



THE EFFECTS OF CREATINE SUPPLEMENTATION ON MUSCLE SYNTHESIS AND FITNESS LEVELS IN DROSOPHILA MELANOGASTER USING A MODEL OF MUSCLE ATROPHY

By Kevin Williams

RELEVANCE

Supplement Industry - \$100+ Billion/ yr

Creatine - \$450+ Million/ yr

Goal - Improve Livelihood

Current Studies on Muscle Atrophy



WHAT WILL BE TESTED?



HYPOTHESIS:

Creatine supplementation can increase the fitness of *Drosophila melanogaster* using a muscle atrophy model as proved by an increase in muscle size and exercise stamina when compared to a non-supplemented fly.

WHO AM I?

DROSOPHILA MELANOGASTER

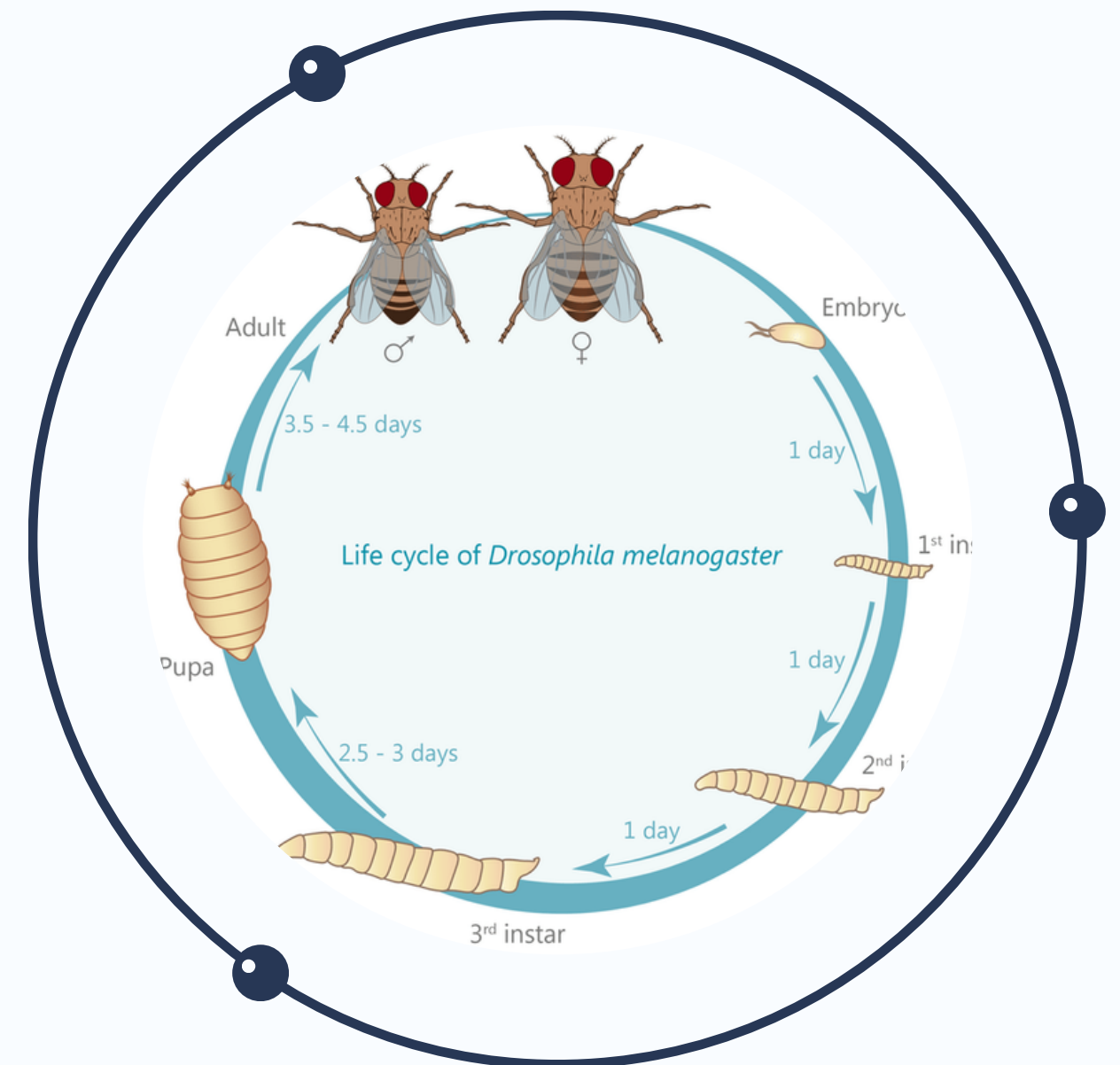
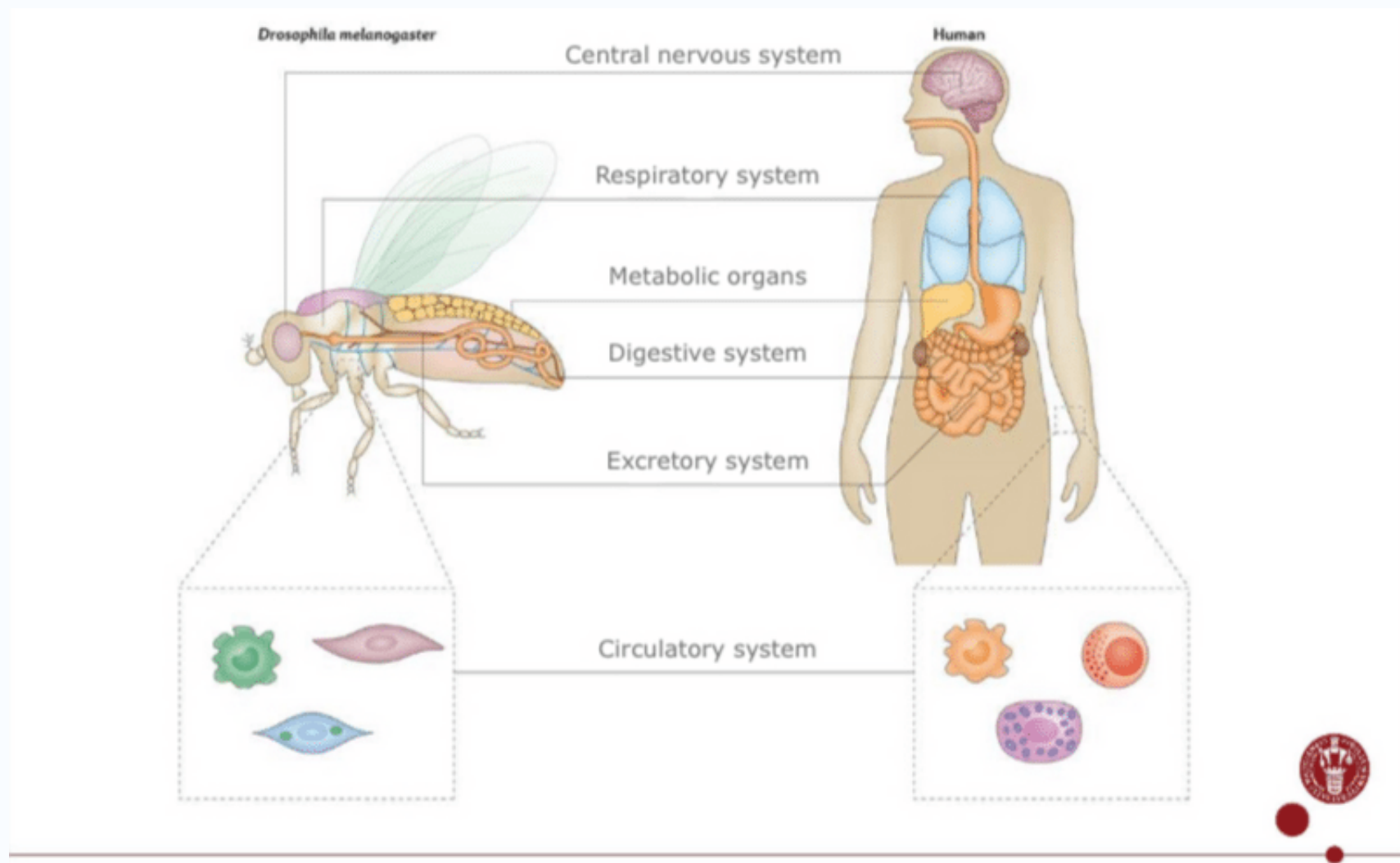


