

# Impact of physical activity levels on cardiometabolic parameters and autonomic modulation in elderly: a cross-sectional study

**Ariane Viana**

Universidade Nove de Julho - Campus Vergueiro

**Neide Bugliani**

Universidade Sao Judas Tadeu

**Michelle Sartori**

Universidade Sao Judas Tadeu

**Camila Martins da Silva**

Universidade Sao Judas Tadeu

**Iris Callado Sanches**

Universidade Sao Judas Tadeu

**Marília Velardi**

Universidade Sao Judas Tadeu

**Maria Luiza de Jesus Miranda**

Universidade Sao Judas Tadeu

**Katia De Angelis** (✉ [prof.kangelis@yahoo.com.br](mailto:prof.kangelis@yahoo.com.br))

Universidade Federal de São Paulo, UNIFESP <https://orcid.org/0000-0002-3640-9049>

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## Research article

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# Abstract

**Background:** By 2030, approximately 20% of the population worldwide will be 65 years of age or older. In this age group, cardiovascular diseases (CVD) will result in 40% of all deaths. Heart rate variability (HRV) appears to be reduced under stress conditions and in many chronic diseases as cardiovascular diseases that appear to be associated with autonomic changes that usually include decreased parasympathetic activation and/or increased sympathetic modulation. Thus, aging is very important for altering neurohumoral mechanisms that control the cardiovascular system.

**Objective:** To analyze the influence of level of physical activity in the HRV in elderly people.

**Methods:** Cross-sectional analytical study which included 49 subjects (60 – 86 years), which answered the Baecke questionnaire.

**Results:** The less physically active group had lower levels of plasma HDL ( $42 \pm 2.43 \text{mg/dL}$  vs  $52 \pm 3.8 \text{mg/dL}$ ) and higher systolic BP ( $131 \pm 117 \text{mmHg}$  vs  $117 \pm 4.07 \text{mmHg}$ ) when compared to the most active group. The more physically active group showed higher PI of variance ( $1849.6 \pm 387.7 \text{ms}^2$ ) when compared to the moderate ( $892.1 \pm 141.9 \text{ms}^2$ ) and the less physically active groups ( $972.3 \pm 285.2 \text{ms}^2$ ). Moreover, the most physically active group presented lower LF/HF balance ( $1.6 \pm 0.2$ ) when compared to the moderate ( $2.6 \pm 0.3$ ) and the less active groups ( $2.5 \pm 0.2$ ).

**Discussion:** The results demonstrated the importance of physical activity to promote hemodynamic, autonomic and lipid profile improvement, suggesting a lower risk of developing or progressing the disease in the elderly.

## 1. Introduction

Aging can lead to a gradual loss of physiological reserves, increasing the risk of contracting various diseases and a general decline in the individual's intrinsic capacity (1).

By 2030, approximately 20% of the population worldwide will be 65 years of age or older. In this age group, cardiovascular diseases (CVD) will result in 40% of all deaths (2). CVD accounts for 29.6% of deaths worldwide (3), and for 30% of deaths in Brazil (4). Data show that 40.6% of the deaths were caused by high blood pressure, 13.7% because of smoking, 13.2% related to poor diet, 11.9% as a consequence of sedentary lifestyle and 8.8% due to changes in the blood glucose level (5).

Heart rate variability (HRV) appears to be reduced under stress conditions and in many chronic diseases (6) as cardiovascular diseases that appear to be associated with autonomic changes that usually include decreased parasympathetic activation and/or increased sympathetic modulation. Thus, aging is very important for altering neurohumoral mechanisms that control the cardiovascular system (7). Although studies on the effects of exercise are still conflicting, there is evidence that aerobic training may increase HRV and cardiac vagal tone both in healthy individuals and in patients with chronic diseases (6,7,8).

Considering the population's aging increase and the consequent increase in cardiovascular risk, in addition to the increasingly sedentary changes in the lifestyle of the population, it is necessary to study the influence of the level of physical activity on the aging process and factors that influence the development of cardiovascular diseases. Thus, in this study, we tested the hypothesis that the level of physical activity in the elderly is associated with different values of HRV. In this sense, our objective was to analyze the influence of the level of physical activity on the modulation of the autonomic nervous system and on cardiometabolic variables in the elderly.

## **2. Materials And Methods**

This is a cross-sectional analytical study. The subjects of this study were individuals participating in a program called "Project Senior with Active Life of São Judas Tadeu University". The Project was submitted to the Research Ethics Committee (COEP/USJT) under number 0067.0.219.000-09. All the participants signed Informed Consent Form.

