EVEN MODEST CHANGES IN DIET COMPOSITION IMPACT METABOLIC AND SKELETAL OUTCOMES

IN THE YELLOW KUO KONDO (KK-A^y) Murine model of type 2 diabetes (T2D)

ON White¹, RK Kohler¹, KR Gallagher¹, DM Segvich¹, K Rajesh¹, and JM Wallace¹

¹Weldon School of Biomedical Engineering, Purdue University, Indianapolis, IN

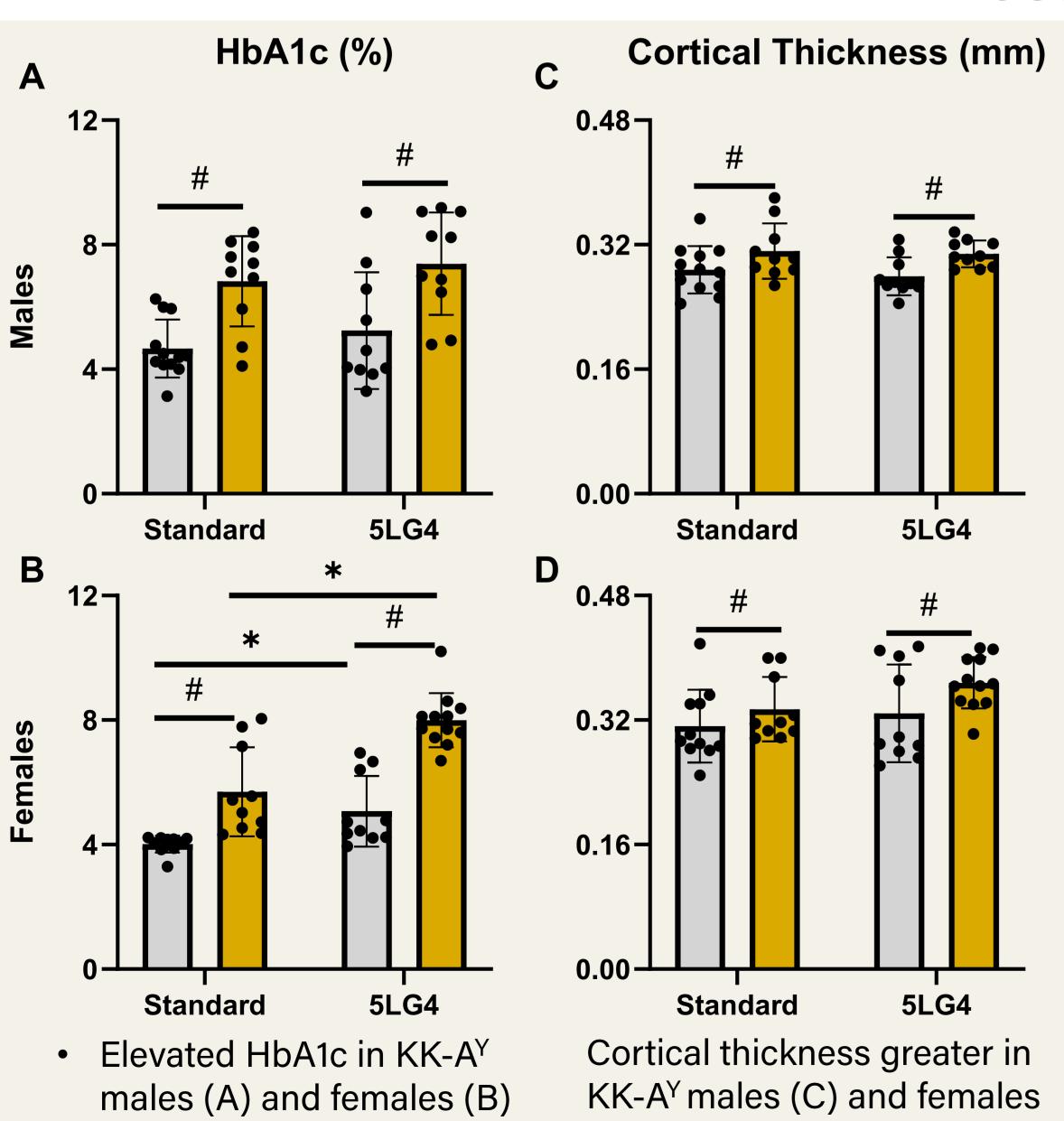
Bone Biology and Mechanics Lab Purdue University

