

**Research Article** 

# Effects of a High Carbohydrate and High Protein Formula Diet On Body Composition and Metabolic Risk Parameters in Obese Subjects

Katharina Möller\*, Inga Schneider, Janina Willers and Andreas Hahn

Institute of Food Science and Human Nutrition, Leibniz University Hannover, Germany

## Abstract

**Background**: In obese subject's weight loss is known to improve blood lipid profiles, glycemic control and other conditions that may contribute to the development of metabolic syndrome or cardiovascular diseases. However, the optimal dietary carbohydrate and protein composition to facilitate weight loss and improving potential adverse effects is still in debate. Therefore, the aim of this study was to compare the effect of two low-fat formula diets either high in carbohydrate or high in protein, on body composition and metabolic risk factors.

**Methods**: 154 obese (BMI  $32.5 \pm 0.14 \text{ kg/m}^2$ ) men and women were included in this randomized clinical trial and classified in two groups (high carbohydrate formula diet (HC) and high protein formula diet (HP)) of 80 matched subjects. They underwent an intervention for eight weeks, which consisted of two phases: (1) week 1 and 2: total replacement of three meals by a formula diet and (2) six week partial formula diet (replacement of 1-2 meals). Measurements were taken prior and post intervention for analysis of body composition and parameters of lipid and glucose metabolism.

**Results**: After eight weeks both groups lost significantly body fat mass (HC:  $-5.11 \pm 0.51$  kg, p<0.001; HP:  $-5.81 \pm 0.54$  kg, p<0.001), while only for subjects of HP group no change of lean body mass and body cell mass was observed. Metabolic risk parameters were reduced in both the HC and HP group; however, subjects in the HC group showed a higher reduction in triacylglycerol concentration (-29.1 mg/dl vs. -14.0 mg/dl, p<0.04). Further, the prevalence of the metabolic syndrome was reduced in both groups without difference (HC: -17.9%, p=0.004; HP: -18.4%, p=0.003).

**Conclusion**: Our data demonstrate, that even in a short period of time, a low-fat meal replacement diet high in carbohydrate or high in protein is effective in improving body composition and reducing metabolic risk parameters.

**Keywords:** Carbohydrate; Protein; Formula diet; Obesity; Body composition; Metabolic risk parameters

**Abbreviations:** BFM: Body Fat Mass; BCM: Body Cell Mass; BMI: Body Mass Index; DBP: Diastolic Blood Pressure; ECM: Extracellular Mass; HC: High Carbohydrate Formula Diet; HDL-C: High-Density Lipoprotein; HipC: Hip Circumference; HOMA-IR: Homeostasis Model Assessment for Insulin Resistance; HP: High Protein Formula Diet; hsCRP: High-Sensitivity C-Reactive Protein; LDL-C: Low-Density Lipoprotein Cholesterol; SBP: Systolic Blood Pressure; SD: Standard Deviation; TAG: Triacylglycerol; TC: Total Cholesterol; TW: Total Body Water; WC: Waist Circumference.

## Introduction

High body weight is associated with an increased risk in hyperglycaemia, atherogenic dyslipidaemia and hypertension [1]. These risk factors potentiate the development of the metabolic syndrome (MetS), which is one of the most prevalent conditions that predispose to cardiovascular complications and diabetes mellitus type 2 [2]. Recommendations agree that people who are overweight and obese need to be advised and given perspective and practical strategies for weight reduction [3] due to prevent developing risk [4].

The rising levels of overweight and obesity have resulted in an increase of weight loss diets. Modest weight loss results from a reduction in energy intake, which can be achieved by a reduction in dietary fat [5]. However, the optimal dietary carbohydrate and protein composition to facilitate weight loss and improving potential adverse effects is still in debate [6-8]. Low fat/ high carbohydrate diets, which mainly of vegetable and fruit origin were still recommended cause of its effects on weight control [9,10] and cardio-protective potential [11,12]. Higher protein diets with replacing carbohydrate by protein have positive effects on glycemic control [13,14] and prevent lean body mass loss [15-18]. However, concerns have also been expressed that diets with protein from animal sources are often high in saturated fat, which have a tendency to increase risk of cardiovascular diseases [19,20]. Therefore, selecting low-calorie food with provided macronutrient composition to realize beneficial high carbohydrate or high protein diets may be difficult to maintain by conventional dietary approaches [21,22].

In addition, the use of meal replacements coupled with lowcalorie diets represent strategies to facilitate an energy deficit for weight reduction [23]. Studies suggest that formula diets considerably increase the number of responder [24,25] and improved weight-related risk factors such as waist circumferences (WC), glucose, insulin, lipid profile, and blood pressure [21,25-30].

