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The Effect of Eight Weeks Functional Exercise Training and Low Carbohydrate Diet on The Levels of Adiponectin, CRP and Lipid Profile in Overweight Women

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ABSTRACT

Introduction: Exercise training and a low-carbohydrate diet have shown favorable effects on inflammatory cytokines. This study examines the combined impact of an eight-week functional exercise training program and a low-carbohydrate (LC) diet on adiponectin, CRP, and lipid profiles in overweight women.

Material & Methods: Forty-eight overweight women, aged 25-35, were randomly selected for the intervention and allocated into control (n=12), LC diet (n=12), training (n=12), and training+ LC diet (n=12) groups. Functional training, conducted three times weekly for eight weeks at 50-80 percent of maximum heart rate, was paired with an LC diet restricting carbohydrate intake to 50g/d. Blood samples collected before and 48 hours after the final session were analyzed for adiponectin and CRP levels using the ELISA method. Data were analyzed using SPSS software with analysis of covariance and Bonferroni post hoc tests.

Results: Adiponectin levels did not significantly change following the eightweek interventions (p=0.135). However, CRP levels significantly decreased in the LC diet group compared to the control group (p=0.003). Cholesterol, LDL, and triglyceride levels significantly decreased in all experimental groups compared to the control group (p<0.05), while HDL levels significantly increased in the LC diet (p=0.009) and training+ LC diet (p=0.001) groups compared to the control group.

Conclusion: The study underscores the positive impact of LC diet and functional training, either alone or in combination, on improving lipid profiles. However, there seems to be no synergistic effect of LC diet with functional training on changing CRP and adiponectin levels.

Keywords: Exercise Therapy, Adiponectin, C Reactive Protein, Overweight

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Introduction

Low-grade inflammation is a wellrecognized characteristic of the obese state, with adipose tissue releasing various inflammatory mediators into circulation (1). This chronic inflammation is a key player in the pathogenesis of several chronic diseases associated with obesity, such as metabolic syndrome, cardiovascular disease, non-alcoholic fatty liver disease (NAFLD), diabetes, hypertension, and certain cancers (2). During adipose tissue expansion and obesity, there is an alteration in the secretion profile of adipokines. This involves the upregulation of adipokines, inflammatory including interleukin (IL-6),monocyte chemoattractant protein-1 (MCP-1),tumor necrosis factor-alpha (TNF-α), alongside a decrease in the levels of antiinflammatory adipokines such adiponectin and IL-10 (3). Human adiponectin, a 30-kDa protein abundantly expressed in adipose tissue (4), exerts local actions on various body tissues, leading to the suppression of hepatic gluconeogenesis and improved insulin sensitivity, thus enhancing whole-body energy homeostasis (5). Adiponectin is recognized as a crucial anti-inflammatory adipokine, and a negative correlation between proinflammatory factors and adiponectin has been established (6). Nevertheless, plasma adiponectin levels decrease in obese individuals, showing a negative correlation with plasma insulin and correlation positive triglycerides, highlighting its importance in the link between insulin resistance and obesity (7). Due to its protective effects, some researchers propose adiponectin as

a potential therapeutic target for obesity, diabetes, and endothelial dysfunction (8).

In contrast to adiponectin, C-reactive protein (CRP) serves as a significant inflammatory marker, with elevated levels observed in overweight and obese individuals. Higher body mass index (BMI) is associated with increased CRP concentrations (9). CRP appears to play a crucial role in various pathological conditions, being significantly associated with elevated insulin levels and insulin resistance. Consequently, **CRP** considered a major risk factor for insulin resistance and diabetes (10). Despite the pathological effects of inflammation, exercise training has been shown to attenuate chronic inflammation (11). The positive effects of exercise training on cardiometabolic diseases, such as diabetes and cardiovascular diseases, are partly attributed to its anti-inflammatory effects Additionally, low-carbohydrate (LC) diets have proven to be effective interventions for treating obesity and type 2 diabetes (13). However, conflicting findings regarding the effects of LC diets on the levels of inflammatory mediators have been reported, with some studies showing no changes (14) and others indicating a decrease (15) in inflammation following LC diets. Furthermore. comparing the effects of LC diets with or without exercise training has demonstrated a greater reduction in percent body fat, triglycerides, glucose, and inflammation, as well as a more significant increase in cardiorespiratory fitness in the LC diet combined with exercise training compared to exercise training alone (16). This emphasizes the synergistic effect of combining an LC diet with exercise training. Nevertheless, limited information exists regarding the combined effects of LC diets and exercise training, especially on inflammatory and anti-inflammatory mediators. Consequently, our study aims to investigate the changes in CRP and adiponectin levels following an eightweek LC diet and functional training intervention in overweight women.