The sample was composed by 49 individuals, 9 men and 40 women that attended the inclusion criteria: to pertain to the lower socio-economic status (earning till 5 minimum wages per month); to belong to levels 3 and 4 of functional status: independent and inactive (level 3) and active (level 4); being between 60 and 86 years old and with read and write skills.

### **2.1 Habitual Physical Activity Questionnaire (Baecke Questionnaire) and Anamnese**

To measure the level of physical activity, the individuals answered the Baecke questionnaire. This questionnaire is composed by 16 questions and measures the habitual physical activity in the last 12 months (Alex Antonio Florindo & Latorre, 2003; (10). The anamnese questionnaire, composed by 12 questions, measured the lifestyle, medical history and physical activity throughout life.

### **2.2 Anthropometric data**

The body mass index (BMI= the weight divided by height squared) was calculated to verify if the individuals were overweight or obese.

### **2.3 Blood arterial pressure**

The individual remained at rest for 15 minutes in a closed room, sitting in a comfortable position, with the legs uncrossed and feet flat. After this period, the blood pressure was measured 3 times at intervals of 02 minutes. The method used was the auscultation following the recommendations of the Brazilian Society of Hypertension and Guideline to American College of Cardiology (2017), using stethoscope and sphygmomanometer of Litman® (4, 11).

### **2.4 Biochemical Analysis**

The participants attended other day in the laboratory of chemical analyzes at University São Judas Tadeu to collect blood samples, state fasting 12 hours. Tubes were collected for biochemical analyzes (blood without anticoagulant) and determination of blood glucose level (fluoride anticoagulant). The blood without anticoagulant was centrifuged and from the serum were analyzed: LDL, HDL, VLDL and triglycerides.

## 2.5 Heart Rate Variability

The HRV was measured by the record of the R-R interval (ms), through the Polar frequency meter, model RS800®. In that HR monitor, the transmitter belt detects the electrocardiographic signal beat-by-beat and transmits it through an electromagnetic wave to the Polar wrist receptor, where this information is digitalized, displayed and stored. This system detects the ventricular depolarization corresponding to the R wave of the electrocardiogram, with sampling frequency of 500 Hz and time resolution of 1 ms which was previously validated against standard electrocardiograph by Holter (12). The recording files were transferred to the Polar Precision Performance Software through the Interface Infrared, or IrDA, which allows the exercise data bidirectional exchange with a microcomputer for subsequent analysis of the interval variability of cardiac pulse in the different recorded situations. The R-R intervals (PI) originated from the frequency meter were converted and stored in an Excel file and it was visually verified to identify and/or correct some incorrect marking. The data were analyzed and charted through the MATLAB program in the format of the Fast Fourier Transform (FFT). After this mathematical remodeling, the absolute potencies were obtained in the respective pre-set frequency bands: low frequency (LF, 0.04-0.15 Hz) and high frequency (HF, 0.15-0.4 Hz). The data are expressed in normalized units. The LF component is used as an index of sympathetic modulation. The HF component is used as an index of the parasympathetic modulation. The LF/HF ratio indicates the sympathovagal balance. The detection of the R-R intervals obtained in the frequency meter will follow the same criteria previously described for the design of the time sets of the variability in the frequency domain (13). For this study the total variance was used as an index in the time domain.

## 2.6 Statistical analysis

The InStat® software (Version 3.0) was used to analyze parametric data. The results are presented in mean  $\pm$  standard error of the mean. ANOVA followed by Newman Keuls Student's post hoc was used for statistical analysis. The significance level adopted was 5%. The power of the sample was calculated *a posteriori* considering the variances of the groups for the Variance of IP obtaining a  $\beta$  of 1.0.

## 3. Results

The sample consisted for 49 individuals, with 9 men and 40 women, aging  $70 \pm 9.97$  years. The BMI was  $27.6 \pm 3.95$  Kg/m<sup>2</sup> (Table 1). There was no difference in BMI considering the different levels of physical activity (Table 1). Between the individuals analyzed, we noted that 3.43% (7) were taking medication for dyslipidemia, 1.96% (4) for diabetes and 9.31% (19) for hypertension.

