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Pre-treatment fasting glucose concentration predicts weight loss on a high fiber, low glycemic load diet: The Healthy Weight for Living Study



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Lorien E. Urban¹, Susan B. Roberts², Yishai Zohar¹, Arne Astrup³, Mads Fiil Hjorth³, Anastassios G. Pittas², Edward Saltzman², and Sai Krupa Das².

1. Gelesis, Boston MA, USA. 2. Tufts University Human Nutrition Research Center on Aging, Boston MA, USA. 3. University of Copenhagen, Denmark.

INTRODUCTION

- The struggle to curb the current obesity epidemic with an optimal diet has largely failed, giving rise to numerous insufficient approaches.¹
- Caloric restriction, although effective in the shortterm, is frequently derailed by biologic feedback mechanisms that stimulate appetite, reduce dietary compliance, and ultimately lead to a rebound of energy intake and weight gain.² Increasing dietary fiber intake has been employed as a means of influencing subjective appetite and preventing weight gain.³
- Low-carbohydrate or low-glycemic load diets have demonstrated differential success according to insulin resistance.⁴⁻⁷

- While glycemic parameters may help predict weight loss success, attempts to establish whether pretreatment glycemic status predicts or determines weight loss response have been inconsistent.⁸⁻⁹
- The purpose of this analysis was to explore fasting plasma glucose concentrations as an objective predictor of weight loss in participants of the Healthy Weight For Living Study: a 24-week comprehensive intervention based on a high-fiber, low glycemic-load diet.

METHODS

- This was a re-analysis of a previously reported 24week, group-based behavioral weight loss study.
 - The primary goal of the diet intervention was to facilitate portion-controlled menus containing ≥ 40 g/day of dietary fiber and a low glycemic load. Macronutrient targets were 25% protein, 27% fat, and 48% low-glycemic index carbohydrates.
- For this re-analysis, subjects were dichotomized by baseline fasting blood glucose (FBG) concentrations as low (FBG < 90 mg/dL) and high (FBG ≥ 90 to 125 mg/dL).
- Differences in weight loss between low and high FBG groups were examined using linear mixed models adjusted for baseline body mass index, age, gender, and site, or Fisher's exact tests ($\alpha = 0.05$).

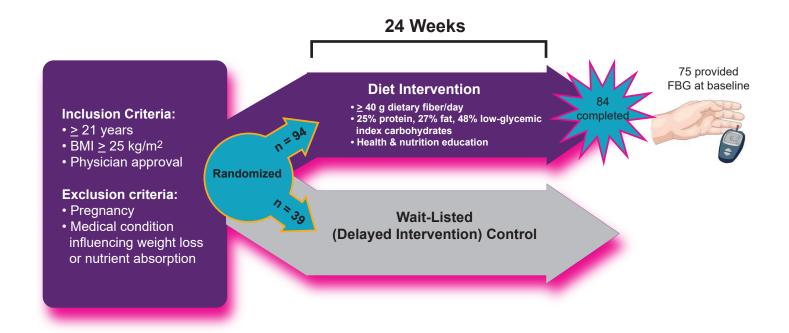


Figure 1: Six-month study design.¹⁰ Only subjects who completed the diet¹¹ intervention (purple arrow) and provided baseline fasting glucose measurements were included in the current analysis. BMI = Body Mass Index, FBG = fasting blood glucose at baseline.

RESULTS

- In the original 24-week study¹⁰, the overall mean weight loss in the intervention group was 8.0 ± 0.7 kg (**Figure 2**).
- Subjects with high FBG lost significantly more weight (-9.4%, range: -7.9% to -10.9%) at 24 weeks compared to subjects with low FBG (-4.1%, range: -1.4% to -6.9%) (P = 0.038, Figure 3). Trajectories between the two FBG groups appeared to diverge by 10 weeks and persisted thereafter.
- By 24-weeks, 79% of participants with high FBG achieved a 5% reduction in body weight, compared with 50% achievement among low FBG participants (P = 0.064, Figure 4).

- Similarly, 36% of participants with high FBG achieved a 10% reduction compared to only 8% with low FBG (P = 0.088).
- When FBG was examined as a continuous predictor of weight loss at 24 weeks, a trend for an inverse relationship was observed (β -0.12%; CI: -0.26% to .011%; P = 0.072; **Figure 5**).

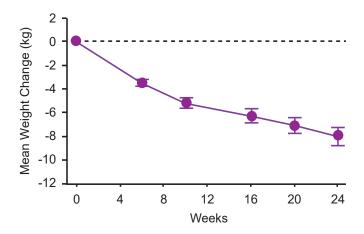
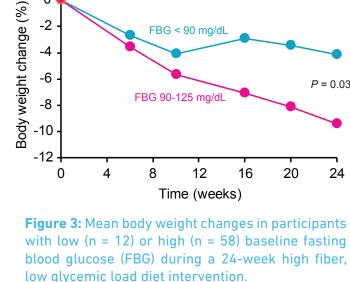


Figure 2: Body weight changes in all participants (n = 84) who completed a high fiber, low glycemic load diet intervention for 24 weeks. 10 Data presented as mean ± SEM.