In the current study, we aimed to clarify if formula diets high in carbohydrate or high in protein have different effects on body composition and decreasing metabolic risk. We conducted an 8-week randomized trial to compare two relatively low fat meal replacement diets – one high in carbohydrate and the other high in protein- on

\*Corresponding author: Katharina Möller, Leibniz University of Hannover, Institute of Food Science and Human Nutrition, Am Kleinen Felde 30, 30167 Hannover, Germany, Tel: 49-0-511-762 3317; Fax: 49-0-511-762 5729; E-mail: moeller@nutrition.uni-hannover.de

Received December 18, 2015; Accepted December 28, 2015; Published December 31, 2015

**Citation:** Möller K, Schneider I, Willers J, Hahn A (2015) Effects of a High Carbohydrate and High Protein Formula Diet On Body Composition and Metabolic Risk Parameters in Obese Subjects. J Obes Weight Loss Ther 5: 291. doi:10.4172/2165-7904.1000291

**Copyright:** © 2015 Möller K, et al. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

Page 2 of 7

weight loss, body composition and metabolic risk parameters in overweight and obese subjects.

# Materials and Methods

## Study design and population

The study protocol, subject selection and randomization procedure as well as anthropometric measurements methodology was described in detail in ref. [31]. In brief, 154 subjects between 30 and 65 years were randomized to two different treatment groups: (1) carbohydrate-rich formula diet (HC) and (2) protein-rich formula diet (HP). The formula diet was administered daily as a drink meal replacement. Shakes were prepared by combining 30 g of powdered HC meal replacement mix and 34 g of powdered HP meal replacement mix with 300 ml milk (low fat) and 5 g vegetable oil. The meal energy macronutrient composition per recommended preparation of the HC meal replacement was approximately 24% protein, 49% carbohydrate, 25% fat and 2% fiber. The macronutrient composition of the HP meal replacement was approximately 38% protein, 34% carbohydrate and 28% fat. Nutritional composition of both formula diets are shown in Table 1. All subjects were advised to replace three meals a day in the first two weeks of intervention. In week 3 and 4, the subjects consumed two meal replacements for both lunch and dinner. During the last intervention period (week 5 to 8), the subjects were encouraged to replace either lunch or dinner (depending on the daily routine) with the formula diet.

## Body composition and blood sampling

The body composition was determined by means of bioelectrical impedance analysis (BIA) using a calibrated device, the Nutriguard-M (Data Input GmbH, Darmstadt, Germany) at baseline week 2 and 8 weeks ( $t_0$ ,  $t_2$ ,  $t_8$ ).

Fasting blood samples (approximately 50 ml at each time point) were collected using sealed Blood Collection Tubes and System S-Monovettes\* (Sarstedt, Germany). The samples were directly sent to an external laboratory (LADR laboratory, Hanover, Germany) for analysis of serum lipids, insulin and high-sensitivity C-reactive protein (hsCRP) as well as plasma glucose according to standardized procedures. The serum triacylglycerol (TAG) concentration, total cholesterol (TC) and high-density lipoprotein cholesterol (HDL-C) levels were measured enzymatically (Beckman Coulter, Inc.). Lowdensity lipoprotein cholesterol (LDL-C) was calculated using the Friedewald equation [32]. Fasting plasma glucose measurements were performed by using the hexokinase method, an ultraviolet (UV) enzymatic (in vitro) assay (Beckman Coulter, Inc.). Serum insulin concentrations were determined by immunoassays (cobas\*, Roche Diagnostics, Mannheim, Germany). An immunoturbidimetric method was used for determination of serum hsCRP.

## Metabolic risk profile/parameters

The NCEP ATP-III criteria was used for defining the MetS, subjects were required to fulfill at least three of the following five criteria: WC >102 (men) or >88 cm (women), blood pressure 130/85 mmHg or treatment of hypertension, fasting blood glucose  $\geq$ 100 mg/dl, TAG  $\geq$  150 mg/dl, HDL-C <40 mg/dl (men) and < 50 mg/dl (women) [33]. To enable us to estimate insulin resistance of subjects, Homeostasis model assessment of insulin resistance (HOMA-IR) were used. HOMA-IR derived from fasting glucose and insulin concentrations as follows (fasting plasma glucose x fasting serum insulin)/405) [34]. In addition, hsCRP levels  $\geq$ 3 mg/L was used for classification to assess an inflammation state [35,36].

	High carb formu	oohydrate la diet	High protein formula diet		
Nutritional values	per 100 g	per serving size*	per 100 g	per serving size**	
Energy (kcal)	371.4	313	364	311	
Carbohydrate (g)	65.6	34.1	34.0	26.0	
from Sugar (g)	64.0	33.6	33.0	25.6	
Protein (g)	22.5	16.9	56.4	29.2	
Fat (g)	0.03	7.8	0.2	9.8	
Total fiber (g)	7.4	2.2	0	0	

'Serving size as recommended preparation: 300 ml milk (1.5% fat)+30 g carbohydrate-rich 5 powder+5 g vegetable oil; "Serving size as recommended preparation: 300 ml milk (1.5% fat)+34 g protein-rich 7 powder+5 g vegetable oil.

 Table 1: Nutritional composition of the formula diets with recommended preparation.