**Table 1.** Demographics and clinical characteristics.

	<b>Active (n=13)</b>	<b>Moderate Active (n=19)</b>	<b>Less Active (n=17)</b>
<b>Age (years)</b>	70.83 ± 2.28	69.68 ± 1.67	69.18 ± 1.60
<b>BMI (Kg/m<sup>2</sup>)</b>	27.11 ± 1.26	28.13 ± 1.21	28.40 ± 1.13
<b>Fasting blood glucose (mg/dL)</b>	100 ± 4.33	104 ± 4.76	101 ± 5.55
<b>Total cholesterol (mg/dL)</b>	179 ± 1.75	190 ± 12.12	186 ± 9.80
<b>HDL (mg/dL)</b>	52 ± 3.08	45 ± 2.40	42 ± 2.43*
<b>LDL (mg/dL)</b>	105 ± 3.91	118 ± 10.87	118 ± 9.11
<b>VLDL (mg/dL)</b>	24 ± 2.91	26 ± 2.55	25 ± 3.18
<b>Triglycerides (mg/dL)</b>	105 ± 4.58	144 ± 11.68*	143 ± 10.96*
<b>Systolic BP (mmHg)</b>	117 ± 4.07	125 ± 2.68	131 ± 3.48*
<b>Diastolic BP (mmHg)</b>	72 ± 2.00	78 ± 1.16*	79 ± 1.65*
<b>HR (bpm)</b>	71 ± 3.93	68 ± 1.82	75 ± 3.02

\* p < 0.05 vs. active group; BMI: Body Mass Index; HDL: High-density lipoprotein; LDL: Low-density lipoprotein; VLDL: Very Low Density Lipoprotein; BP: Blood Pressure; HR: Heart rate. Data are represented as mean +/- SEM.

We observed a significant difference in energy expenditure between the active group (9.25 ± 0.18 MET) versus the moderate active group (7.84 ± 0.08 MET) and least active group (6.33 ± 0.34 MET) (Figure 1-A).

There was no significant difference between the groups regarding the fasting blood level of fasting blood glucose, cholesterol, low density lipoprotein (LDL) and very low-density lipoprotein (VLDL) (Table 1). However, we found that the less active group had lower levels of high-density lipoproteins (HDL) (42 ± 2.43 mg/dL) compared to the active group (52 ± 3.08 mg/dL) (Figure 1-D). In addition, the moderate active and the less active groups had higher values of blood triglycerides (144 ± 11.68 mg/dL and 143 ± 10.96 mg/dL respectively) when compared to the active group (105 ± 4.58 mg/dL) (Table 1. Figure 1-C).

The systolic blood pressure was higher in the less active group (131 ± 3.5 mmHg) compared to the active group (117 ± 4.1 mmHg). The same was observed for diastolic blood pressure (DBP). We noticed a reduction in DBP in the active group (72 ± 3.93 mmHg) in comparison to the moderate active (78 ± 1.16

mmHg) and the less active groups ( $79 \pm 1.65$  mmHg) (Figure 1-B). There was no significant difference between groups with respect to heart rate (Table 1).

Regarding HRV, the active group showed an increase in the variance of RR interval ( $1849.6 \pm 387.7$  ms<sup>2</sup>) as compared to the moderate active ( $892.1 \pm 141.9$  ms<sup>2</sup>) and to the less active groups ( $972.3 \pm 285.2$  ms<sup>2</sup>) (Figure 2-A). Moreover, the active group showed higher absolute HF component (ms<sup>2</sup>) as compared to the other groups (Table 2). The sympathetic-vagal balance was higher in the moderate active ( $2.6 \pm 0.3$ ) and the less active groups ( $2.5 \pm 0.2$ ) as compared to the active group ( $1.6 \pm 0.2$ ) (Figure 2-B).

**Table 2.** The Heart Rate Variability in the frequency domain.

	<b>Active (n=13)</b>	<b>Moderate Active (n=19)</b>	<b>Less Active (n=17)</b>
<b>LF (ms<sup>2</sup>)</b>	$331.8 \pm 102.3$	$182.5 \pm 40.3$	$158.1 \pm 41.4$
<b>HF (ms<sup>2</sup>)</b>	$298.9 \pm 80.3$	$78.5 \pm 20.8^*$	$63.2 \pm 16.2^*$
<b>%LF</b>	$57.7 \pm 3.5$	$68.6 \pm 2.4^*$	$69.3 \pm 2.2^*$
<b>%HF</b>	$42.3 \pm 3.5$	$31.4 \pm 2.4^*$	$30.7 \pm 2.2^*$

\*  $p < 0.05$  vs. active group; LF: Low Frequency band of RR; HF: High Frequency band of RR. Data are represented as mean +/- SEM.