Abstract

- 1 in 9 Diabetic Adults Worldwide: Type 2 diabetics (T2D) > 90% of cases [1]
- **T2D Characteristics:** Insulin resistance, hyperglycemia, & elevated fracture risk despite normal or elevated BMD [2], [3]
- KK-A^Y Model [4] used to assess source of fracture risk: Pilot studies suggest diet dependency of KK-A^Y Model

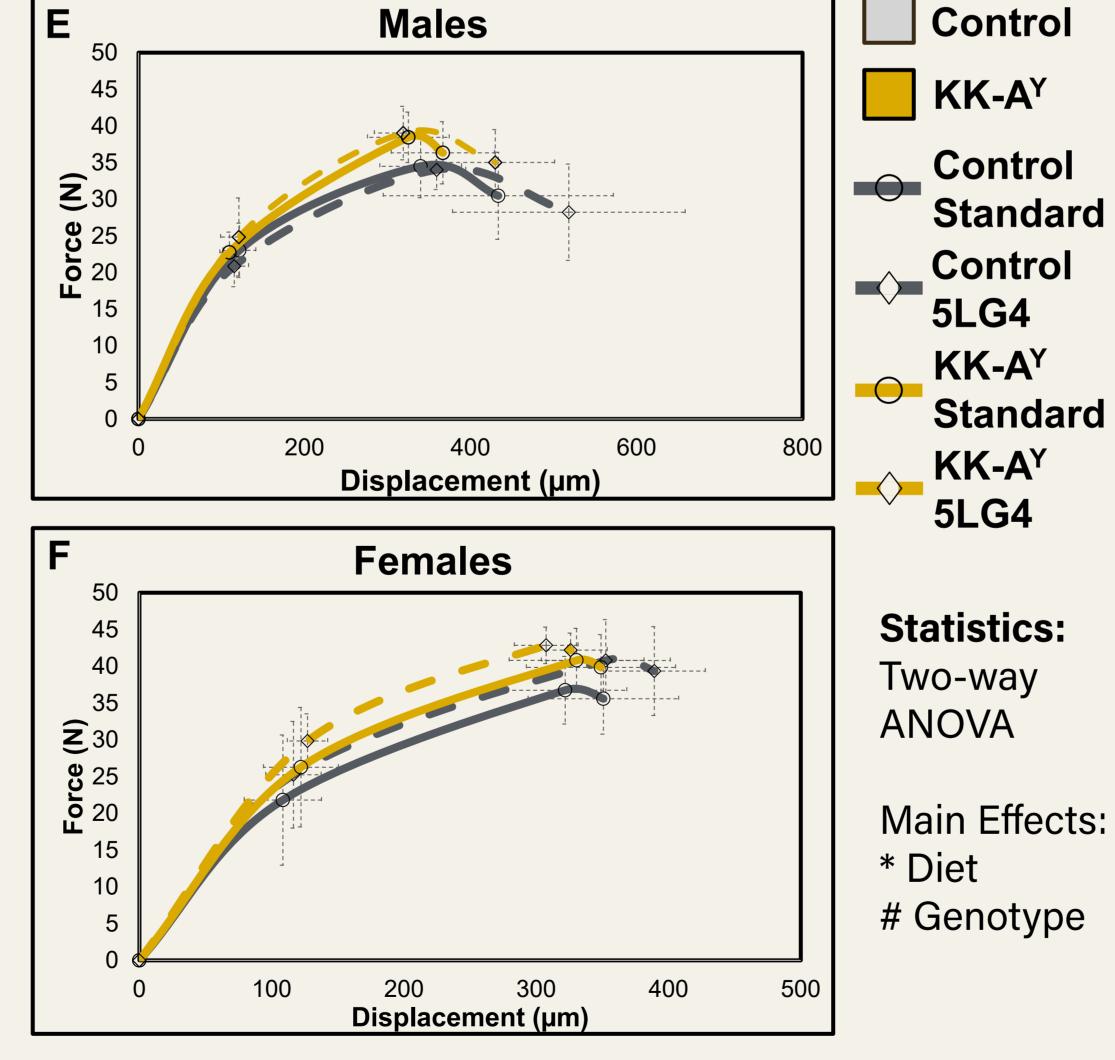
Hypothesis: T2D metabolic (hyperglycemia, obesity, elevated HbA1c) and skeletal characteristics (increased bone size & fracture risk) are dependent on diet in the KK-A^Y murine model of T2D

