

Biomarkers of Glucose-Insulin Homeostasis, Randomized Treatment with Omega-3 and Vitamin D Supplementation, and Incident Type 2 Diabetes: Prospective Analysis From the VITamin D and Omega-3 Trial (VITAL)

Background

- Insulin resistance is the key driver in the development of type 2 diabetes (T2D).¹
- However, data about the association between T2D and specific glucose-insulin homeostasis biomarkers have been inconsistent.
- In addition, whether such associations are influenced by supplementation with vitamin D or omega-3 fatty acids (n-3 FA) is unclear.
- **Objective:** This study examined the association between glucose-insulin biomarkers and risk of incident T2D. It also examined whether supplementation with vitamin D or n-3 FA modified any associations between the biomarkers and incident T2D risk.

Methods

- VITAL is a randomized trial investigating whether supplemental vitamin D (cholecalciferol, 2000 IU/d) or n-3 FA (EPA+DHA, 1 g/d) influence CVD and cancer risk.^{2,3}
- In this study, VITAL trial participants with incident T2D were matched to healthy control participants based on age and sex.
- The associations of incident T2D risk with baseline levels of insulin, C-peptide, and HbA1c, and insulin resistance score (IRS) were assessed with conditional logistic regression.
 - The models adjusted for demographics, T2D risk factors (eg, body mass index), and randomized treatment (ie, vitamin D or n-3 FA supplementation or placebo).
- The interactions between each of the glucose-insulin biomarkers and the treatment groups were also tested.

Results

- The study population included 359 patients with incident T2D matched to 359 healthy control participants.
 - Mean age of the matched cohorts was 66 years; 45% were female and 16% were African American.
- Baseline levels of all 4 biomarkers were associated with risk of incident T2D: adjusted odds ratios (aORs) per incremental standard deviation are shown below.
 - Insulin: aOR, 1.46; 95% CI, 1.10 to 1.94
 - C-peptide: aOR, 1.72; 95% CI, 1.28 to 2.31
 - HbA1c: aOR, 80.43; 95% CI, 25.90-249.77
 - IRS: aOR, 1.59; 95% CI, 1.26-2.01
- Vitamin D and n-3 FA supplementation did not affect the associations of the glucose-insulin biomarkers with incident T2D.

Conclusions

- In this study, higher levels of glucose-insulin homeostasis biomarkers were associated with increased risk of T2D.
- Vitamin D and n-3 FA supplementation did not affect these associations.

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