Materials and methods

Study Subjects

In this semi-experimental research, the Iranian (Tehran) overweight women age ranging 25-35 years old recruited, and 48 women randomly selected among eligible women for conducting present interventions.

Study Design

Following the selection of study participants, they were randomly allocated into four equal groups, each consisting of 12 subjects: control (n=12), LC diet (n=12), training (n=12), and training+LC diet (n=12) groups. All subjects voluntarily participated in this study, and various intervention stages were conducted in accordance with the principles of the Declaration of Helsinki. Participants had a sedentary lifestyle in the preceding years, and all were overweight (BMI: 25-29.9 kg/m²). They were in good health and had no cardiometabolic disorders (including type diabetes, hypertension), with no physical or medical limitations preventing the completion of exercise sessions. Overweight women provided informed consent by signing the consent form and willingly accepted all research conditions, including potential benefits and side effects. Subsequently, baseline testing

(including blood sampling, height, and weight measurements) was performed, and after one week, the eight-week interventions (control, training, training+LC diet, LC diet) commenced as agreed upon by the participants.

Functional Training Program

The exercise training conducted in the present study utilized functional circuit (FCT), which emphasized training functional balance lower-body and strength-based exercises. FCT sessions occurred three times a week over a 12week period. Prior to each training session, participants engaged in a 10minute warm-up by walking at their usual pace, and the training session concluded with a 5-minute cool-down involving stretching exercises. Throughout the FCT program, participants in the control and LC diet groups maintained their daily routine lifestyle without incorporating regular physical activity. The details of the FCT training program are provided in a previous research publication (17).

Low Carbohydrate Diet

Participants in the low-carbohydrate (LC) diet and training+LC diet groups received three pre-prepared meals and two interim mid-meal snacks daily throughout the eight-week intervention. Neither diet group was provided with a specific calorie or energy goal. The pre-prepared meals for the LC diet group did not exceed a total of 50 g of carbohydrates (CHO) per day. Daily food intake, including all foods and quantities, was meticulously recorded for both groups, and adherence to the diet was documented daily using a 24-hour food log. The specific LC diet employed

in this study had previously been utilized by Perissiou and colleagues (16).

Biochemical Measurement

Blood sampling was conducted both before and after the completion of the eight-week experiment. A total of 7 ml of blood was drawn from each participant by a laboratory specialist. The collected blood samples were then transferred into a Falcon tube and centrifuged at 3000 rpm for 10 minutes. Subsequently, the serum samples were carefully separated and stored in the freezer for subsequent biochemical measurements.

The levels of components in the lipid profile were determined using a special kit from Pars Azmoon (Iran). Additionally, fasting glucose and C-reactive protein (CRP) levels were measured using Pars Azmoon kits. The serum levels of adiponectin were quantified using an enzyme-linked immunosorbent assay (ELISA) kit from Cusabio (China), with a sensitivity of 1.102 ng/mL and catalog number CSB-E07270h.

Body composition analysis was carried out using the BOCA-X1 analyzer. This comprehensive approach to blood sampling and biochemical analysis, along with body composition assessments, ensured a thorough evaluation of the participants' physiological responses over the course of the eight-week experiment.

Statistical Analyses

The study data were analyzed using SPSS software version 24. Prior to analysis, the normality of data distribution was confirmed through the Shapiro-Wilk test. As the data exhibited a normal

distribution, parametric tests deemed appropriate for the analysis. Between-group differences among the training, control, LC diet, training+LC diet groups were determined **Analysis** of Covariance using (ANCOVA) and Bonferroni post-hoc tests. Additionally, intragroup changes, reflecting the difference between baseline and final analysis, were assessed using paired t-tests. A significance level of p<0.05 was considered for all statistical analyses, ensuring a robust evaluation of the study's findings.