 Statistical analysis

The results are presented as the mean value  $\pm$  standard error (SE). Statistical comparison between the groups was performed using the nonparametric Mann-Whitney U test for unpaired data. The changes in the parameters in comparison with baseline were analysed using the Wilcoxon test. The chi-square test was used to compare the difference between the frequencies of the groups. Differences were considered significant at p≤0.05. The statistical data analysis was carried out by using the Statistical Package for Social Sciences SPSS 22.0 (SPSS Inc., Chicago, Illinois, USA).

## Results

#### Subjects and baseline characteristics

Included subjects (60 male; 94 female) had an average age of 50.4 years (range: 30-65 years). The anthropometric measurements, body composition and blood parameters were quantified of 154 participants (n=78 of them received HC formula diet and n=76 the HP formula diet) before and after eight weeks of weight reduction. No differences were observed at baseline between both groups regarding to the parameters (Tables 2 and 3). The selected risk parameters did not differ between the HC and the HP group. The prevalence of MetS at baseline was 46.2% in the HC group (n=36) and 52.6% in the HP group (n=40). The prevalence of insulin resistance (according to the definition of HOMA-IR  $\geq 2$  [34]) was 67.9% (n=53) in the HC group and 71.1% (n=54) in the HP group. Further, hsCRP level >3 mg/L was observed for 37.2% of subjects in HC group and 40.8% in HP group (Figure 1).

## Change of body weight and body composition

The efficiency of both formula diets for reduction in body weight and WC has already been described in ref. [31]. In brief, the formula diet resulted in a significant weight loss of  $-6.52 \pm 0.41$  kg ( $6.61 \pm 0.38\%$ ; p<0.001) in the HC group and  $-6.79 \pm 0.42$  ( $-6.96 \pm 0.41\%$ ; p<0.001) in the HP group after eight weeks. Further, both groups showed significant reductions in WC (p<0.001) respectively.

Eight weeks of dietary intervention resulted in a significant reduction of body fat mass (BFM) in both HC group (-5.11  $\pm$  0.51 kg; -14.3  $\pm$  1.51%; p<0.001) and HP group (-5.81  $\pm$  0.54 kg; -16.2  $\pm$  1.51%; p<0.001) without significant difference between the groups. During the eight-week weight loss phase, body cell mass (BCM) and lean body

## Page 3 of 7

		HC group (n=78)	P (f _>f )*	HP group (n=76)	P (f >f )*	P (HC vs.
			· (* <sub>0</sub> * <sub>8</sub> /		· (*° <sup>-,</sup> *8)	HP)**
		Mean ± SE		mean ± SE		
Weight (kg)	to	98.2 ± 1.46		97.6 ± 1.42		n.s
	t <sub>s</sub>	91.7 ± 1.39	<0.001	90.8 ± 1.39	<0.001	n.s
BMI (kg/m²)	to	32.4 ± 0.21		32.5 ± 0.18		n.s
	t <sub>s</sub>	30.3 ± 0.24	<0.001	30.3 ± 0.23	<0.001	n.s
	to	106 ± 1.11		106 ± 1.10		n.s
	t <sub>s</sub>	97.4 ± 0.98	<0.001	97.5 ± 0.95	<0.001	n.s
	to	115 ± 0.73		115 ± 0.73		n.s
	t <sub>s</sub>	109 ± 0.71	<0.001	109 ± 0.66	<0.001	n.s
SBD (mm/Ha)	to	138 ± 1.73		142 ± 2.11		n.s
SBF (mm/rig)	t <sub>s</sub>	129 ± 1.39	<0.001	129 ± 1.58	<0.001	n.s
	$t_o$	84.5 ± 1.15		85.3 ± 1.27		n.s
DBF (mm/ng)	t <sub>s</sub>	79.9 ± 0.91	<0.001	79.7 ± 0.99	<0.001	n.s
	$t_o$	63.1 ± 1.49		61.6 ± 1.44		n.s
LBM (Kg)	t <sub>s</sub>	61.5 ± 1.32	0.007	60.6 ± 1.40	n.s	n.s
BFM (kg)	to	35.3 ± 0.76		36.1 ± 0.72		n.s
	t <sub>s</sub>	30.2 ± 0.79	<0.001	30.2 ± 0.81	<0.001	n.s
ECM (kg)	to	30.3 ± 0.66		30.2 ± 0.64		n.s
	t <sub>s</sub>	29.9 ± 0.62	n.s	29.2 ± 0.61	<0.001	n.s
BCM (kg)	to	32.6 ± 0.90		31.4 ± 0.89		n.s
	t <sub>s</sub>	31.7 ± 0.78	0.009	31.4 ± 0.87	n.s	n.s
ECM/BCM	to	0.95 ± 0.02		0.99 ± 0.02		n.s
	t <sub>s</sub>	0.96 ± 0.01	n.s	$0.95 \pm 0.02$	<0.001	n.s
<b>TW</b> (I)	to	46.1 ± 1.09		45.1 ± 1.05		n.s
	t <sub>s</sub>	45.0 ± 0.97	0.006	44.4 ± 1.02	0.049	n.s

Abbreviations: BFM: body fat mass; BCM: body cell mass; BMI: body mass index; DBP: diastolic blood pressure; ECM: extracellular mass; HC: high carbohydrate formula diet; Hip C: hip circumference; HP: high protein formula diet; SBP: systolic blood pressure; TW: total body water; WC: waist circumference.