FBG < 90 mg/dL

FBG 90-125 mg/dL

P = 0.038

0

-2

-4

-6

-8

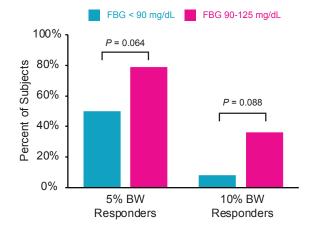


Figure 4: Achievement of 5% and 10% body weight (BW) loss at 24 weeks in participants with low or high fasting blood glucose (FBG).

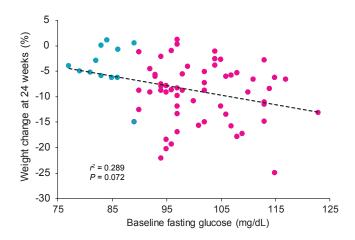


Figure 5: Inverse relationship between body weight change at 24-weeks and baseline fasting blood qlucose (FBG) levels. Blue = FBG < 90 mg/dL, Red = FBG 100-125 mg/dL.

DISCUSSION

- In this re-analysis of the Healthy Weight for Living study¹⁰, our primary observations were that 1) individuals with high FBG demonstrated an enhanced weight loss response to a high fiber, low glycemic load diet, and 2) baseline FBG tended to predict weight loss. Accordingly, a greater percentage of participants with high FBG achieved weight loss milestones of 5% and 10% by the end of the study.
- These results are consistent with other analyses of independent diet studies presented at this congress demonstrating that subjects with prediabetes respond

better to diets that are lower in glycemic load and carbohydrate content (Hjorth ADA#202-OR, Hjorth ADA#73-LB), ad libitum diets that are lower in their relative amounts carbohydrate (Astrup ADA#201-OR), or higher in fiber (Astrup ADA#792-P), and that subjects with elevated fasting plasma glucose responded preferentially to Mediterranean diets that are higher in plant-based fats (and therefore lower in carbohydrate content, Estruch ADA#78-LB).

DISCUSSION CONTINUED

- There are several central (CNS) and peripheral mechanisms by which blood glucose levels may influence satiety and ultimately weight loss; 12,13 however, we cannot conclude if this particular biomarker is responsible for the enhanced weightloss observed or whether there are other markers or mechanisms involved. For example, the diet employed in this study was rich in dietary fibers and strong evidence suggests a role for gut microbiota in their fermentation and production of metabolites such as short-chain fatty acids that may facilitate weight loss. 14
- Over the past several decades, numerous trials have compared various diets for the management of obesity based on the assumption that a single dietary strategy is appropriate for all individuals. Our results clearly demonstrate that failure to consider glycemic status may underestimate true weight loss potential.

CONCLUSIONS

- These novel results, along with other analyses of large, international diet studies presented at this congress (73-LB, 78-LB, 792-P, 201-OR, 202-OR), demonstrate that easily accessible biomarkers such as fasting
- blood glucose are strong predictors of dietary weight loss success and represent a significant step forward in personalized weight management.

REFERENCES

- 1.Freedhoff Y, Hall KD. Weight loss diet studies: we need help not hype. The Lancet. 2016;388(10047):849-851.
- 2.MacLean PS, et al. Biological control of appetite: A daunting complexity. Obesity. 2017;25 Suppl 1:S8-s16.
- 3. Wanders, et al. Effects of dietary fibre on subjective appetitie, energy intake and body weight: a systematic review of controlled trials. Obesity Reviews. 2011;12:724-739.
- 4.McClain AD, et al. Adherence to a low-fat vs. low-carbohydrate diet differs by insulin resistance status. Diabetes, Obesity and Metabolism. 2013;15(1):87-90.
- 5.Gardner CD, et al. Weight loss on low-fat vs. low-carbohydrate diets by insulin resistance status among overweight adults and adults with obesity: A randomized pilot trial. Obesity. 2016;24(1):79-86.
- 6.Pittas AG, et al. A low-glycaemic load diet facilitates greater weight loss in overweight adults with high insulin secretion but not in overweight adults with low insulin secretion in the CALERIE Trial. Diabetes Care. 2005 Dec;28(12):2939-41.
- 7.Ebbeling CB, et al. Effects of a low–glycaemic load vs low-fat diet in obese young adults: a randomized trial. JAMA. 2007:297(19):2092-102.

- 8.Astrup A, et al. Oral administration of Gelesis100, a novel hydrogel, significantly decreases body weight in overweight and obese subjects. In: Program of the Endocrine Society 96th Annual Meeting. June 21-24, 2014; Chicago, IL.
- 9.de Luis DA, et al. Differences in glycaemic status do not predict weight loss in response to hypocaloric diets in obese patients. Clin Nutr. 2006;25(1):117-22.
- 10. Salinardi TC, et al. Lifestyle intervention reduces body weight and improves cardiometabolic risk factors in worksites. Am J Clin Nutr. 2013;97:667–76.
- 11.Roberts SB, Sargent BK. The "I" diet. New York, NY: Workman Publishing, 2010.
- 12.Chandler-Laney PC, et al. Return of hunger following a relatively high carbohydrate breakfast is associated with earlier recorded glucose peak and nadir. Appetite. 2014;80:236-41.
- 13.Luo S, et al. Differential effects of fructose versus glucose on brain and appetitive responses to food cues and decisions for food rewards. Proc Natl Acad Sci. 2015;112(20):6509-14.
- 14.Brahe LK, et al. Can we prevent obesity-related metabolic diseases by dietary modulation of the gut microbiota? Adv Nutr. 2016;7:90-101.