Results

All 48 participants in the present research successfully completed the prescribed interventions, and no subjects were excluded from the study. All participants were included in the statistical analysis. The pre-test and post-test measurements of participants' body weight, BMI, body fat percentage, cholesterol, triglycerides, LDL, HDL, and fasting blood glucose in the different groups are reported as mean±standard deviation in Table 1. This comprehensive inclusion of data allows for a thorough examination of the effects interventions on various physiological parameters across the study groups.

Table 1. The levels of study variables (Mean \pm SD).

Variables	Stage	Control	Training	LC diet	Training+LC Diet	Inter-Group P Value
Glucose (mg/dl)	pre test	87.5±6.54	89.1±10.32	90.9 <u>±</u> 5.74	91.8±5.87	0.976
	post test	88.3±6.27	88.4±4.87	89.6 <u>±</u> 5.14	89.6 <u>±</u> 5.34	
Paired t test		0.999	0.709	0.480	0.011	-
Cholesterol (mg/dl)	pre test	198.4±9.6 5	201.1±9.36	188.1±14.4 8	192.5±9.67	<0.001
	post test	200.1±8.0 3	178.6±9.45	178.3±12.2 5	174.5±10.0 4	
Paired t test		0.316	< 0.001	< 0.001	< 0.001	
Triglyceride (mg/dl)	pre test	176.6±20. 13	179.4±20.6 1	180.3±10.9 3	174.9±9.80	<0.001
	post test	178.1±16. 52	171.4±16.8 4	171.6±7.78	166.6±8.87	
Paired t test		0.555	0.004	< 0.001	< 0.001	-
LDL (mg/dl)	pre test	118.2±10. 61	121.2±7.61	118.7±7.17	118.9±5.40	<0.001
	post test	118.1±9.9 5	115.1±6.99	112.1±6.50	109.1±3.53	
Paired t test		0.960	< 0.001	< 0.001	< 0.001	-
HDL (mg/dl)	pre test	54.1 <u>±</u> 8.39	56.7 <u>±</u> 7.59	57.2 <u>±</u> 6.66	58.8 <u>+</u> 2.12	<0.001
	post test	54.6±6.81	59.7±5.89	61.5±5.55	63.7±3.10	
Paired t test		0.673	0.078	0.001	< 0.001	
Percent Body Fat (%)	pre test	31.7±3.04	33.9 <u>±</u> 3.69	34.6±4.12	33.5 <u>+</u> 3.54	<0.001
	post test	32.3±3.26	30.4±2.87	29.1±3.25	28.2±2.42	
Paired t test		0.47	< 0.001	< 0.001	< 0.001	
BMI (kg.m²)	pre test	28.9±1.04	29.4±0.69	28.9±1.02	28.7±1.10	<0.001
	post test	29.1±1.06	28.6±0.62	27.9±0.96	27.4±0.98	
Paired t test		0.24	0.009	< 0.001	< 0.001	

The levels of cholesterol, triglyceride, LDL, HDL, percent body fat, and BMI exhibited a significant between-group difference as determined by the ANCOVA test (p<0.001). However, there was no significant difference between research groups for glucose levels

(p=0.976). Bonferroni post-hoc test findings indicated a significant decrease in cholesterol levels in the LC diet, training, and training+LC diet groups compared to the control group (p<0.001). Additionally, a significant decrease in cholesterol levels was observed in the

training group compared to the LC diet (p=0.036). Furthermore. group triglyceride and LDL levels significantly decreased in the LC diet, training, and training+LC diet groups compared to the control group (p<0.001). Notably, there was a significant increase in HDL levels in the LC diet (p=0.009) and training+LC diet (p<0.001) groups compared to the control group. Conversely, a significant decrease in percent body fat and BMI was observed in the LC diet, training, and training+LC diet groups compared to the control group (p<0.001) (Table 1).

The ANCOVA test results revealed that the change in adiponectin levels after the eight-week intervention was not significant between the different groups (p=0.135). Additionally, paired t-tests demonstrated no significant changes in adiponectin levels after the eight-week

intervention in the control (p=0.266), LC diet (p=0.358), training (p=1.000), and training+LC diet (p=0.067) groups. The serum levels of adiponectin in the various groups before and after eight weeks are depicted in Figure 1.