**Table 2:** Anthropometric data and body composition in the HC and HP group at baseline ( $t_0$ ) and after eight weeks ( $t_a$ ).

		HC group (n=78)	P (t <sub>0</sub> _>t <sub>8</sub> )*	HP group (n=76) mean ± SE	P (t <sub>0</sub> >t <sub>8</sub> )*	<b>P</b> (HC vs. HP)**
		mean ± SE				
Glucose (mg/dl)	to	90.5 ± 1.14		90.4 ± 0.98		n.s
	t <sub>s</sub>	88.8 ± 1.35	0.006	88.7 ± 0.99	0.045	n.s
<b>Insulin</b> (µU/ml)	to	12.5 ± 0.78		12.5 ± 0.71		n.s
	t <sub>s</sub>	8.94 ± 0.56	<0.001	8.65 ± 0.56	<0.001	n.s
HOMA-IR	to	2.87 ± 0.21		2.83 ± 0.18		n.s
	t <sub>s</sub>	2.01 ± 0.15	<0.001	1.93 ± 1.21	<0.001	n.s
TC-C (mg/dl)	to	234 ± 5.79		226 ± 4.75		n.s
	t <sub>s</sub>	215 ± 5.00	<0.001	214 ± 4.60	<0.001	n.s
LDL-C (mg/dl)	to	151 ± 4.63		148 ± 4.26		n.s
	t <sub>s</sub>	138 ± 4.10	<0.001	138 ± 3.79	0.001	n.s
HDL-C (mg/dl)	to	56.2 ± 1.47		53.9 ± 1.55		n.s
	t <sub>s</sub>	47.2 ± 1.27	<0.001	51.1 ± 1.36	0.001	n.s
LDL/HDL	to	2.77 ± 0.09		2.86 ± 0.11		n.s
	t <sub>s</sub>	2.69 ± 0.08	n.s	2.79 ± 0.08	n.s	n.s
TAG (mg/dl)	to	151 ± 7.99		136 ± 6.52		n.s
	t <sub>s</sub>	124 ± 6.46	<0.001	122 ± 5.76	0.040	n.s
hsCRP (mg/l)	to	3.19 ± 0.47		3.29 ± 0.36		n.s
	t <sub>s</sub>	2.64 ± 0.49	0.007	2.48 ± 0.28	0.002	n.s
Abbroviations: H(	^ high	carbohydrate formula diet: HDL-C-L	high-density linoprotein: HON	ALIP: homeostasis model asses	sment for insulin resistanc	e. HD. high protein

Abbreviations: HC: high carbohydrate formula diet; HDL-C: high-density lipoprotein; HOMA-IR: homeostasis model assessment for insulin resistance; HP: high protein formula diet; hsCRP: high-sensitivity C-reactive protein; LDL-C: low-density lipoprotein cholesterol; TAG: triacylglycerol; TC: total cholesterol.

Table 3: Clinical parameters in the HC and HP group at baseline ( $t_0$ ) and after eight weeks ( $t_8$ ).



mass (LBM) among the HC group reduced by a mean of  $0.99 \pm 0.33$  kg and  $1.42 \pm 0.46$  kg, representing a  $2.12 \pm 0.94\%$  and  $1.73 \pm 0.70\%$ ; decreased from baseline (p=0.009; p=0.007), whereas the HP group has been shown no significant changes from baseline (p=0.928; p=0.051) (Table 2).

## Change of clinical and metabolic risk parameters

At week eight, both groups significantly lowered fasting blood glucose and insulin levels by a mean of  $-1.72 \pm 0.82$  mg/dl glucose  $(-1.78 \pm 0.83\%; p=0.006)$  and  $-3.60 \pm 0.53 \mu U/ml$  insulin  $(-22.3 \pm 3.93\%;$ p<0.001) in HC group as well as -1,70 ± 0.87 mg/dl glucose (-1.56 ± 0.95%; p=0.045) and  $-3.86 \pm 0.52 \mu U/ml$  insulin (-27.57  $\pm 3.57\%$ ; p<0.001) in HP group. Similar, both groups experienced significant changes in low TC, LDL-C but also HDL-C at eight weeks (each p<0.005, respectively), without significant differences between the groups. During the eight week intervention, serum concentrations of TAG decreased in the HC (-12.10 ± 3.26%) and HP (-4.29 ± 3.68%) group (p<0.001 and p<0.040). This difference reflected a significant higher reduction in TAG concentration after the HC formula diet (p<0.040). Regarding the metabolic risk profile, after eight week dietary intervention the prevalence of subjects with HOMA-Index  $\geq 2$  and hsCRP levels  $\geq 3$  mg/ dl decreased in both groups significantly (p<0.001, respectively) (Figure 1). In addition, after eight week of dietary intervention, the prevalence of MetS decreased in the HC group by 17.9% (n=14; p=0.004) and in the HP group by 18.4% (n=14; p=0.003). Further, subjects with MetS showed significant reduction of TC concentration by a mean of (-26.3  $\pm$  6.75 mg/dl; p) in HC and (-10.6  $\pm$  5.29 mg/dl) in HP group. Only the subjects in HC group with Mets reduced significantly LDL-C (-15.4  $\pm$ 5.51 mg/dl). The decreased TAG concentration was significantly higher (p=0.001) in subjects with MetS of HC group (-65.1  $\pm$  11.06 mg/dl) compared to HP group (-23.3  $\pm$  10.56 mg/dl) (Figure 2).