MODEL ORGANISM

Translate to humans

EASE OF CARE

Short life cycles



TESTING MECHANISM

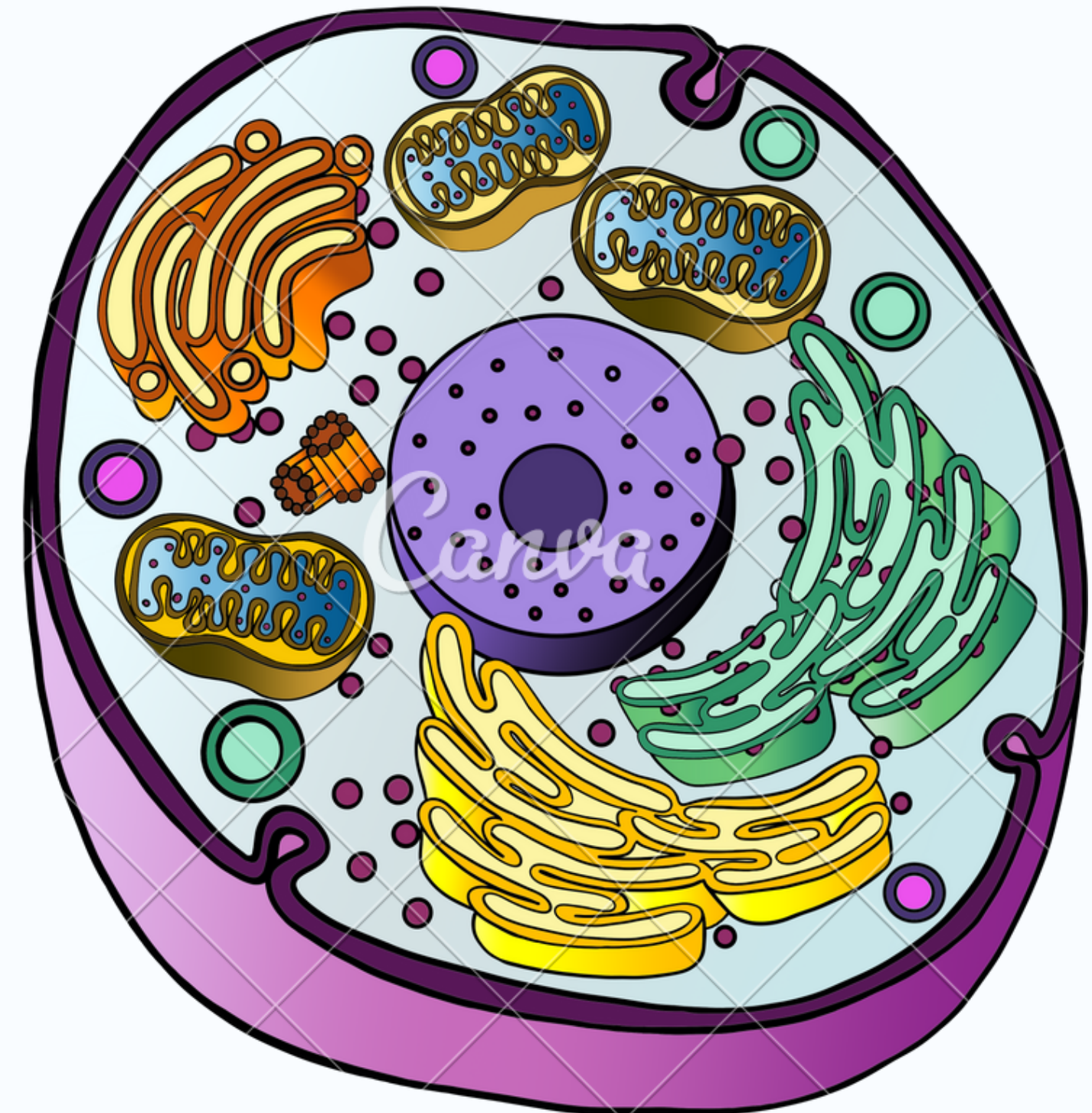


Cell

Many Organelles

Functioning body

Mitochondria



TESTING MECHANISM

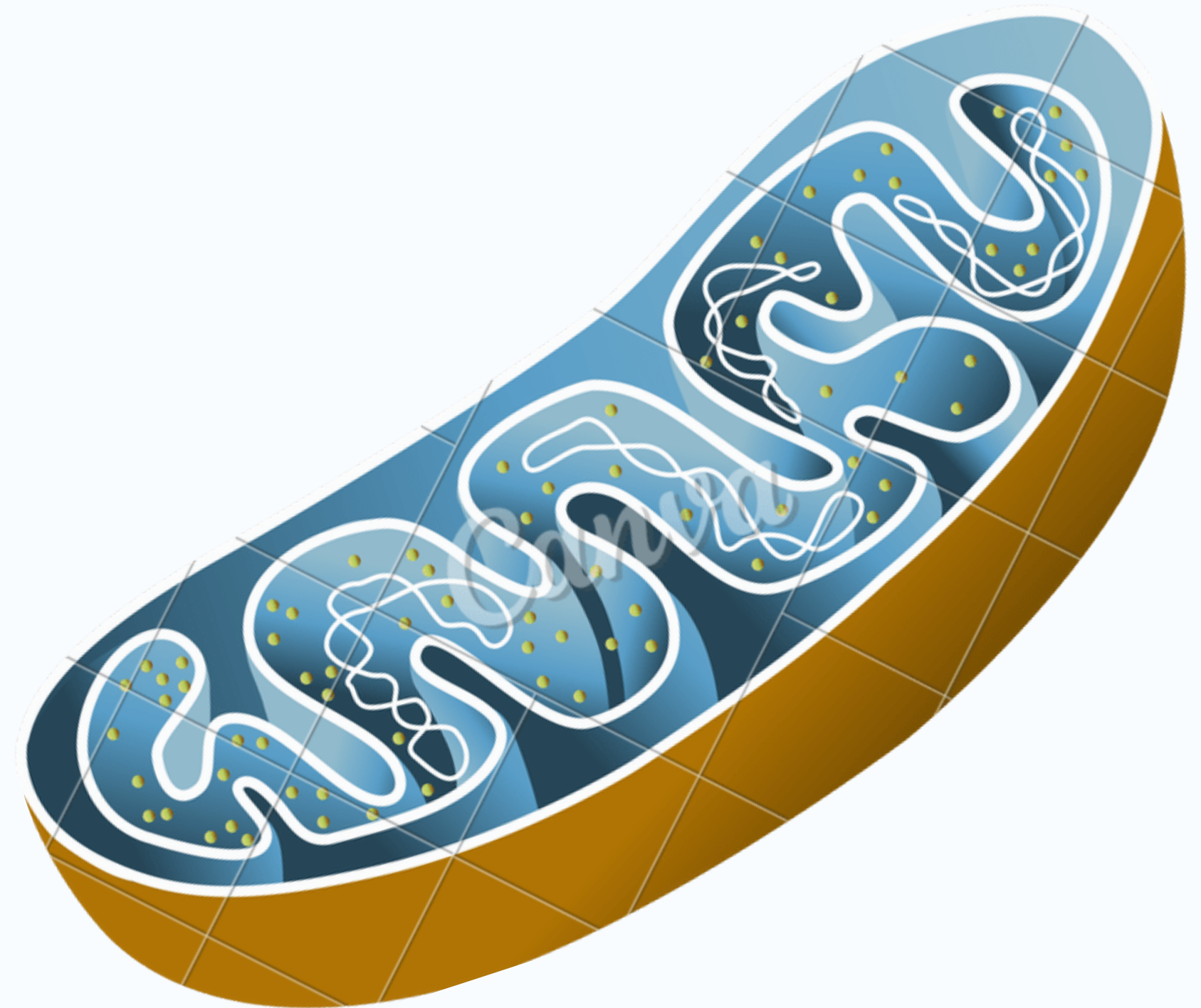


Mitochondria

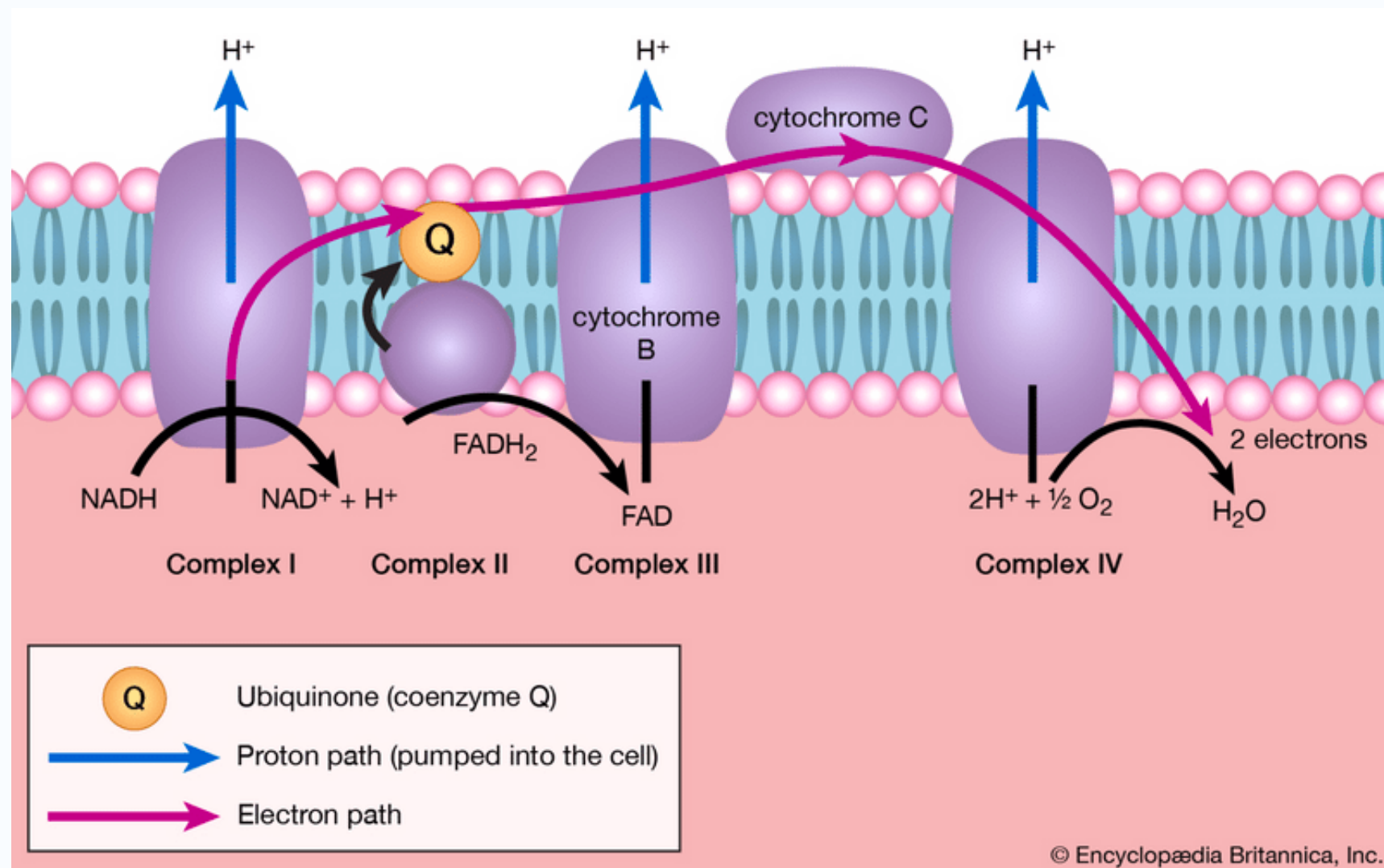
"Powerhouse" of the cell

Respiration and Energy production

Mechanisms to produce this energy



TESTING MECHANISM



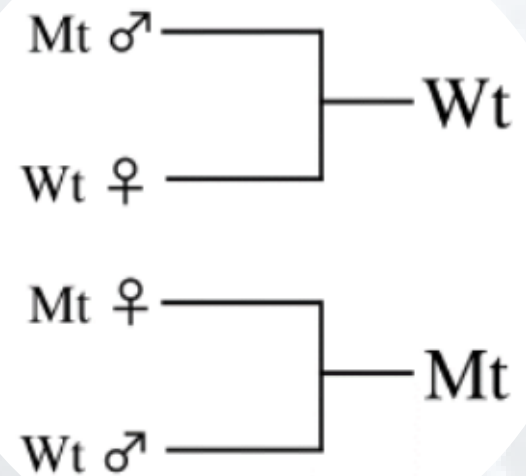
ELECTRON TRANSPORT CHAIN

An inhibited cytochrome oxidase

Phosphocreatine acts as a carrier for transferring energy from the mitochondria to the cytosol



