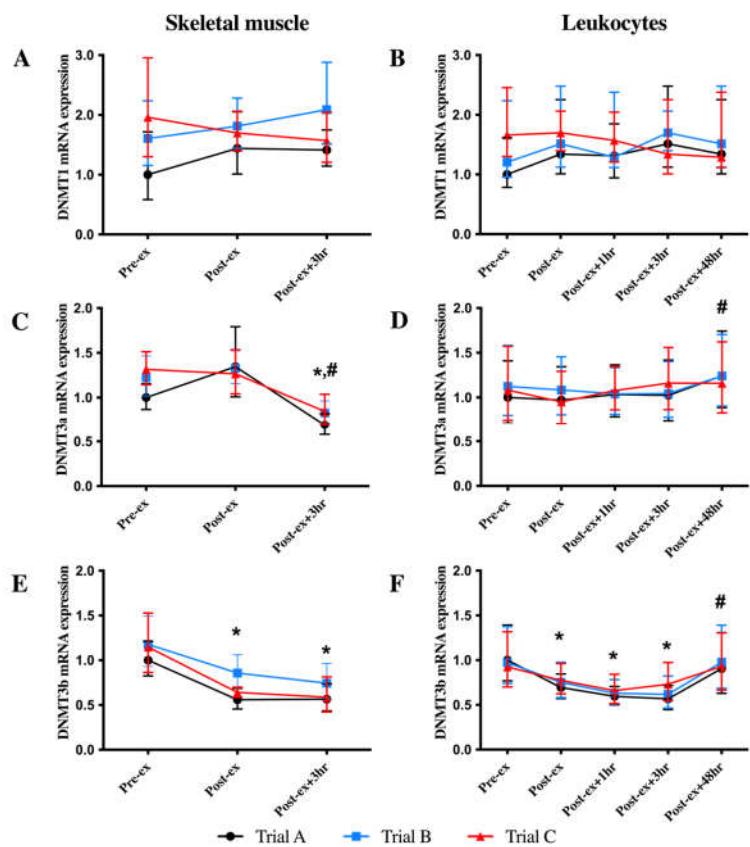
**Supplementary Table S1.** Serum concentrations of protein markers associated with inflammation and muscle damage. Data presented as the mean of each trial ± standard deviations. LDH; Lactate dehydrogenase, Mb; Myoglobin, CK; Creatine Kinase.

| Marker                   | Trial | Pre-ex         | Post-ex         | Post-ex+3hr     | Post-ex+48hr    |
|--------------------------|-------|----------------|-----------------|-----------------|-----------------|
| TNF- $\alpha$<br>(pg/mL) | A     | 0.26 ± 0.25    | 0.23 ± 0.19     | 0.29 ± 0.27     | 0.25 ± 0.22     |
|                          | B     | 0.26 ± 0.29    | 0.16 ± 0.13     | 0.21 ± 0.06     | 0.21 ± 0.23     |
|                          | C     | 0.12 ± 0.06    | 0.2 ± 0.11      | 0.27 ± 0.32     | 0.30 ± 0.27     |
| IL-6<br>(pg/mL)          | A     | 0.36 ± 0.27    | 3.73 ± 1.8      | 3.4 ± 2.73      | 2.16 ± 2.86     |
|                          | B     | 0.5 ± 0.39     | 2.66 ± 1.88     | 2.68 ± 1.15     | 0.66 ± 0.83     |
|                          | C     | 0.52 ± 0.19    | 4.91 ± 4.64     | 2.64 ± 1.3      | 0.7 ± 0.68      |
| LDH<br>(U/L)             | A     | 214.74 ± 71.44 | 238.16 ± 66.88  | 293.28 ± 84.59  | 291.44 ± 114.51 |
|                          | B     | 226.48 ± 92.25 | 240.49 ± 85.9   | 259.74 ± 77.61  | 254.97 ± 84.96  |
|                          | C     | 226.64 ± 65.12 | 243.23 ± 63.16  | 265 ± 65.04     | 247.95 ± 67.79  |
| Mb<br>( $\mu$ g/L)       | A     | 42.49 ± 22.35  | 438.18 ± 314.56 | 549.96 ± 399.67 | 364.14 ± 611.31 |
|                          | B     | 35.31 ± 11.26  | 201.43 ± 121.66 | 210.97 ± 134.8  | 34.23 ± 11.81   |
|                          | C     | 58.35 ± 72.42  | 214.03 ± 126.92 | 225.1 ± 114.79  | 32.02 ± 9.15    |
| CK<br>(U/L)              | A     | 140.88 ± 26.47 | 280.69 ± 115.18 | 610.14 ± 424.22 | 889.98 ± 714.5  |
|                          | B     | 163.38 ± 65.72 | 264.35 ± 111.26 | 397.45 ± 200.04 | 429.48 ± 220.21 |
|                          | C     | 144.5 ± 49     | 281.31 ± 118.28 | 431.61 ± 218.18 | 440.04 ± 234.83 |