Analysis of CRP findings suggested a significant statistically difference between the various groups (p=0.004). The Bonferroni post-hoc test indicated that CRP levels significantly decreased only in the LC diet group compared to the control group (p=0.003). Intragroup analysis revealed a significant decrease in CRP levels only in the LC diet group (p=0.002). However, there was significant change in CRP levels in the control (p=0.339), training (p=0.551), and groups training+LC diet (p=0.256)(Figure 2).

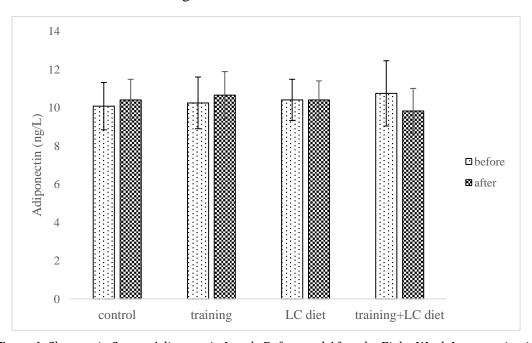


Figure 1. Changes in Serum Adiponectin Levels Before and After the Eight-Week Intervention in Various Groups (Mean±SD).

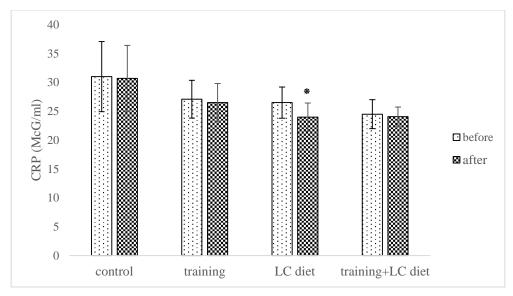


Figure 2. Changes in Serum CRP Levels Before and After the Eight-Week Intervention in Different Groups (Mean±SD). * Indicates a significant decrease compared to the control group.

Discussion

The present study aimed to investigate the effect of eight weeks of functional training and an LC diet on the levels of CRP, adiponectin, and lipid profile in overweight women. The main finding was that functional training alone and with an LC diet had no effect on CRP and adiponectin levels, but the eight-week LC diet led to a significant decrease in CRP levels compared to the control group. Adiponectin, an important adipokine, plays a crucial role in the regulation of glucose and lipid metabolism and insulin sensitivity, with these positive effects attributed to its anti-inflammatory, antifibrotic, and antioxidant properties (18). Various molecular mechanisms, such as upregulation of adenosine the monophosphate-activated protein kinase (AMPK) in the liver and skeletal muscles, seem to be involved in the positive effects of adiponectin (19).

Due to the benefits of adiponectin, it is considered a therapeutic target for obesity and its related disorders (20). Exercise training and nutritional diets are effective interventions in increasing adiponectin levels; however, the present findings showed that functional training alone and in combination with an LC diet did not have a significant effect on adiponectin levels. Exercise training has different effects on the levels of adiponectin, with an increase (21), decrease (22), and no change (23) reported after exercise training. These contradictory findings may be caused by various factors such as the type, intensity, and duration of exerted training programs, as well as the physical characteristics, age, and gender of participants. Ando et al. (2009) confirmed our findings and suggested that 12 weeks of combined training (enduranceresistance) in healthy subjects had no effect on adiponectin levels. concluded that combined training without reducing body weight cannot have a significant effect on adiponectin levels, or its effect is very limited (24).

In another study, researchers observed the positive effects of aerobic training,

specifically improvement in lipid profile, independent of changes in adiponectin levels (25).Despite adiponectin's recognized role in enhancing insulin sensitivity, some studies suggest that exercise-induced reductions in insulin resistance may occur independently of changes in adiponectin levels. instance. Ahmadizad et al. (2007)reported a significant decrease in insulin resistance after 12 weeks of aerobic and resistance training in overweight men, but the change in adiponectin levels was not significant. The researchers concluded that the lack of change in adiponectin levels was likely due to the low intensity of the training program (26).

In line with these findings, Fatouros et al. (2005) demonstrated that the increase in adiponectin levels in overweight and obese elderly populations is dependent on the intensity of resistance training. A sixmonth low-intensity resistance training program (45-50% 1RM) had no effect on adiponectin levels, while moderate (60-65% 1RM) and high-intensity (85-85% 1RM) resistance training was associated with a significant increase in adiponectin levels. Notably, the high-intensity group exhibited a further increase in adiponectin levels. The researchers attributed the upregulation of adiponectin to downregulation of inflammatory mediators (IL-6 and TNF- α) levels (27), although, unfortunately, these circulating levels were not measured in our study.