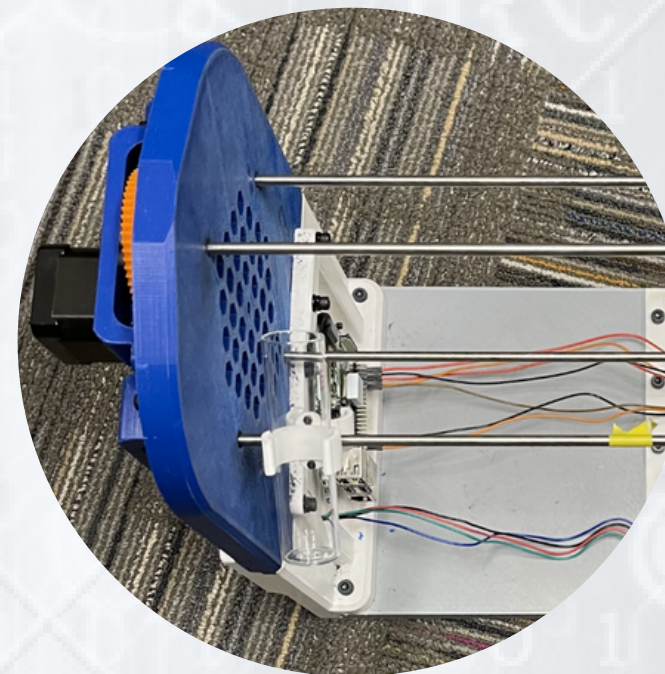
METHODS



Cross



Measure Before

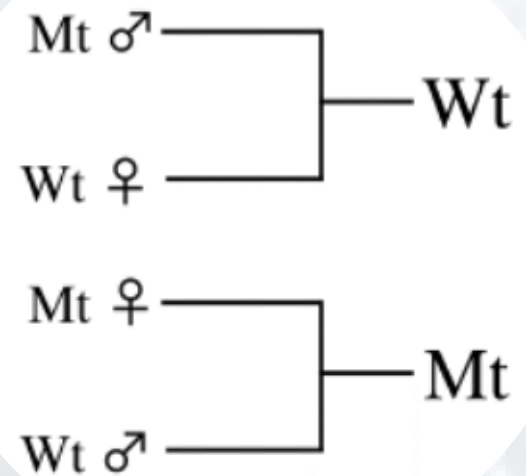


Supplement and Exercise

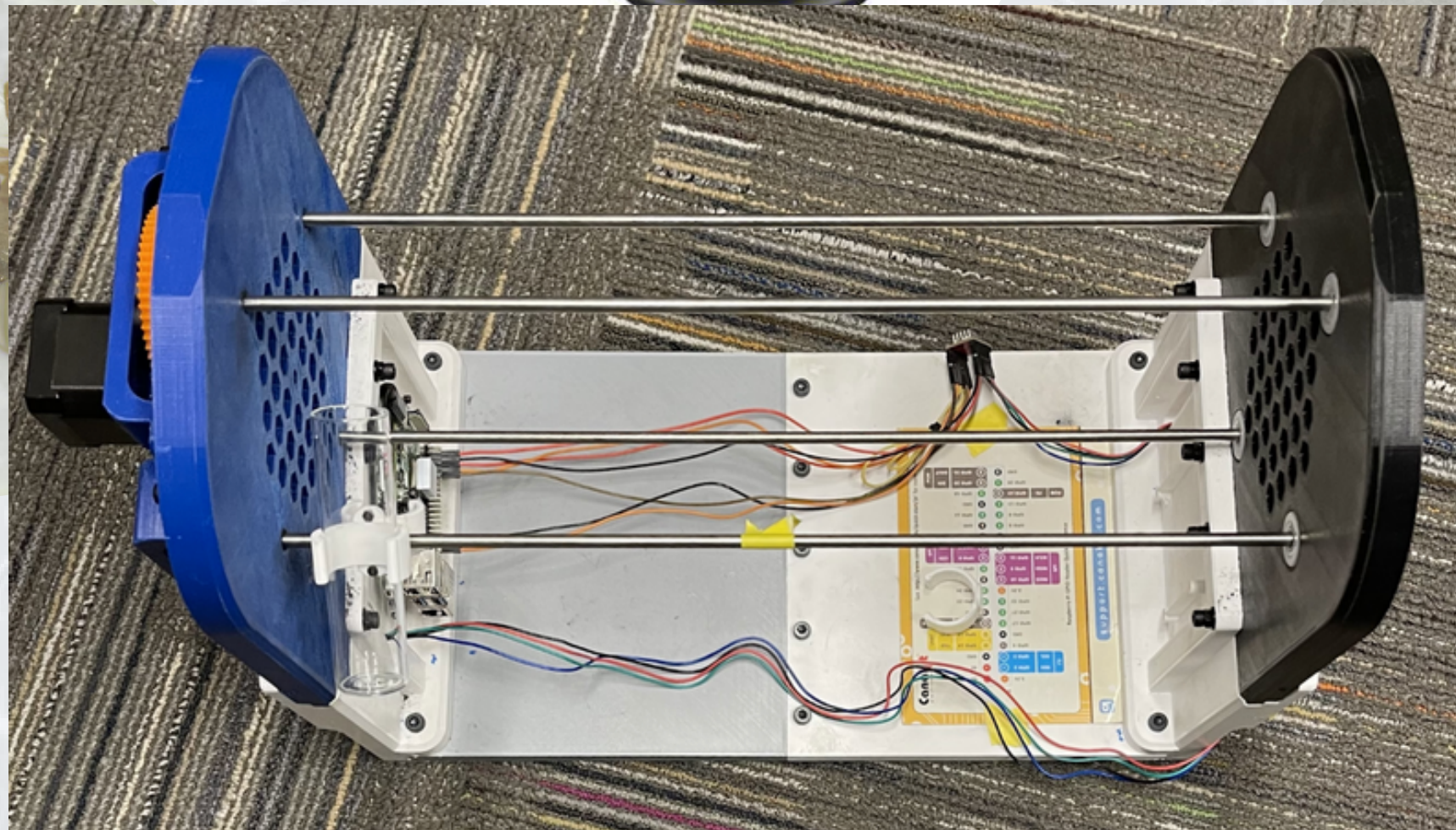


Measure After

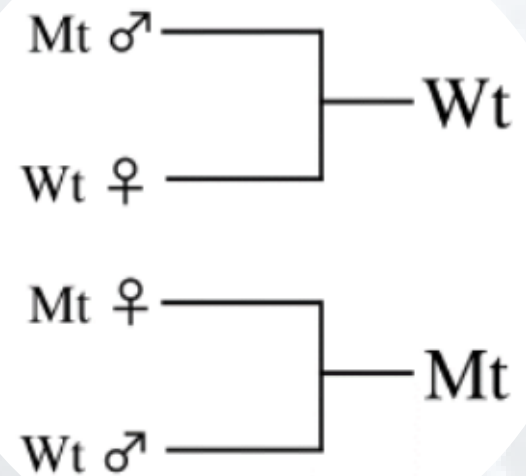