## 4. Discussion

Considering the increase in the age of the population and consequent increase in cardiovascular risk, in addition to the increasingly sedentary changes in the lifestyle of the population, our findings showed that the less active group had higher SBP and DBP values compared to the active group. The most active group showed higher VAR RR values and the IP HF band in relation to the less active group, in addition to a lower sympathovagal balance, indicating lower cardiac sympathetic modulation.

The habitual physical activity questionnaire (PAQ), used in this study, allowed us to verify the energy expenditure dispensed in daily activities during the 12-month period. Through this questionnaire, we divided the participants of this study into three groups to analyze the influence of physical activity on quality of life and health of people over 60 years of age (9). The least active group presented the lowest value of energy expenditure, being considered as a sedentary group. It is worth noting that in the Western world, sedentary lifestyle, as a risk factor for health, is only surpassed by tobacco consumption and is highly representative (14, 15). The global challenge of physical inactivity is further amplified by the risk it carries. We have observed in some studies enough evidence that 6-10% of all deaths from noncommunicable diseases worldwide can be attributed to physical inactivity, and this percentage is even higher for specific diseases (eg. 30% for the ischemic heart disease). In 2007, 5.3 to 5.7 million of

deaths worldwide from noncommunicable diseases could have been avoided if, in theory, people who were inactive were sufficiently active (16).

In addition, some serious health problems are associated with obesity, including type II Diabetes and cardiovascular disease. White adipose tissue is not only a place for energy storage, but it is also an active endocrine organ that secretes more than 50 cytokines/chemokines and bioactive mediators called adipocytokines that are involved in lipid metabolism, insulin sensitivity, immunity, angiogenesis and inflammation (17).

According to Santanasto (2017) with aging there is a reduction in muscle mass and an increase in the accumulation of body fat, which may be related to food and cultural habits (18). The least active group presented higher values of SBP and DBP compared to the active group. In this sense, it is worth remembering that CVD mortality increases considerably with elevated blood pressure.

Regarding the results obtained from the lipid profile, the less active group presented a high level of triglycerides and a reduction in the level of HDL, if changes in LDL and VLDL in relation to the other groups. Velloso et al. (2013) observed that the decrease in the plasma concentration of HDL is strongly related to the development of endothelial dysfunction and, consequently, to CAD (19).

Furthermore, some studies have shown that heart rate variability (HRV) is reduced in stress conditions and in many chronic diseases, and may even predict the development and prognosis of some diseases (6). In a recent study of our group, we showed that sedentary women presented higher heart rate, lower variance of RR interval and RMSSD and higher cardiac sympathovagal balance (LF/HF) both at rest and in response to the mental stress test in comparison to physically active women (20). In the HRV spectral analysis, the less active group showed lower IP VAR and IP HF band in relation to the active group, besides a higher sympathovagal balance, indicating a higher cardiac sympathetic modulation.

According to our findings, some authors mention that there are losses in HRV with aging and a less active life (21, 22). It is important to emphasize that the reduction in HRV is related to a decrease in vagal activity over the sinus node and may be associated with other risk factors and increased sympathetic activity related to high plasma noradrenaline concentration in individuals over 65 years (23).

Studies show that higher HRV values indicate better adaptation and control of the sympathetic and parasympathetic nervous system, and active individuals have a better index in the time and frequency domain due to the increase in vagal tone caused by the practice of physical activity (24). Recent study showed that sedentary offspring of hypertensives, in contrast with the strength-trained group, presented impairment of total variance of RR interval, as well as an increase in cardiac sympathovagal balance (25).

Our results corroborate these findings, since the active group showed higher VAR RR and AF component ( $m^2$ ), as well as a better sympathovagal balance. In addition, studies have demonstrated that cardiovascular autonomic dysfunction, evidenced by a reduction in HRV, is associated with higher levels

of inflammatory mediators such as TNF- $\alpha$  and IL-6 (26). In addition, a study of our group showed that cardiovascular autonomic dysfunction precedes metabolic dysfunction in male mice submitted to chronic consumption of fructose (27). Finally, studies show that the vagus nerve can modulate the inflammatory response and oxidative stress in some pathophysiological situations (28, 29).

In this sense, the results of the present study suggest that the best cardiac autonomic modulation observed in the active elderly group may be associated with the reduction of inflammatory mediators and oxidative stress, which together could be related to the reduction of cardiovascular and metabolic risk in this phase of life.