**Supplementary Table S2.** Methylation of skeletal muscle and leukocytes at baseline (Trial A, Pre-ex). p<0.05 indicates a significant difference between tissues. Data presented as mean ± SD.

|            | CpG Site        | Tissue       | Methylation (%) | p |
|------------|-----------------|--------------|-----------------|---|
| <b>TNF</b> |                 |              |                 |   |
| CpG1       | Skeletal Muscle | 30.82 ± 5.11 | <0.001          |   |
|            | Leukocytes      | 13.17 ± 2.70 |                 |   |
| CpG2       | Skeletal Muscle | 25.03 ± 3.88 | <0.001          |   |
|            | Leukocytes      | 10.53 ± 2.5  |                 |   |
| CpG3       | Skeletal Muscle | 30.41 ± 3.17 | <0.001          |   |
|            | Leukocytes      | 12.62 ± 3.42 |                 |   |
| CpG4       | Skeletal Muscle | 50.87 ± 5.64 | <0.001          |   |
|            | Leukocytes      | 14.65 ± 3.31 |                 |   |
| CpG Mean   | Skeletal Muscle | 34.28 ± 4.38 | <0.001          |   |
|            | Leukocytes      | 12.74 ± 2.58 |                 |   |
| <b>IL6</b> |                 |              |                 |   |
| CpG1       | Skeletal Muscle | 72.41 ± 6.93 | 0.002           |   |
|            | Leukocytes      | 91.10 ± 1.92 |                 |   |
| CpG2       | Skeletal Muscle | 76.87 ± 4.46 | <0.001          |   |

|          |                 |                  |                  |
|----------|-----------------|------------------|------------------|
|          | Leukocytes      | $91.12 \pm 1.19$ |                  |
| CpG3     | Skeletal Muscle | $82.94 \pm 3.97$ | <b>0.008</b>     |
|          | Leukocytes      | $91.17 \pm 3.09$ |                  |
| CpG4     | Skeletal Muscle | $66.08 \pm 5.03$ | <b>&lt;0.001</b> |
|          | Leukocytes      | $88.89 \pm 1.95$ |                  |
| CpG5     | Skeletal Muscle | $72.23 \pm 4.03$ | <b>0.007</b>     |
|          | Leukocytes      | $82.01 \pm 3.58$ |                  |
| CpG6     | Skeletal Muscle | $74.09 \pm 5.38$ | <b>&lt;0.001</b> |
|          | Leukocytes      | $88.99 \pm 2.3$  |                  |
| CpG Mean | Skeletal Muscle | $74.10 \pm 4.14$ | <b>&lt;0.001</b> |
|          | Leukocytes      | $88.87 \pm 1.42$ |                  |



**Supplementary Figure S1.** Effect of exercise on the mRNA expression of (A&B) DNMT1, (C&D) DNMT3a and (E&F) DNMT3b in skeletal muscle (left-hand column) and leukocytes (right-hand column). \*Indicates significantly different from Pre-ex; # indicates significantly different from Post-ex.