METHODS



Cross



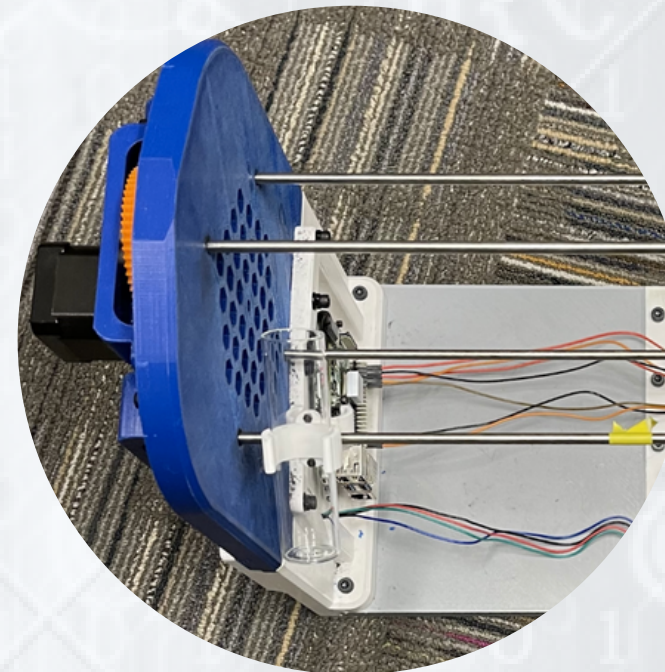
METHODS



Cross



Measure Before



Supplement and Exercise



Measure After

RESULTS

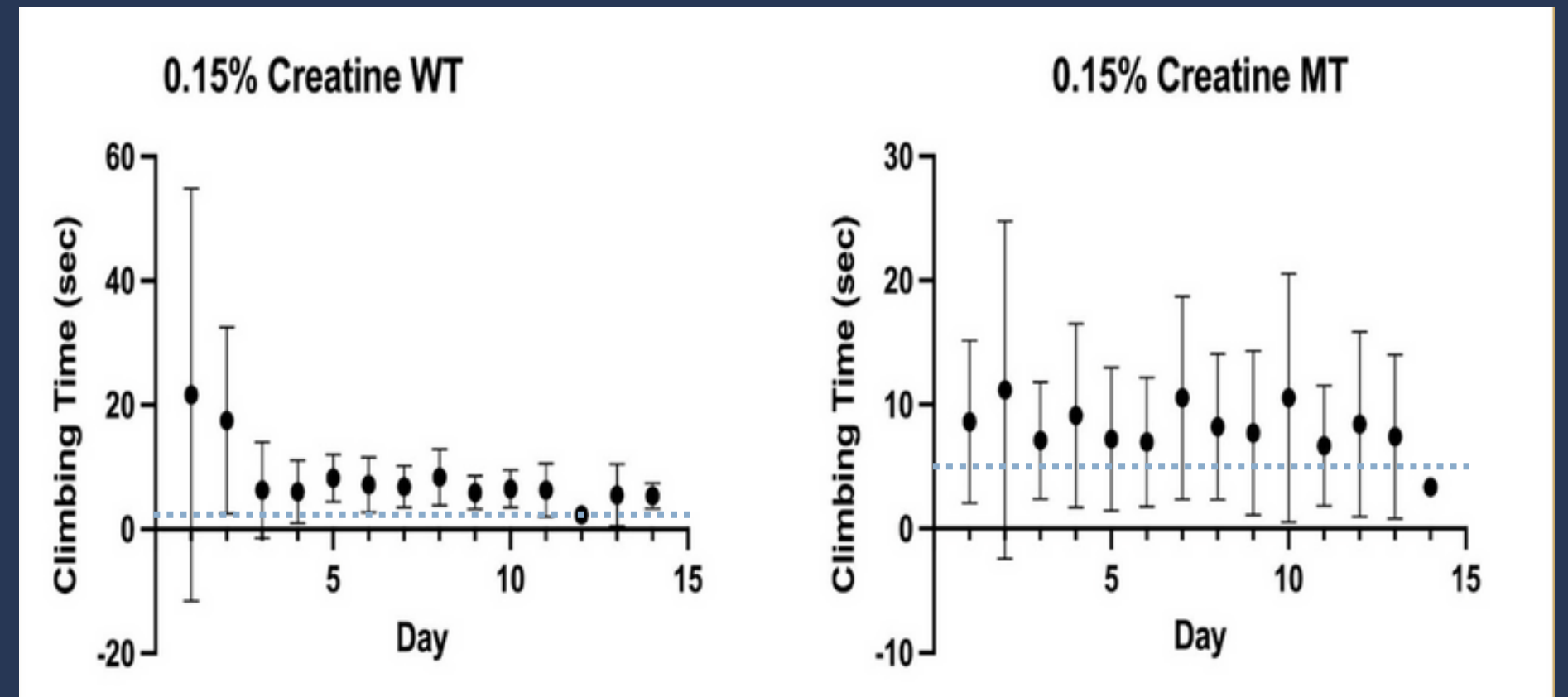
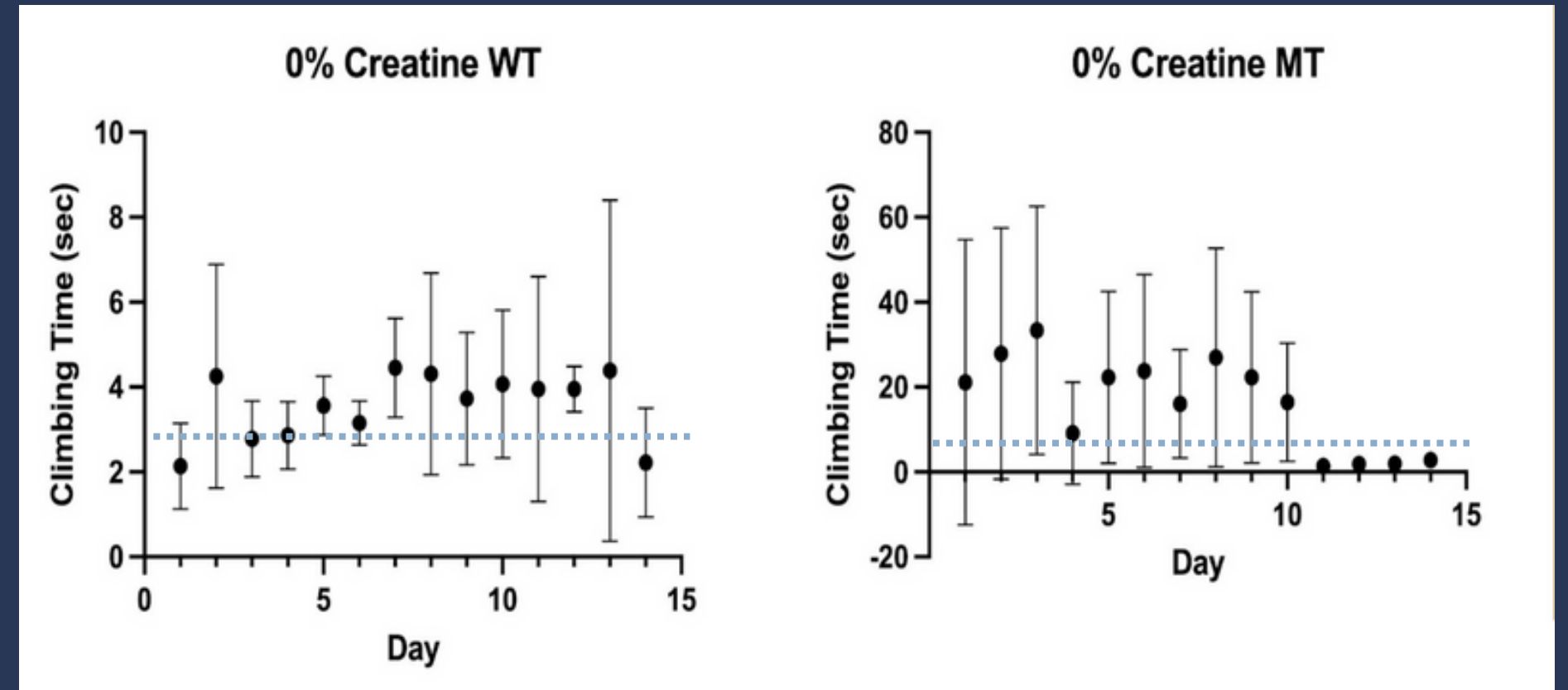
CLIMBING TIME

