

Comment on: Very-Low-Calorie Ketogenic Diet as a Safe and Valuable Tool for Long-Term Glycemic Management in Patients with Obesity and Type 2 Diabetes.

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With interest, we have read the article of Moriconi et al ¹ entitled “Very-Low-Calorie Ketogenic Diet as a Safe and Valuable Tool for Long-Term Glycemic Management in Patients with Obesity and Type 2 Diabetes.” A very low carbohydrate ketogenic diet (VLCHKD; <10% energy from CHO/day) is a popular, often effective dietary intervention for people with obesity related-type 2 diabetes mellitus (T2DM) ²⁻⁶. However, ketogenic diets applied in T2DM studies differ significantly in nutrient composition (different percentages of carbohydrate, protein, fat and types of fatty acids), resulting in a heterogeneity that influences reproducibility and interpretation of their results.

Moriconi et al ¹, did not provide data on the nutritional composition of the VLCHKD during the study period, making the interpretation and translation of the results into everyday practice difficult. Furthermore, the title of the paper indicates that ketosis was reached, but data on the achieved ketosis are not presented. As studies among adults with diabetes often opt for a diet with a ratio of fat to carbohydrates and protein of no more than 1:1, the question is whether substantial ketosis actually occurred during the intervention in the study of Moriconi and co-workers. This is in contrast to diets used e.g. in children with epilepsy, in which ratios of 3:1 and higher are often achieved, resulting in a considerable level of ketosis ^{7,8}

In addition, we have some concerns about the methodology used in the study by Moriconi et al¹. First of all, the significant differences in baseline anthropometric measures between the cohorts certainly influence the results. For example, the VLCHKD group had 20 kg more excess in bodyweight and thus presumably this cohort had a higher motivation and potential for improvement. It is also unclear whether participants were free to choose the dietary intervention. Also details on the time period in which the data from the historical cohort were collected are missing.

The use of VLCHKDs is an important and promising area of research in obesity-related T2DM. However, interpretation of the available published results is currently hampered by differences in diet composition, incomplete reporting of diet composition, and differences in nomenclature as also indicated by Trimboli et al ⁹. We therefore strongly advocate a consensus on these issues in the field of obesity-related T2DM in order to improve the quality of study designs, interpretation and reproducibility of data. Although the results reported by Moriconi et al ¹ are of interest, we believe that they should be interpreted with caution.

References

1. Moriconi E, Camajani E, Fabbri A, Lenzi A, Caprio M. Very-Low-Calorie Ketogenic Diet as a Safe and Valuable Tool for Long-Term Glycemic Management in Patients with Obesity and Type 2 Diabetes. *Nutrients*. 2021 Feb 26;13(3).
2. van Zuuren EJ, Fedorowicz Z, Kuijpers T, Pijl H. Effects of low-carbohydrate- compared with low-fat-diet interventions on metabolic control in people with type 2 diabetes: a systematic review including GRADE assessments. *Am J Clin Nutr*. 2018 Aug 1;108(2):300-31.
3. Korsmo-Haugen HK, Brurberg KG, Mann J, Aas AM. Carbohydrate quantity in the dietary management of type 2 diabetes: A systematic review and meta-analysis. *Diabetes Obes Metab*. 2019 Jan;21(1):15-27.10.1111/dom.13499
4. McArdle PD, Greenfield SM, Rilstone SK, Narendran P, Haque MS, Gill PS. Carbohydrate restriction for glycaemic control in Type 2 diabetes: a systematic review and meta-analysis. *Diabet Med*. 2019 Mar;36(3):335-48.10.1111/dme.13862
5. Sainsbury E, Kizirian NV, Partridge SR, Gill T, Colagiuri S, Gibson AA. Effect of dietary carbohydrate restriction on glycemic control in adults with diabetes: A systematic review and meta-analysis. *Diabetes Res Clin Pract*. 2018 May;139:239-52.S0168-8227(17)31171-3
6. Goldenberg JZ, Day A, Brinkworth GD, Sato J, Yamada S, Jonsson T, et al. Efficacy and safety of low and very low carbohydrate diets for type 2 diabetes remission: systematic review and meta-analysis of published and unpublished randomized trial data. *BMJ*. 2021 Jan 13;372:m4743.10.1136/bmj.m4743
7. Klepper J, Leiendecker B. Glut1 deficiency syndrome and novel ketogenic diets. *J Child Neurol*. 2013 Aug;28(8):1045-8.0883073813487600
8. Longo R, Peri C, Cricri D, Coppi L, Caruso D, Mitro N, et al. Ketogenic Diet: A New Light Shining on Old but Gold Biochemistry. *Nutrients*. 2019 Oct 17;11(10).nu11102497
9. Trimboli P, Castellana M, Bellido D, Casanueva FF. Confusion in the nomenclature of ketogenic diets blurs evidence. *Rev Endocr Metab Disord*. 2020 Mar;21(1):1-3.10.1007/s11154-020-09546-9