Another finding of the present study was that CRP levels in the functional training, LC diet, and functional training+LC diet groups decreased by 2.21%, 9.43%, and 8.55%, respectively. However, the reduction in CRP was statistically

significant only in the LC diet group. It is well-documented that CRP, IL-6, and other inflammatory markers increase in obese populations compared to those with normal weight. Conversely, the reduction of visceral adipose tissue has been associated with a decrease inflammatory marker levels (28). Libardi et al. (2012) supported the present findings and suggested that 16 weeks of aerobic, resistance, and combined training do not affect pro-inflammatory cytokines such as CRP, IL-6, and TNF-α, although improvement in lipid profile observed in the different exercise training groups (29).

In contrast to the present findings, it has been reported that eight weeks of aerobic training leads a significant to downregulation of CRP, concluding that the reduction in CRP can independently of changes (decrease) in body weight (30). The researchers attributed this reduction to the possible mechanisms of reducing adipose tissue, given the role of adipose tissue-secreted cytokines (IL-6 and TNF-α) in CRP secretion. Increasing antioxidant capacity and improving endothelial function are also suggested mechanisms for reducing CRP levels with exercise training (30, 31). In the present study, a decrease in CRP levels with functional training was observed. though statistically not significant. Considering that the antiinflammatory effects of exercise training depend on the intensity and duration of exercise sessions (32),insufficient intensity and duration in our training program could be a possible reason for the non-significant changes in CRP levels. However, the exact mechanism needs to be determined in future studies.

The anti-inflammatory effects of exercise training are exerted through various mechanisms. Some studies attribute these effects to the reduction of visceral fat mass, leading to a decrease in adipokine secretion from adipose tissue and the induction an anti-inflammatory of environment (33). Additionally, release of heat shock proteins, changes in the phenotype of immune cells, and the reduction of tissue hypoxia are other crucial mechanisms contributing to the anti-inflammatory effects of exercise training (34).

Another finding of present study was that eight weeks LC diet alone and combined with functional training did not have a significant effect on adiponectin levels. There is insufficient evidence for positive effects of LC diet on the levels of adiponectin and leptin, and researchers suggested that LC diet does not have a significant effect on adiponectin levels (35). Some researchers indicated that despite LC diet role in upregulation of adiponectin, the observed changes wasn't significant statistically (36), and some researchers showed that six weeks LC, low-fat or hypocaloric diets in overweight subjects don't affect adiponectin levels despite weight loss (37). On the other hand, the present findings showed that LC diet cause to a significant decrease in CRP levels, but LC diet with functional training wasn't associated with significant change in the CRP level. Luscombe et al (2005) confirmed our findings and reported that 12 weeks restricted carbohydrate diet led to a significant (24%) reduction in CRP levels in overweight and obese individuals, and researchers considered weight loss as effective mechanisms for CRP reduction

(38). In another study, Sharman et al (2004) confirmed present findings and showed that LC diet in overweight men can exert anti-inflammatory effects and significantly reduce the levels inflammatory cytokines such as CRP, TNF- α and IL-6, They attributed the antiinflammatory properties of LC diet to the reduction of adipose tissue as a main source for inflammatory cytokine, as well as decrease in fasting and post-meal triacylglycerol levels, which these changes in turn is associated with reducing the formation of other lipoproteins such as LDL and VLDL (39). The anti-inflammatory effects ketogenic diet or caloric restriction can be related to the action of β-hydroxybutyrate (βHB) . It has been proven that βHB inhibits histone deacetylase enzymes, free fatty acid receptors and NLRP3 inflammasome, which results in inhibiting inflammatory pathways, oxidative stress and preventing the progression of chronic diseases (40). Cipryan et al (2021) indicated that 12 weeks (three to five sessions per week) high intensity interval training (HIIT) combined with LC diet result significant increase in in adiponectin, significant decrease in leptin levels and increase in the adiponectin/leptin ratio. Moreover, they suggested that LC diet with HIIT has a and synergistic effects further inflammatory markers, and considered LC diet as an effective intervention to prevent chronic diseases (41). The short duration of intervention (eight weeks versus 12 weeks), the different type of exerted exercise training program, and the low volume of training program (low training sessions per week) in the present study compared to the above-mentioned study can be possible reasons for observed contradiction findings regarding changes in the adiponectin level. According to the findings of present study, functional training alone and together with LC diet does not have a significant effect on CRP and adiponectin levels. Nevertheless, functional training, LC diet and their combination led to a significant improvement in lipid profile. It seems that functional training and LC diet alone could not provide the enough stimulation for significant changes in the levels of inflammatory (CRP) and antiinflammatory (adiponectin) adipokines in overweight women, which may be related to insufficient duration or intensity of functional training.