WILD TYPE (WT)

Increase with addition of creatine

MUTANT (MT)

Decrease with addition of creatine



RESULTS

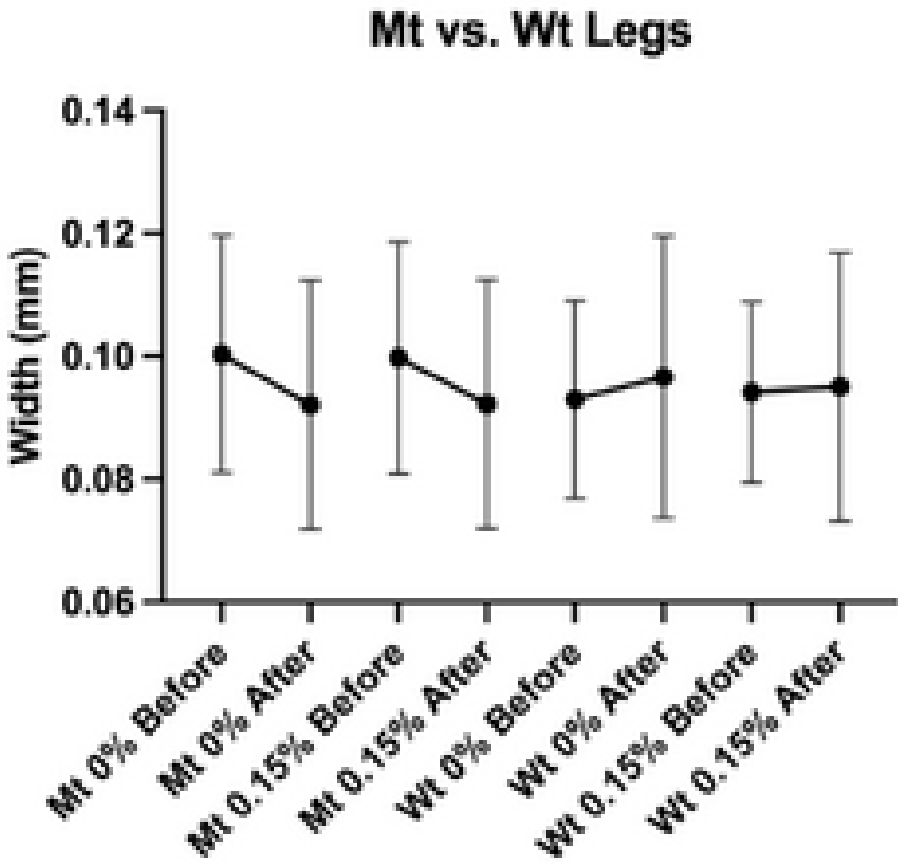
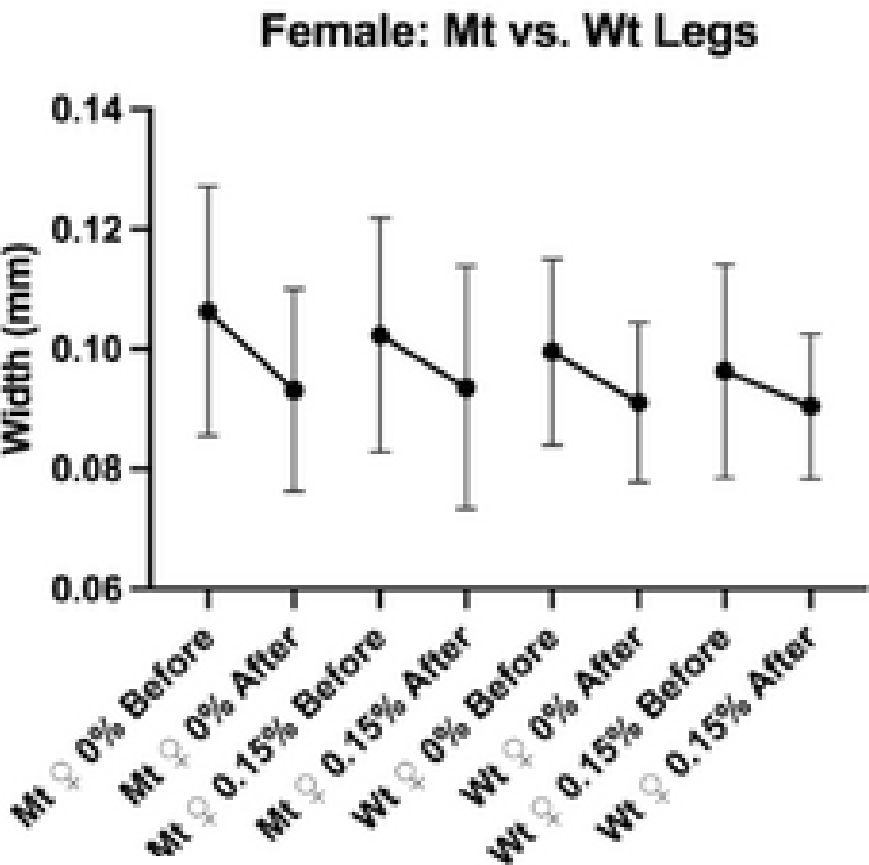
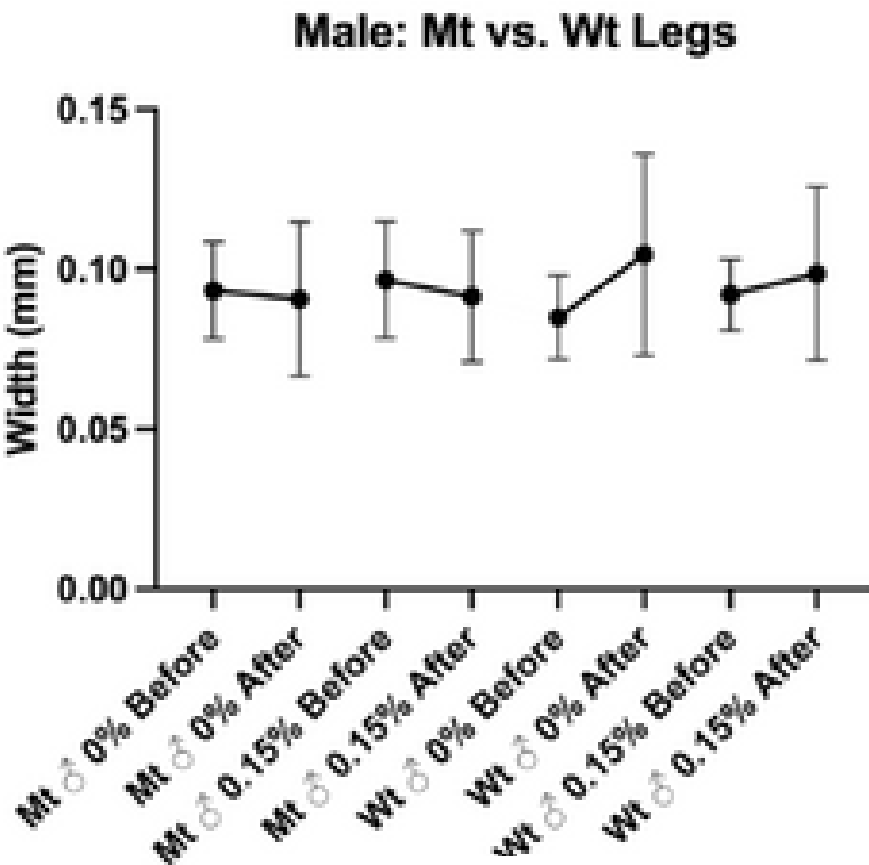
LEG WIDTH