# **Dietary Intake**

At baseline, the reported energy intake was similar in both groups (HC: 2454  $\pm$  88.2 kcal/d; HP: 2371  $\pm$  95.6 kcal/d). In addition, the percent energy intake from protein (HC: 16.1  $\pm$  0.41 %; HP 16.9  $\pm$ 

0.42%), carbohydrate (HC: 42.4  $\pm$  1.04%; HP: 42.1  $\pm$  0.94%) and fat (HC: 36.7  $\pm$  0.68%; HP: 38.3  $\pm$  0.78%) did not differ between both groups at baseline. After the eight week intervention energy intake was significantly reduced within each group but not between both groups (HC: 1530  $\pm$  61.8 kcal/d; HP: 1559  $\pm$  56.8 kcal/d). Percent energy intake from carbohydrates and proteins in HC group (44.8  $\pm$  0.96% and 20.7  $\pm$  0.52%) was significantly different compared to HP group (42.1  $\pm$  0.91%; p=0.034 and 23.7  $\pm$  0.50%; p<0.001).

# Discussion

We previously described that both formula diets are effective strategies for weight loss and WC reduction in overweight and obese subjects [31]. In this context the influence of energy restriction and macronutrients composition of both formula diets on body composition and metabolic risk parameters will be discuss.

A rapid weight loss induced by energy restricted diets has often been associated with loss of BFM in addition to the reduction of LBM [15,37]. LBM may lead to retention of resting energy expenditure, which is essential for weight maintaining after weigh loss [38]. We observed that both low-fat formula diets induce comparable reductions of BFM, but only subjects of HP group showed retention of LBM and BCM. This is in accordance to studies, which suggest that high protein diets have been associated with no change of LBM compared with high carbohydrate diets [14,17,18] even when there has been no differences in change of body weight. The required daily protein intake is 0.83 g/ kg of body weight. However, this reference protein intake per day may not be directly applied to overweight and obese subjects, because they are closely associated to LBM and this LBM is lower than in normal weight persons [39]. Therefore, the European Food Safety Authority (EFSA) suggested a daily protein intake during energy restriction diets of 75 g/day to preserve LBM in obese subjects [39]. Further, a metaanalysis of 87 weight loss diets comparing different macronutrient compositions, suggested that protein intake >1.05 g/kg body weight per day is associated with greater retention of LBM compared with lower protein intake [17]. During the partial meal replacement plan the daily protein intake in HC group was about 0.82 g/kg (75.2 g protein/day)

Page 4 of 7



and in HP group 0.96 g/kg (87.3 g protein/day). Therefore, the daily protein intake in HP meal replacement plan achieved more than 75 g protein/day and was nearly the cut off level >1.05 g/kg, thus it may be possible to attribute the observed difference of LBM retention by different protein intake. Nevertheless, to avoid LBM and BCM loss and increase energy expenditure, dieting should be combined with physical exercise [16]. However, our results demonstrate that the retention in BCM and LBM in HP group was probably not caused by an increase in physical activity, because these levels were similar in both groups [31]. Studies suggest that high GL diets and elevated insulin response might preserve the release of free fatty acids from adipose tissue due to the lipogenic actions of insulin, which maintain the BFM [40]. Considering the HC formula diets was classified as high GL food and the HP as low GL food [41] we found similar reduction of BFM after consumption of both formula diets.

In the present study, both formula diets were similar in their effect on blood lipid reduction. Although a beneficial effect of weight loss on blood lipids was found in both intervention groups, more favorable improvements were seen in the HC group. There was a significantly greater reduction in serum TAG levels after eight weeks in obese subjects with and without MetS. These changes in TAG levels were in disagreement with other studies, which found that higher carbohydrate diets are associated to increase production of TAG [15,26,42]. Diets containing lower amounts of carbohydrates would lead to a lower synthesis in very low density lipoproteins (VLDL) and potentially lower TAG storage in the liver [43]. The inconsistence between our results and others might be because the consumed meals during partial meal replacement led to a lower difference in macronutrient intake of both groups resulting in an underestimation effect of the HC and HP diet plan on TAG concentrations. The reduced HDL-C levels during the intervention of this low-fat diets is well known [30]. Improvements of HDL-C have commonly been observed in studies lasting, more than 12 weeks [37] or combined increased physical activity [44-46].