Conclusion

The present findings highlight the positive effects of an LC diet, with or without functional training, in improving lipid profiles. According to the results, it can be inferred that the beneficial outcomes of the interventions applied (LC diet, functional training, or both), including improvements in lipid profiles, body weight loss, and reduced body fat percentage, may occur independently of changes in adiponectin and CRP levels. The exact mechanisms underlying these effects should be investigated in future studies.

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Conflict of interest

The authors declare no conflict of interest.

Authors' contributions

M SH and M GH designed, executed, and monitored the research protocol and interventions. All authors participated in data collection. F GH conducted the statistical analysis. Additionally, the draft and final manuscript were edited and approved by all authors.

References

- Calder PC, Ahluwalia N, Brouns F, Buetler T, Clement K, Cunningham K, et al. Dietary factors and low-grade inflammation in relation to overweight and obesity. Br J Nutr. 2011; 106(S3):S1-S78. doi: 10.1017/S0007114511005460.
- Rodríguez-Hernández H, Simental-Mendía LE, Rodríguez-Ramírez G, Reyes-Romero MA. Obesity and inflammation: epidemiology, risk factors, and markers of inflammation. Int J Endocrinol. 2013; 678159. doi: 10.1155/2013/678159.
- 3. Aguilar-Valles A, Inoue W, Rummel C, Luheshi GN. Obesity, adipokines and neuroinflammation. Neuropharmacology. 2015; 96:124-34. doi: 10.1016/j.neuropharm.2014.12.023.
- 4. Raucci R, Rusolo F, Sharma A, Colonna G, Castello G, Costantini S. Functional and structural features of adipokine family. Cytokine. 2013; 61(1):1-14. doi: 10.1016/j.cyto.2012.08.036.
- 5. Wang ZV, Scherer PE. Adiponectin, the past two decades. J Mol Cell Biol. 2016;8(2):93-100. doi: 10.1093/jmcb/mjw011.
- Fantuzzi G. Adiponectin and inflammation: consensus and controversy. J Allergy Clin Immunol. 2008; 121(2):326-30. doi: 10.1016/j.jaci.2007.10.018.
- 7. Gil-Campos M, Cañete R, Gil A. Adiponectin, the missing link in insulin resistance and obesity. Clin Nutr. 2004; 23(5):963-74. doi: 10.1016/j.clnu.2004.04.010.
- 8. Achari AE, Jain SK. Adiponectin, a therapeutic target for obesity, diabetes, and endothelial dysfunction. Int J Mol Sci. 2017; 18(6):1321. doi: 10.3390/ijms18061321.
- 9. Visser M, Bouter LM, McQuillan GM, Wener MH, Harris TB. Elevated C-reactive protein levels in overweight and obese adults. JAMA. 1999; 282(22):2131-5. doi: 10.1001/jama.282.22.2131.
- Gelaye B, Revilla L, Lopez T, Suarez L, Sanchez SE, Hevner K, et al. Association between insulin resistance and c-reactive protein among Peruvian adults. Diabetol Metab Syndr. 2010;2(1):1-6. doi: 10.1186/1758-5996-2-30.
- 11. Beavers KM, Brinkley TE, Nicklas BJ. Effect of exercise training on chronic inflammation. Clin Chim Acta. 2010; 411(11-12):785-93. doi: 10.1016/j.cca.2010.02.069.
- 12. Pedersen BK. Anti-inflammatory effects of exercise: role in diabetes and cardiovascular disease. Eur J Clin Invest.