In fact, regardless of age, the practice of physical activity seems to be beneficial and helps increase the standard curve of physiological aging (18). In addition, changes in lifestyle are recommended in the primary prevention of hypertension, and in individuals with borderline BP. These changes may reduce BP as well as cardiovascular mortality. The main non-medicinal recommendations for the primary prevention of hypertension are: healthy eating, controlled consumption of sodium and alcohol, potassium intake, sedentary lifestyle and smoking (11).

## 5. Conclusion

Our findings highlight the importance of a physically active life to prevent cardiometabolic dysfunctions and for a healthier aging, enabling improvements in quality of life and fewer comorbidities associated with ageing. However, more studies are needed to consolidate our findings, which indicate a major influence of physical activity level on increased cardiovascular risk, assessed by classic markers such as BMI, lipid profile and BP, but also by heart rate variability.

## 6. Abbreviations

**HRV**- Heart Rate Variability

**BP**- Blood Pressure

**CVD**- Cardiovascular Disease

**BMI**- Body Mass Index

**MET** – Energy Expenditure

**LDL**- Low Density Lipoproteins

**VLDL**- Very Low Density Lipoproteins

**HDL**- High Density Lipoproteins

**DBP**- Diastolic Blood Pressure

**LF-** Low Frequency band of RR

**HF-** High Frequency band of RR

**SBP-** Systolic Blood Pressure

**VAR RR-** Variance of RR interval

**IP HF-** Pulse Interval of High Frequency

**PAQ-** Habitual Physical Activity Questionnaire

**CAD-** Coronary Artery Disease

**RMSSD-** Square Root Of The Mean Squared Differences Of Successive N-N Intervals

**LF/HF-** Sympathovagal Balance

**TNF- $\alpha$ -** Tumor Necrosis Factor

**IL-6-** Interleukin

**HR-** Heart Rate

**FFT-** Fast Fourier Transform

## 7. Declarations

**Ethics approval and consent to participate:** The study participants were informed about the procedures and signed a free and informed consent form. The Project was submitted to the Research Ethics Committee (COEP/USJT) under number 0067.0.219.000-09.

**Consent for publication:** “Not applicable”

**Availability of data and materials:** The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

**Conflict of Interest:** The authors declare that the results of the study are presented clearly, honestly, and without fabrication, falsification, or inappropriate data manipulation. There is no conflict of interest between authors.

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**Author Contributions:** AV, NB, MS, CMS, ICS, MV, MLJM and KA contributed conception and design of the study; AV, MS, NB and CMS organized the database; AV, MV, MLJM and KA performed the statistical

analysis; AV wrote the first draft of the manuscript; MS, MV, MLJM and KA wrote sections of the manuscript. All authors contributed to manuscript revision, read and approved the submitted version.