For metabolic risk estimation also the fat distribution is useful [47]. Subjects of this study exceed the cut of value for WC of 102 cm in men  $(112 \pm 0.73 \text{ cm})$  and 88 cm  $(101 \pm 0.98 \text{ cm})$  in women. The accumulation of visceral adipose tissue is closely associated with decrease of insulin sensitivity and low-grade inflammation state [48], which were considered primary factors in the development of insulin resistance, type 2 diabetes and MetS [49]. The value of HOMA-IR <2 was regarded normal ≥2 was pathologic and values >4 represented the prediabetic phase [50]. The prevalence of subjects with HOMA-IR ≥2 was 67.9% in HC and 71.1% in HP group implying a pathological to prediabetic stage. Visceral obesity is further viewed as a key factor responsible for the up regulation of inflammation in obese subjects [48,51]. Regarding the mean hsCRP levels, subjects of both groups demonstrate values of hsCRP ≥3 mg/L (37.2% in HC and 40.8% in HP group), which are thought to reflect an elevated inflammation state and considered as predictor for future cardiovascular and metabolic diseases [35,36]. In contrast to studies which shown that HP diets lowered CRP more effectively in women with high TAG concentrations [26] hsCRP levels in this study were not influenced by dietary composition. Nevertheless, in the present study the significant decrease of body weight and WC in both groups suggests a change in abnominal fat distribution, which would explain the reduction of HOMA-IR and hsCRP levels independent of dietary macronutrient composition [52].

Further, we demonstrated that consumption of the HC and HP formula diets led to a significant reduction in the prevalence of MetS,

which is one of the most prevalent conditions that predict diabetes and cardiovascular diseases [2,53]. Some studies found that an energy-restricted high protein diet or high carbohydrate diet resulted in a significant decrease of MetS prevalence [54], while others found no effect of different macronutrient distribution on MetS prevalence [55,56]. Therefore, the influence of macronutrients on MetS prevalence remains unclear. Given the importance of abdominal fat and lipid metabolism in the pathogenesis of MetS [57], it may be speculated that the observed reduction of WC and improvements of lipid profile in this study could be the reasons for reduced prevalence of MetS even in this short time period.

#### Conclusions

In addition to similar weight loss, these data demonstrate that both low-calorie meal replacement diets - high in carbohydrate or high in protein - were effective strategies for improvement of body composition and metabolic risk parameters. In this short term, the HC formula diet produced greater improvement in TAG concentration, while the HP formula diet led to a better maintenance of LBM. On this basis, subjects on energy restricted weight loss diets could have been able to choose meal replacements high in carbohydrate or high in protein to facilitate the required macronutrient ratio of the diet.

#### References

- Alberti K, Eckel RH, Grundy SM, Zimmet PZ, Cleeman JI, et al. (2009) Harmonizing the metabolic syndrome: a joint interim statement of the international diabetes federation task force on epidemiology and prevention; national heart, lung, and blood institute; american heart association; world heart federation; international atherosclerosis society; and international association for the study of obesity. Circulation 120: 1640-1645.
- Mottillo S, Filion KB, Genest J, Joseph L, Pilote L, et al. (2010) The metabolic syndrome and cardiovascular risk a systematic review and meta-analysis. J Am Coll Cardiol 56: 1113-1132.
- Astrup A (2008) Dietary management of obesity. JPEN J Parenter Enteral Nutr 32: 575-577.
- Harder H, Dinesen B, Astrup A (2004) The effect of a rapid weight loss on lipid profile and glycemic control in obese type 2 diabetic patients. Int J Obes Relat Metab Disord 28: 180-182.
- Hall KD, Heymsfield SB, Kemnitz JW, Klein S, Schoeller DA, et al. (2012) Energy balance and its components: implications for body weight regulation. Am J Clin Nutr 95: 989-994.
- Sacks FM, Bray GA, Carey VJ, Smith SR, Ryan DH, et al. (2009) Comparison of weight-loss diets with different compositions of fat, protein, and carbohydrates. N Engl J Med 360: 859-873
- Abete I, Parra MD, Zulet MA, Martínez JA (2006) Different dietary strategies for weight loss in obesity: role of energy and macronutrient content. Nutr Res Rev 19: 5-17.
- Abete I, Astrup A, Martínez JA, Thorsdottir I, Zulet MA (2010) Obesity and the metabolic syndrome: role of different dietary macronutrient distribution patterns and specific nutritional components on weight loss and maintenance. Nutr Rev 4: 214-231.
- Du H, van der A DL, Boshuizen HC, Forouhi NG, Wareham NJ, et al. (2010) Dietary fiber and subsequent changes in body weight and waist circumference in European men and women. Am J Clin Nutr 91: 329-336.
- Jebb SA (2015) Carbohydrates and obesity: from evidence to policy in the UK. Proc Nutr Soc 74: 215-220.
- Barnard ND, Levin SM, Yokoyama Y (2015) A systematic review and metaanalysis of changes in body weight in clinical trials of vegetarian diets. J Acad Nutr Diet 115: 954-969.
- Buckland G, Bach A, Serra-Majem L (2008) Obesity and the Mediterranean diet: a systematic review of observational and intervention studies. Obes Rev 9: 582-593.