- 2017; 47(8):600-11. doi: 10.1111/eci.12781.
- 13. Hall KD, Chung ST. Low-carbohydrate diets for the treatment of obesity and type 2 diabetes. C Curr Opin Clin Nutr Metab Care. 2018; 21(4):308-12. doi: 10.1097/MCO.00000000000000470.
- 14. Cipryan L, Maffetone PB, Plews DJ, Laursen PB. Effects of a four-week very low-carbohydrate high-fat diet on biomarkers of inflammation: non-randomised parallel-group study. Nutr health. 2020; 26(1):35-42. doi: 10.1177/0260106020903206.
- 15. Jonasson L, Guldbrand H, Lundberg AK, Nystrom FH. Advice to follow a low-carbohydrate diet has a favourable impact on low-grade inflammation in type 2 diabetes compared with advice to follow a low-fat diet. Ann Med. 2014; 46(3):182-7. doi: 10.3109/07853890.2014.894286.
- 16. Perissiou M, Borkoles E, Kobayashi K, Polman R. The effect of an 8 week prescribed exercise and low-carbohydrate diet on cardiorespiratory fitness, body composition and cardiometabolic risk factors in obese individuals: a randomised controlled trial. Nutrients. 2020; 12(2):482. doi: 10.3390/nu12020482.
- 17. Giné-Garriga M, Guerra M, Unnithan VB. The effect of functional circuit training on self-reported fear of falling and health status in a group of physically frail older individuals: a randomized controlled trial. Aging Clin Exp Res. 2013; 25(3):329-36. doi: 10.1007/s40520-013-0048-3.
- Nguyen TMD. Adiponectin: role in physiology and pathophysiology. Int J Prev Med. 2020;11: 136. doi: 10.4103/ijpvm.IJPVM_193_20.
- 19. Yamauchi T, Kamon J, Minokoshi Ya, Ito Y, Waki H, Uchida S, et al. Adiponectin stimulates glucose utilization and fattyacid oxidation by activating AMP-activated protein kinase. Nat Med. 2002;8(11):1288-95. doi: 10.1038/nm788.
- 20. Kim JY, Barua S, Jeong YJ, Lee JE. Adiponectin: The potential regulator and therapeutic target of obesity and Alzheimer's disease. Int J Mol Sci. 2020; 21(17):6419. doi: 10.3390/ijms21176419.
- 21. Brooks N, Layne JE, Gordon PL, Roubenoff R, Nelson ME, Castaneda-Sceppa C. Strength training improves muscle quality and insulin sensitivity in Hispanic older adults with type 2 diabetes. Int J Med Sci. 2006; 4(1):19-27. doi: 10.7150/ijms.4.19.
- 22. Van Berendoncks AM, Beckers P, Hoymans VY, Possemiers N, Wuyts FL, Vrints CJ, et al. Exercise training reduces circulating adiponectin levels in patients with chronic

- heart failure. Clin Sci. 2010; 118(4):281-9. doi: 10.1042/CS20090213.
- 23. Park T-G, Hong H-R, Lee J, Kang H-S. Lifestyle plus exercise intervention improves metabolic syndrome markers without change in adiponectin in obese girls. Ann Nutr Metab. 2007; 51(3):197-203. doi: 10.1159/000104137.
- 24. Ando D, Hosaka Y, Suzuki K, Yamagata Z. Effects of exercise training on circulating high molecular weight adiponectin and adiponectin oligomer composition: a randomized controlled trial. J Atheroscler Thromb. 2009; 16(6):733-9. doi: 10.5551/jat.2089.
- 25. Guo W, Kawano H, Piao L, Itoh N, Node K, Sato T. Effects of aerobic exercise on lipid profiles and high molecular weight adiponectin in Japanese workers. Intern Med. 2011; 50(5):389-95. doi: 10.2169/internalmedicine.50.4380.
- Ahmadizad S, Haghighi AH, Hamedinia MR. Effects of resistance versus endurance training on serum adiponectin and insulin resistance index. Eur J Endocrinol. 2007; 157(5):625-31. doi: 10.1530/EJE-07-0223.
- 27. Fatouros I, Tournis S, Leontsini D, Jamurtas A, Sxina M, Thomakos P, et al. Leptin and adiponectin responses in overweight inactive elderly following resistance training and detraining are intensity related. Th J Clin Endocrinol Metab. 2005; 90(11):5970-7. doi: 10.1210/jc.2005-0261.
- 28. Selvin E, Paynter NP, Erlinger TP. The effect of weight loss on C-reactive protein: a systematic review. Arch Intern Med. 2007; 167(1):31-9. doi: 10.1001/archinte.167.1.31.
- 29. Libardi CA, De Souza GV, Cavaglieri CR, Madruga VA, Chacon-Mikahil M. Effect of resistance, endurance, and concurrent training on TNF-a, IL-6, and CRP. Med Sci Sports Exerc. 2012;44(5):50-6. doi: 10.1249/MSS.0b013e318229d2e9.
- 30. Okita K, Nishijima H, Murakami T, Nagai T, Morita N, Yonezawa K, et al. Can exercise training with weight loss lower serum C-reactive protein levels? Arterioscler Thromb Vasc Biol. 2004; 24(10):1868-73. doi:
 - 10.1161/01.ATV.0000140199.14930.32.
- 31. Bastard J-P, Jardel C, Bruckert E, Blondy P, Capeau J, Laville M, et al. Elevated levels of interleukin 6 are reduced in serum and subcutaneous adipose tissue of obese women after weight loss. J Clin Endocrinol Metab. 2000; 85(9):3338-42. doi: 10.1210/jcem.85.9.6839.

