Influence of Football on Physiological Cardiac Indexes in Professional and Young Athletes

Cristian V. Francavilla†, Francesco Sessa†, Monica Salerno², Giuseppe D. Albano², Ines Villano³, Giovanni Messina², Fabio Triolo⁴, Lorenzo Todaro⁵, Maria Ruberto⁶, Gabriella Marsala⁷, Orazio Cascio⁸, Maria P. Mollica³, Vincenzo Monda³, Giuseppe Cibelli², Anna Valenzano³, Christian Zammit⁹, Marcellino Monda³ and Antonietta Messina³*

Background: After long-term intensive training, considerable morphological and functional heart changes occur in professional athletes. Such changes arise progressively and regress upon interruption of the physical activity. Morphological and functional alterations on heart are known as “Athlete’s heart” condition.

Objective: This study aims to compare echocardiographic parameters in two different groups of professional athletes. Furthermore, a prospective study is performed analyzing the echocardiographic changes occurring in 12 professional players in 3 years of follow-up.

Materials and Methods: 78 football players were examined from July 2011 to May 2016 (40 enrolled in Group A and 38 in Group B). Twelve players of GROUP A were followed for 3 consecutive seasons. The general clinical examination, the cardiopulmonary evaluation, the ECG, the ergometer stress test, the spirometric examination and the standard cardiac echo color doppler test were recorded.

Results: Left ventricle dimensions, left atrium dimensions, and interventricular septum dimensions were higher in A players than in B players. Moreover, following up 12 players for 3 years, a statistically significant increase of such values was observed.

Discussion: In A players, higher dimensions of the left chambers and the interventricular septum were observed, compared to B players. No statistically significant difference was found regarding the ejection fraction. The 3 years follow-up showed a statistically significant increase of both left chambers and interventricular septum dimensions, particularly in the second and third year.

Conclusions: These findings demonstrated that A players have higher echocardiographic parameters respect to B players. The results of this study support the scientific theory that long-term intensive training influences heart function, inducing “athlete’s heart” with morphological adaptations. No significant echocardiographic
variation within the examined sample was observed for different roles (goalkeeper, defender, midfielder, or attacker) or skills of individual players.

Keywords: professional football player, echocardiographic parameters, heart adaptation to exercise, left ventricle dimensions, athlete’s heart

INTRODUCTION

Physical activity is an essential tool in order to prevent cardiovascular, musculoskeletal and metabolic diseases. Sedentary lifestyle is still widespread among both youths and adults, especially in the western countries (Francavilla et al., 2007).

Long-term intensive training program leads to several morphological and functional cardiovascular changes that occur progressively, and regress upon interruption of the physical activity (Atchley and Douglas, 2007; Zeppilli, 2008; D’Andrea et al., 2009; Neri et al., 2009a).

“Athlete’s heart” is defined as the adaptation of the heart to long-term intensive physical activity, and is characterized by symmetric and harmonic increase of intracavitary diameters and wall thickness (Fagard, 2003; Turillazzi et al., 2010).

Such adaptation is strictly dependent on a wide spectrum of factors: (1) genotype (hereditary factors) (Sessa et al., 2011); (2) age of onset of physical activity; (3) type and intensity of training program (Lavie et al., 2001).

Two different intensive physical activity models were initially described, both related to different ventricular hypertrophy models.

Resistance training is characterized by isotonic-dynamic muscular activity and requires aerobic energy expenditure. It gradually leads to a decrease in both systemic arterial resistance and venous return, and a telediastolic left ventricle volume increase, leading to an increase in systolic output. Resistance athletes (swimming, long distance running, cycling etc…) show an increase in all the intracavitary dimensions, and also in wall thickness. These features lead to eccentric hypertrophy due to the volume overload, with a proportional increase in myocardial mass (Neri et al., 2009b; Lamotte et al., 2010; Forlano et al., 2011).

Power training is characterized by static muscular activity and requires anaerobic energy expenditure. It leads to a consistent increase in myocardial mass, due to the increase in wall thickness. The increase in intracavitary diameters is insignificant, and therefore results in a concentric ventricular hypertrophy. These changes are frequently found in athletes practicing weightlifting, short distance running, etc….

Such adaptations are involved in isometric muscular work. They are characterized by a pressure response to prolonged muscular tensions, with no increase in regional perfusion. The increase in peripheral resistance leads to a substantial raise in left ventricular afterload during exercise, resulting in left ventricle pressure overload (Messina et al., 2016; Chieffi et al., 2017; Maté-Muñoz et al., 2017).

The above-mentioned model, characterized by two different training mechanisms, is no longer reputed accurate. Recent studies suggest a unique heart adaptation model, characterized by a ventricular hypertrophy correlated to body size and exercise intensity, not to the nature of the physical activity (Pelliccia et al., 1991, 1999; Spirito et al., 1994; Avola et al., 2004; Naylor et al., 2008; Zeppilli et al., 2008; Cianci et al., 2016).

The physiological demands of soccer are complex and they are related to the nature of the exercise pattern (Morgans et al., 2014). According to the Mitchell et al. classification (Mitchell et al., 2005), football players usually perform a physical activity that includes both a dynamic and a static component. The requirement for frequent changes in both the speed of movement (e.g., walking, jogging, high intensity running, and sprinting) and direction, makes the activity profile intermittent. In fact, football, like most other sports, is characterized by both aerobic and anaerobic energy expenditure, with an overload in both left ventricular volume and pressure. In athletes who practice this sport, there is an increase in both intracavitary diameters and walls thickness. This results in an intermediate ventricular hypertrophy model, between resistance exercise and power exercise (Bangsbo et al., 2006; Bramanti et al., 2012; Mielgo-Ayuso et al., 2015; Precenzano et al., 2016a; Zurutuza et al., 2017).