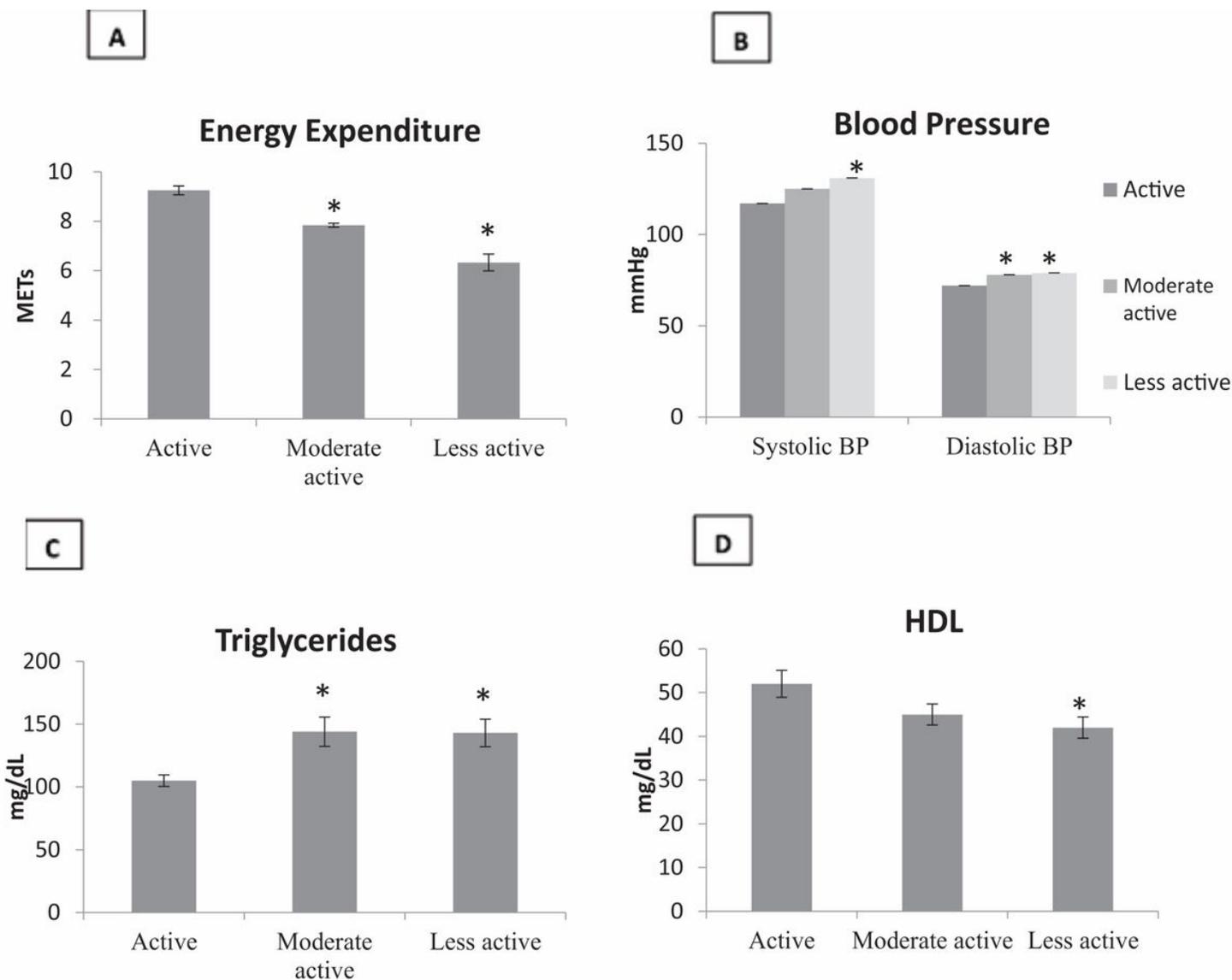
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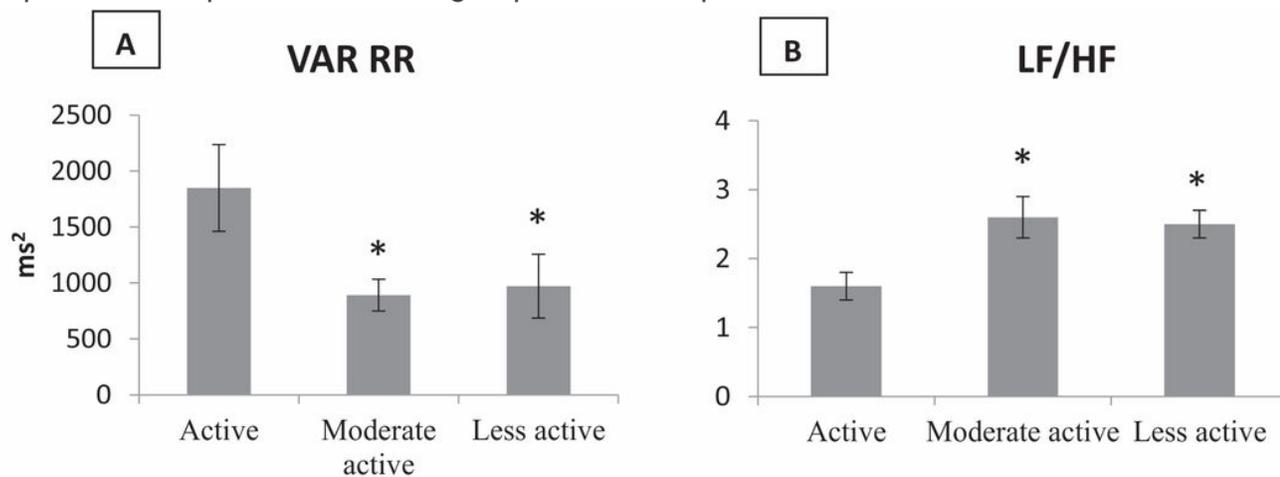
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## Figures



**Figure 1**

Energy Expenditure (A), Blood Pressure (B), Triglycerides (C) and High-density Lipoprotein (D) in the groups studied. \*  $p < 0.05$  vs. active group. Data are represented as mean  $\pm$  SEM.



## Figure 2

Variance of RR (VAR RR) (A) and sympathetic-vagal balance (LF/HF) (B) in the groups studied. \*  $p < 0.05$  vs. active group. Data are represented as mean  $\pm$  SEM.