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## The effect of a low carbohydrate high fat diet on apolipoproteins and cardiovascular risk

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## Abstract

Apolipoproteins (apo) regulate lipoprotein characteristics and lipid metabolism. ApoC-III is a regulator of triglyceride-rich lipoprotein (TRL) metabolism and apolipoproteins are important biomarkers for cardiovascular disease (CVD) risk prediction. A low carbohydrate high fat (LCHF) diet improves cardiometabolic risk, especially via reduction of TRL. However, few studies have compared a LCHF vs. a high carbohydrate (HC), lower fat diet under ad libitum conditions on apoC-III levels. The objectives of this investigation were to measure the effect of a LCHF vs. a HC diet on apoC-III, apoA1, apoB and apoB/apoA1 in 16 healthy Caucasian adults aged 19-64. Ethical approval: Liverpool John Moores University Research Ethics Committee (16/ELS/029); registered with ClinicalTrials. gov (Ref. NCT03257085). Participants randomly assigned to a HC diet (UK Eatwell guidelines;  $\geq$  50% of energy from carbohydrates) (n = 8), or a LCHF diet (consume < 50 g/day of carbohydrates) (n = 8) provided plasma samples at 0, 4 and 8 weeks. ApoA1 and apoB were analysed by an automated chemistry analyser (Daytona, Randox Laboratories Ltd, UK). ApoC-III was analysed via ELISA (Thermo Fisher Ltd, USA). Factorial 2×3 ANOVA and ANCOVA (IBM SPSS 25®) were undertaken to investigate significant differences and to control for variables influenced by baseline measures and visceral adipose tissue (VAT). Results show 0, 4, and 8 weeks respectively: ApoC-III (LCHF: 19.12 ± 9.14, 16.05 ± 7.95, 15.11 ± 3.17 mg/dl; HC: 22.13 ± 8.38, 28.22 ± 13.85, 22.22 ± 7.7 mg/dl) showed no significant (P = 0.319) change. No significant (P = 0.23) change was also observed in ApoB (LCHF:  $107.25 \pm 20.35$ ,  $111.38 \pm 24.81$ ,  $111.43 \pm 19.93$  mg/dl; HC:  $94.38 \pm 20.79$ ,  $105.00 \pm 20.13$ ,  $99.00 \pm 29.09$  mg/dl). Similarly apoA1 (LCHF:  $158.71 \pm 10.93$  mg/dl) 14.27,  $166.50 \pm 23.09$ ,  $173.00 \pm 29.42$  mg/dl; HC:  $164.71 \pm 30.25$ ,  $172.50 \pm 29.44$ ,  $174.00 \pm 32.83$  mg/dl) showed no significant change (P = 0.76). This resulted in a relatively unchanged apoB/A1 throughout the study in both diets (P = 0.30). No significant (P > 0.05) differences were found after 4 weeks or between groups also. ANCOVA revealed a trend (P = 0.06) in apoC-III for a difference between groups (LCHF:  $\Delta$ -6.6 mg/dl vs. HC:  $\Delta$ 1.2 mg/dl) after 8 weeks but no significant (P > 0.05) changes in other apolipoproteins were detected. These preliminary data reveal that a LCHF diet does not improve the apolipoprotein profile; however, when accounting for other metabolic risk factors (i.e. VAT) there was a trend towards lowering apoC-III levels (P = 0.06). Modulation of apoC-III may lead to improved lipid metabolism, but higher-powered studies are warranted before any improvement on CVD risk can be inferred.

## **Conflict of Interest**

There is no conflict of interest.