- 32. Petersen AMW, Pedersen BK. The antiinflammatory effect of exercise. J Appl Physiol. 2005; 98(4):1154-62. doi: 10.1152/japplphysiol.00164.2004.
- 33. Gleeson M, Bishop NC, Stensel DJ, Lindley MR, Mastana SS, Nimmo MA. The anti-inflammatory effects of exercise: mechanisms and implications for the prevention and treatment of disease. Nat Rev Immunol. 2011; 11(9):607-15. doi: 10.1038/nri3041.
- 34. Flynn MG, McFarlin BK, Markofski MM. State of the art reviews: The anti-inflammatory actions of exercise training. Am J Lifestyle Med. 2007; 1(3):220-35. doi: 10.1177/1559827607300283.
- 35. Morshedzadeh N, Ahmadi AR, Tahmasebi R, Tavasolian R, Heshmati J, Rahimlou M. Impact of low-carbohydrate diet on serum levels of leptin and adiponectin levels: a systematic review and meta-analysis in adult. J Diabetes Metab Disord. 2022: 21(1):979-990. doi: 10.1007/s40200-021-00952-7.
- Shemirani F, Golzarand M, Salari-Moghaddam A, Mahmoudi M. Effect of low-carbohydrate diet on adiponectin level in adults: a systematic review and doseresponse meta-analysis of randomized controlled trials. Crit Rev Food Sci Nutr. 2022; 62(14):3969-78. doi: 10.1080/10408398.2021.1871588.
- 37. Youssef M. The effect of dietary intervention by low carbohydrate diet, and low fat diet, on weight loss, leptin and adiponectin. Life Sci J. 2015;12:33-42. doi: 10.7537/marslsj120415.05.
- 38. 38. Luscombe-Marsh ND, Noakes M, Wittert GA, Keogh JB, Foster P, Clifton PM. Carbohydrate-restricted diets high in either monounsaturated fat or protein are equally effective at promoting fat loss and improving blood lipids. Am J Clin Nutr. 2005; 81(4):762-72. doi: 10.1093/ajcn/81.4.762.
- 39. Sharman MJ, Volek JS. Weight loss leads to reductions in inflammatory biomarkers after a very-low-carbohydrate diet and a low-fat diet in overweight men. Clin Sci. 2004; 107(4):365-9. doi: 10.1042/CS20040111.
- 40. Møller N. Ketone body, 3-hydroxybutyrate: minor metabolite-major medical manifestations. J Clin Endocrinol Metab. 2020; 105(9):2884-92. doi: 10.1210/clinem/dgaa370.
- 41. Cipryan L, Dostal T, Plews DJ, Hofmann P, Laursen PB. Adiponectin/leptin ratio increases after a 12-week very lowcarbohydrate, high-fat diet, and exercise

training in healthy individuals: A non-randomized, parallel design study. Nutr Res. 2021;87:22-30. doi: 10.1016/j.nutres.2020.12.012.