Results



5LG4 elevated HbA1c in

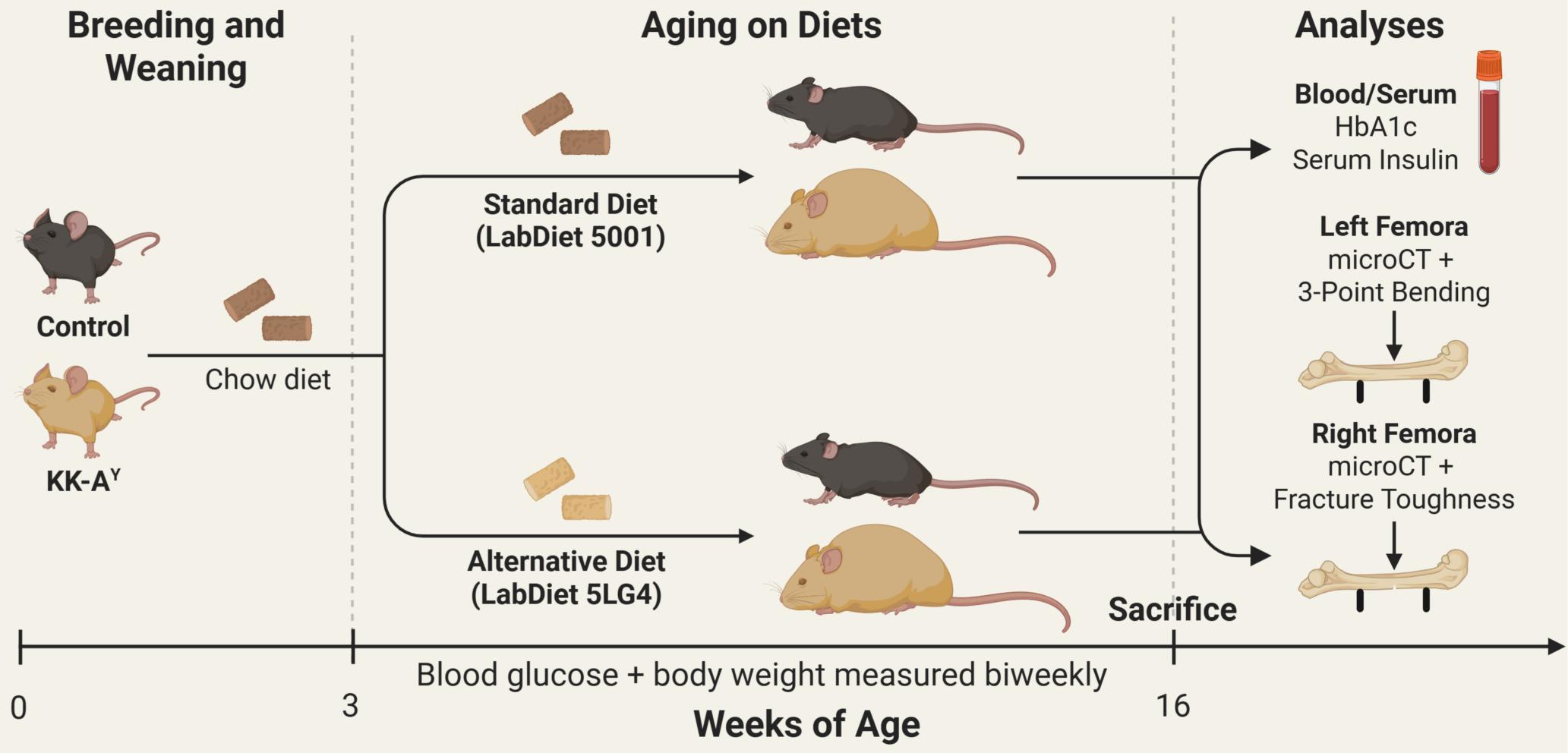
females only (B)



Three-Point Bending to Failure

- Improved strength at the expense of ductility in KK-A^Y males
 (E) and females (F)
- 5LG4 diet improved strength and ductility in each genotype in males (E) and females (F)

Methods



Conclusions

The KK-A^Y model is impacted by diet in a sexdependent manner

- KK-A^Y, 5LG4 females had the most compelling metabolic T2D outcomes
- Control, Standard females were the only normoglycemic (Blood Glucose < 250 mg/dL) group
- Improved cortical properties with lower mechanics model T2D bone phenotype
- Females are ideal in the use of the KK-A^Y with implementation of KK-A^Y, 5LG4 as a model of T2D and Control, Standard as the non-diabetic control
- Quality effects should be assessed further

Acknowledgements

This project was funded by the NIH R01 Grant (AR072609).

Special thanks to all members of the Bone Biology and Mechanics Lab.

References