WILD TYPE (WT)

Less of a decrease

MUTANT (MT)

More of a decrease



RESULTS

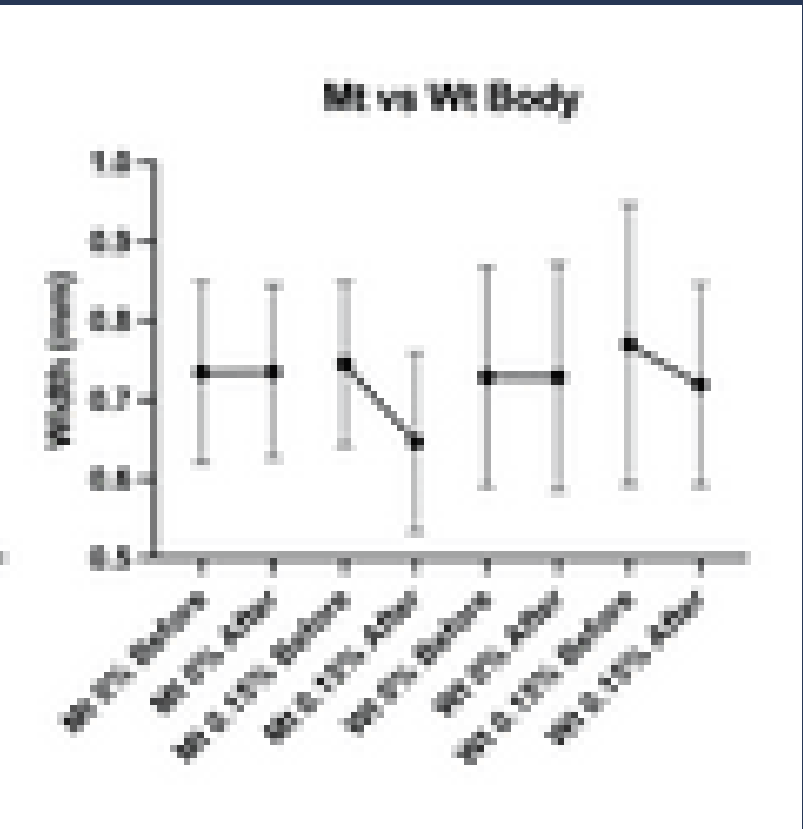
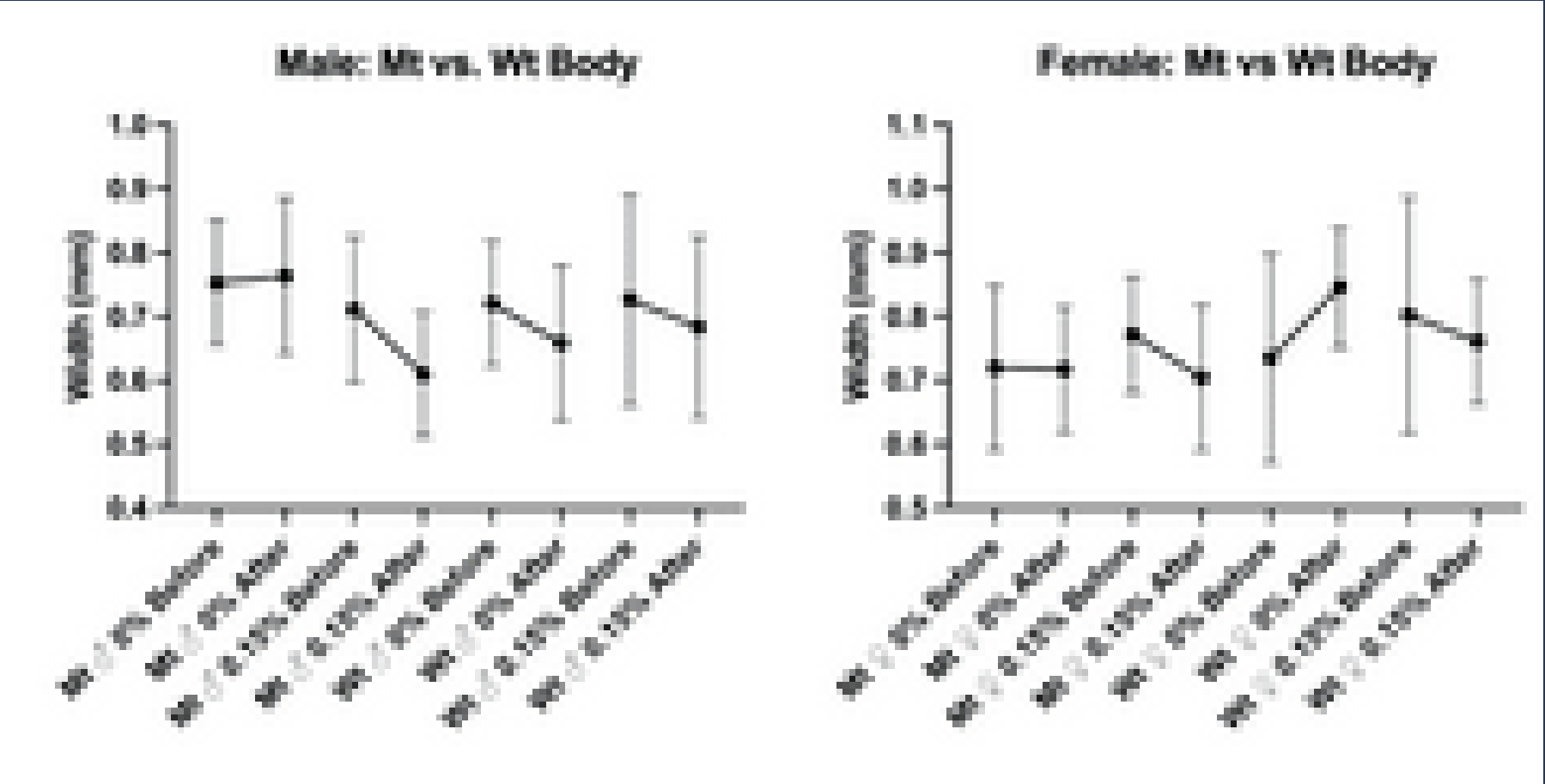
BODY WIDTH