Page 6 of 7

- Layman, Donald K, Baum, Jamie I (2004) Dietary protein impact on glycemic control during weight loss. J Nutr 134: 968-973.
- 14. Farnsworth E, Luscombe Natalie D, Noakes M, Wittert G, Argyiou E, et al. (2003) Effect of a high-protein, energy-restricted diet on body composition, glycemic control, and lipid concentrations in overweight and obese hyperinsulinemic men and women. Am. J Clin Nutr 78: 31-39.
- Layman DK, Boileau RA, Erickson DJ, Painter, James E, et al. (2004) A reduced ratio of dietary carbohydrate to protein improves body composition and blood lipid profiles during weight loss in adult women. J Nutr 133: 411-417.
- Deibert P, König D, Schmidt-Trucksaess A, Zaenker KS, Frey I, et al. (2004) Weight loss without losing muscle mass in pre-obese and obese subjects induced by a high-soy-protein diet. Int J Obes Relat Metab Disord 28: 1349-1352.
- Krieger James W, Harry S Sitren, Michael J Daniels, and Bobbi Langkamp-Henken (2006) Effects of variation in protein and carbohydrate intake on body mass and composition during energy restriction: a meta-regression 1. Am J Clin Nutr 83: 260-274.
- Leidy HJ, Carnell NS, Mattes RD, Campbell Wayne W. (2007) Higher protein intake preserves lean mass and satiety with weight loss in preobese and obese women. Obesity 15 2: 421-429.
- Foo SY, Heller ER, Wykrzykowska J, Sullivan CJ, Manning-Tobin JJ, et al. (2009) Vascular effects of a low-carbohydrate high-protein diet. Proc Natl Acad Sci U S A 106: 15418-15423.
- Lagiou P, Sandin S, Lof M, Trichopoulos D, Adami HO, et al. (2012) Low carbohydrate-high protein diet and incidence of cardiovascular diseases in Swedish women: prospective cohort study. BMJ 344: e4026.
- Ditschuneit HH, Flechtner-Mors M, Johnson TD, Adler G (1999) Metabolic and weight-loss effects of a long-term dietary intervention in obese patients. Am J Clin Nutr 69: 198-204.
- 22. Ashley JM, Herzog H, Clodfelter S, Bovee V, Schrage J, et al. (2007) Nutrient adequacy during weight loss interventions: a randomized study in women comparing the dietary intake in a meal replacement group with a traditional food group. Nutr J 6: 12.
- Leeds AR (2014) Formula food-reducing diets: A new evidence-based addition to the weight management tool box. Nutr Bull 39: 238-246.
- 24. Cheskin LJ, Mitchell AM, Jhaveri AD, Mitola AH, Davis LM, et al. (2008) Efficacy of meal replacements versus a standard food-based diet for weight loss in type 2 diabetes: a controlled clinical trial. Diabetes Educ 34: 118-127.
- 25. Metzner CE, Folberth-Vögele A, Bitterlich N, Lemperle M, Schäfer S, et al. (2011) Effect of a conventional energy-restricted modified diet with or without meal replacement on weight loss and cardiometabolic risk profile in overweight women. Nutr Metab (Lond) 8: 1-9.
- 26. Noakes M, Keogh JB, Foster PR, Clifton PM (2005) Effect of an energyrestricted, high-protein, low-fat diet relative to a conventional high-carbohydrate, low-fat diet on weight loss, body composition, nutritional status, and markers of cardiovascular health in obese women. Am J Clin Nutr 81: 1298-1306.
- 27. Lee K, Lee J, Bae WK, Choi JK, Kim HJ, et al. (2009) Efficacy of low-calorie, partial meal replacement diet plans on weight and abdominal fat in obese subjects with metabolic syndrome: a double-blind, randomised controlled trial of two diet plans one high in protein and one nutritionally balanced. Int J Clin Pract 63: 195-201.
- Flechtner-Mors M, Boehm BO, Wittmann R, Thoma U, Ditschuneit HH (2010) Enhanced weight loss with protein-enriched meal replacements in subjects with the metabolic syndrome. Diabetes Metab Res Rev 26: 393-405.
- Frestedt JL, Young LR, Bell M (2012) Meal Replacement Beverage Twice a Day in Overweight and Obese Adults (MDRC2012-001). Curr Nutr Food Sci 8: 320-329.
- König D, Deibert P, Frey I, Landmann U, Berg A (2008) Effect of meal replacement on metabolic risk factors in overweight and obese subjects. Ann Nutr Metab 52: 74-78.
- Möller K, Willers J, Hahn A (2015) Efficacy of high carbohydrate versus high protein meal replacements on weight reduction - A randomized controlled trial. J Obes Weight Loss Ther 05: 1-9.
- Friedewald WT, Levy RI, Fredrickson DS (1972) Estimation of the concentration of low-density lipoprotein cholesterol in plasma, without use of the preparative ultracentrifuge. Clin Chem 18: 499-502.