WILD TYPE (WT)

Decrease slightly more with 0.15%

MUTANT (MT)

Decrease with 0.15%



INCONCLUSIVE

DATA IS LARGELY STATISTICALLY INSIGNIFICANT

FUTURE STUDIES



Additions/Changes

Better camera

More trials

More flies per trial

RNA Expression



REFERENCES

1. TOROK ZA, BUSEKRUS RB, HYDOCK D. EFFECTS OF TWO CREATINE MONOHYDRATE FEEDING PROTOCOLS ON DOXORUBICIN-INDUCED SKELETAL MUSCLE DYSFUNCTION. *FASEB J.* 2019;33(S1). DOI:10.1096/FASEBJ.2019.33.1_SUPPLEMENT.868.18
2. KREIPKE RE, KWON YV, SHCHERBATA HR, RUOHOLA-BAKER H. DROSOPHILA MELANOGASTER AS A MODEL OF MUSCLE DEGENERATION DISORDERS. IN: CURRENT TOPICS IN DEVELOPMENTAL BIOLOGY. VOL 121. ELSEVIER; 2017:83-109. DOI:10.1016/BS.CTDB.2016.07.003
3. KIM S, HONG KB, KIM S, SUH HJ, JO K. CREATINE AND TAURINE MIXTURES ALLEVIATE DEPRESSIVE-LIKE BEHAVIOUR IN DROSOPHILA MELANOGASTER AND MICE VIA REGULATING AKT AND ERK/BDNF PATHWAYS. *SCI REP.* 2020;10(1):11370. DOI:10.1038/S41598-020-68424-1
4. CREATINE. IN: PUBCHEM. NIH. [HTTPS://PUBCHEM.NCBI.NLM.NIH.GOV/COMPOUND/CREATINE](https://pubchem.ncbi.nlm.nih.gov/compound/Creatine).
5. WYSS M, KADDURAH-DAOUK R. CREATINE AND CREATININE METABOLISM. *PHYSIOL REV.* 2000;80(3):1107-1213. DOI:10.1152/PHYSREV.2000.80.3.1107
6. ANTONIO J, CANDOW DG, FORBES SC, ET AL. COMMON QUESTIONS AND MISCONCEPTIONS ABOUT CREATINE SUPPLEMENTATION: WHAT DOES THE SCIENTIFIC EVIDENCE REALLY SHOW? *J INT SOC SPORTS NUTR.* 2021;18(1):13. DOI:10.1186/S12970-021-00412-W
7. DALBO VJ, ROBERTS MD, STOUT JR, KERKSICK CM. PUTTING TO REST THE MYTH OF CREATINE SUPPLEMENTATION LEADING TO MUSCLE CRAMPS AND DEHYDRATION. *BR J SPORTS MED.* 2008;42(7):567-573. DOI:10.1136/BJSM.2007.042473
8. DAUTANT A, MEIER T, HAHN A, TRIBOUILLARD-TANVIER D, DI RAGO JP, KUCHARCZYK R. ATP SYNTHASE DISEASES OF MITOCHONDRIAL GENETIC ORIGIN. *FRONT PHYSIOL.* 2018;9:329. DOI:10.3389/FPHYS.2018.00329
9. XIONG Q, DU F, ZHU X, ET AL. ATP PRODUCTION RATE VIA CREATINE KINASE OR ATP SYNTHASE IN VIVO: A NOVEL SUPERFAST MAGNETIZATION SATURATION TRANSFER METHOD. *CIRC RES.* 2011;108(6):653-663. DOI:10.1161/CIRCRESAHA.110.231456
10. GARCÍA-AGUILAR A, CUEZVA JM. A REVIEW OF THE INHIBITION OF THE MITOCHONDRIAL ATP SYNTHASE BY IF1 IN VIVO: REPROGRAMMING ENERGY METABOLISM AND INDUCING MITOHORMESIS. *FRONT PHYSIOL.* 2018;9:1322. DOI:10.3389/FPHYS.2018.01322
11. KULEESHA Y, PUAH WC, WASSER M. A MODEL OF MUSCLE ATROPHY BASED ON LIVE MICROSCOPY OF MUSCLE REMODELLING IN DROSOPHILA METAMORPHOSIS. *R SOC OPEN SCI.* 2016;3(2):150517. DOI:10.1098/RSOS.150517
12. SMITH WW, THOMAS J, LIU J, LI T, MORAN TH. FROM FAT FRUIT FLY TO HUMAN OBESITY. *PHYSIOL BEHAV.* 2014;136:15-21. DOI:10.1016/J.PHYSBEH.2014.01.017
13. PAN Y, BAKER BS. GENETIC IDENTIFICATION AND SEPARATION OF INNATE AND EXPERIENCE-DEPENDENT COURTSHIP BEHAVIORS IN DROSOPHILA. *CELL.* 2014;156(1-2):236-248. DOI:10.1016/J.CELL.2013.11.041
14. LOVERO D, GIORDANO L, MARSANO RM, ET AL. CHARACTERIZATION OF DROSOPHILA ATPSYNC MUTANTS AS A NEW MODEL OF MITOCHONDRIAL ATP SYNTHASE DISORDERS. SKOULAKIS EMC, ED. *PLOS ONE.* 2018;13(8):E0201811. DOI:10.1371/JOURNAL.PONE.0201811
15. KAMIKOUCHI A, INAGAKI HK, EFFERTZ T, ET AL. THE NEURAL BASIS OF DROSOPHILA GRAVITY-SENSING AND HEARING. *NATURE.* 2009;458(7235):165-171. DOI:10.1038/NATURE07810
16. LAU MT, LIN YQ, KISLING S, ET AL. A SIMPLE HIGH THROUGHPUT ASSAY TO EVALUATE WATER CONSUMPTION IN THE FRUIT FLY. *SCI REP.* 2017;7(1):16786. DOI:10.1038/S41598-017-16849-6
17. LÜERSEN K, RÖDER T, RIMBACH G. DROSOPHILA MELANOGASTER IN NUTRITION RESEARCH—THE IMPORTANCE OF STANDARDIZING EXPERIMENTAL DIETS. *GENES NUTR.* 2019;14(1):3. DOI:10.1186/S12263-019-0627-9
- MENDEZ S, WATANABE L, HILL R, ET AL. THE TREADWHEEL: A NOVEL APPARATUS TO MEASURE GENETIC VARIATION IN RESPONSE TO GENTLY INDUCED EXERCISE FOR DROSOPHILA. IIJIMA KM, ED. *PLOS*