- 33. National Cholesterol Education Program Committee (2002) Third report of the national cholesterol education program (NCEP) expert panel on detection, evaluation, and treatment of high blood cholesterol in adults (Adult Treatment Panel III) 02-5215.
- 34. Matthews D.R, Hosker JP, Rudenski AS, Naylor BA, Treacher DF, et al. (1985) Homeostasis model assessment: insulin resistance and beta-cell function from fasting plasma glucose and insulin concentrations in man. Diabetologia 28: 412-419.
- Ridker PM (2007) C-reactive protein and the prediction of cardiovascular events among those at intermediate risk: moving an inflammatory hypothesis toward consensus. J Am Coll Cardiol 49: 2129-2138.
- Ridker PM, Cook N (2004) Clinical usefulness of very high and very low levels of C-reactive protein across the full range of Framingham Risk Scores. Circulation 109: 1955-1959.
- Layman DK, Evans EM, Erickson D, Seyler J, Weber J, et al. (2009) A moderate-protein diet produces sustained weight loss and long-term changes in body composition and blood lipids in obese adults. J Nutr 139: 514-521.
- Vogels N, Diepvens K, Westerterp-Plantenga MS (2005) Predictors of longterm weight maintenance. Obes Res 13: 2162-2168.
- EFSA NDA Panel (2015) Scientific Opinion on the essential composition of total diet replacements for weight control. EFSA Journal 1: 3957 52p.
- 40. Goss AM, Goree LL, Ellis AC, Chandler-Laney PC, Casazza K, et al. (2013) Effects of diet macronutrient composition on body composition and fat distribution during weight maintenance and weight loss. Obesity (Silver Spring) 21: 1139-1142.
- Möller K, Willers J, Schneider I, Hahn A (2015) Glycemic index and glycemic load of a carbohydrate-rich and protein-rich formula diet. J Nutr Health Sci 2: 404.
- 42. Skov AR, S Toubro, B Rùnn, L Holm and A Astrup (1999) Randomized trial on protein vs carbohydrate in ad libitum fat reduced diet for the treatment of obesity. Int J Obes 23: 528-536.
- 43. Clifton PM, Bastiaans K, Keogh JB (2009) High protein diets decrease total and abdominal fat and improve CVD risk profile in overweight and obese men and women with elevated triacylglycerol. Nutr Metab Cardiovasc Dis 19: 548-554.
- 44. Kodama S, Tanaka S, Saito K, Shu M, Sone Y, et al. (2007) Effect of aerobic

exercise training on serum levels of high-density lipoprotein cholesterol: a meta-analysis. Arch Intern Med 167: 999-1008.

Page 7 of 7

- 45. Ahmed HM, Blaha MJ, Nasir K, Rivera JJ, Blumenthal RS (2012) Effects of physical activity on cardiovascular disease. Am J Cardiol 109: 288-295.
- 46. Pattyn N, Cornelissen VA, Eshghi SR, Vanhees L (2013) The effect of exercise on the cardiovascular risk factors constituting the metabolic syndrome: a metaanalysis of controlled trials. Sports Med 43: 121-133.
- Després JP (2012) Body fat distribution and risk of cardiovascular disease: an update. Circulation 126: 1301-1313.
- Van Gaal LF, Mertens IL, De Block CE (2006) Mechanisms linking obesity with cardiovascular disease. Nature 444: 875-880.
- Chen L, Chen R, Wang H, Liang F (2015) Mechanisms Linking Inflammation to Insulin Resistance. Int J Endocrinol 2015: 508409.
- 50. Sink Angelica 2007 Insulin resistance in patients with chronic hepatitis C. TMJ 4: 1-5.
- Alexopoulos N, Katritsis D, Raggi P (2014) Visceral adipose tissue as a source of inflammation and promoter of atherosclerosis. Atherosclerosis 233: 104-112.
- Belza A, Toubro S, Stender S, Astrup A (2009) Effect of diet-induced energy deficit and body fat reduction on high-sensitive CRP and other inflammatory markers in obese subjects. Int J Obes (Lond) 33: 456-464.
- 53. Ford ES (2005) Risks for all-cause mortality, cardiovascular disease, and diabetes associated with the metabolic syndrome: a summary of the evidence. Diabetes Care 28: 1769-1778.
- Mc Keown NM, Meigs JB, Simin Liu, Saltzman E, Wilson PW, Jacques Paul F. (2004) Carbohydrate nutrition, insulin resistance, and the prevalence of the metabolic syndrome in the framingham offspring cohort. Diabetes Care 27: 538-546
- 55. Rajaie S, Azadbakht L, Khazaei M, Sherbafchi M, Esmaillzadeh A (2014) Moderate replacement of carbohydrates by dietary fats affects features of metabolic syndrome: A randomized crossover clinical trial. Nutrition 30: 61-68.
- 56. Papadaki A, Linardakis M, Plada M, Larsen TM, Damsgaard CT, et al. (2014) Impact of weight loss and maintenance with ad libitum diets varying in protein and glycemic index content on metabolic syndrome. Nutrition 30: 410-417.
- 57. Després JP, Lemieux I (2006) Abdominal obesity and metabolic syndrome. Nature 444: 881-887.