After a long-term intensive exercise, professional athletes undergo substantial morphological and functional cardiovascular changes, which arise progressively and regress upon reduction of physical activity. During the last decades, sports culture has changed consistently. Professional athletes follow rigorous training programs with daily practice sessions, leading to a wide range of cardiovascular and musculoskeletal adaptation mechanisms. These changes are related to other factors, such as genotype, age, and age onset of physical activity.

One of the most important adaptations is named “Athlete's heart,” involving hearth changes. Several studies showed how correct interpretation of electrocardiography could lead to a correct interpretation of the features present in the “Athlete's heart” (Braschi et al., 2009, 2011, 2012; Neri et al., 2013; Vyshta and Vacchiano, 2014; Francavilla et al., 2016).

This study aims to compare echocardiographic parameters in two different groups of professional athletes. Furthermore, a prospective study is performed analyzing the echocardiographic changes occurring in 12 professional players in 3 years of follow-up.

MATERIALS AND METHODS

Seventy eight professional football players were enrolled subdivided into two groups. Group A is composed of 40 “Serie A” players (“Serie A” is a professional league competition for football clubs located at the top of the Italian football league system); Group B is constituted by 38 “Primavera” players of the respective team (“Primavera” is named the Italian youth league, under 19 y.o. football players). All evaluations were recorded between
July 2011 and May 2016. The principal biological parameters for each group are summarized in Table 1. All athletes were evaluated at the beginning of the regular season; 12 players of GROUP A were followed for 3 consecutive seasons. No lifestyle changes were reported. A complete cardiovascular examination was performed on all athletes. The general clinical examination, the cardiopulmonary evaluation, the ECG, the ergometer stress test, the spirometric examination and the standard cardiac echo doppler test were recorded.

All subjects gave written informed consent in accordance with the latest version of the declaration of Helsinki. The protocol was approved by the Human Ethical Review Committee (University of Palermo).

Training Program in Professional Football Players
All athletes were trained following the typical training protocol for each season. A regular season consists of 11 working months and is preceded by 3 weeks of intensive training characterized by training twice daily. In a typical training week, the players undergo six practice sessions and 1 official game, which usually take place on Sunday. Each player is trained for a total amount of 12–14 h of physical activity per week. Most of the physical activity is performed on the football field, with a once weekly training session taking place in the gym.

Body Composition and Anthropometric Parameters
For the body weight measurement the Seca 877 Mechanical Scale (Hamburg, Germany) weight scale was used. Body length measurement was performed using SECA 240. Bioimpedance analysis was performed using BIA 101 ASE (Akern) instrumentation. Resistance and reactance were measured in standard condition, at rest and 12 h after the last training session. The subcutaneous tissue thickness was also measured using the Wither formula to estimate the fat mass percentage.

Echo Color Doppler Evaluation
The echocardiographic evaluation was conducted by an experienced cardiologist using a Philips Ie33 device, with an X-Matrix 2.5 MHz probe. The exam was always conducted by the same operator to eliminate inter-operator variability. In accordance with the American Society of Echocardiography guidelines, both systolic and ventricular diastolic parameters were measured at rest.

Echocardiographic evaluation included measuring of wall thickness and intracavitary diastolic and telediastolic diameters. Cardiac chambers dimensions were calculated in ml/mq. These data were differentiated according to the training mechanism involved.

Both systolic and diastolic interventricular septum thickness, posterior wall thickness, left ventricle diameter, aortic root diameter and left atrium dimensions were calculated. The Devereux formula was used in order to calculate left ventricular mass index (g/m²) (Ganau et al., 1992). The Simpson formula was used in order to calculate ejection fraction. Diastolic function was evaluated by calculating E and A waves and deceleration time (TD). In addition, by using tissue doppler, E', A', S', and E/E' ratio were calculated, to estimate left ventricular filling pressures.

Statistic Analysis
For each of the variables related to the analysis, the mean, standard deviation, median, minimum and maximum, were calculated. The left ventricle size, the left atrium, the interventricular septum, and the ejection fraction were compared between Group A and B using the T Student test.

A team variables comparison was performed by using the ANOVA model. If requested, Tukey post-hoc test was calculated for multiple comparing.

RESULTS
All data were expressed as mean and standard deviation values (SD). The assessment of body composition showed that the Group A weighed more than the B group, with a higher active cellular mass and a lower fat mass percentage. Group A players also had a better hydration state. The principal parameters for each group were summarized in Table 1.

In the examined cohort, the diastolic left ventricle size was between 40 and 61 mm, and the left atrium volume was between 21.3 and 40 ml/m². The interventricular septum size was between 7 and 11.9 mm, and showed a directly proportional trend with the athlete’s age and professional physical activity duration.

The ejection fraction had values between 60 and 70%. A higher than normal regional diastolic longitudinal functional myocardial pattern was observed in 85% of athletes, and it was characterized by high myocardial pro-diastolic velocities, as well as an increased Em/Am ratio. Em velocity is directly correlated to diastolic diameter and systolic output under physical effort. Cardiac valves function was also evaluated; no structural valvular changes were observed. In seven athletes a low-grade tricuspid insufficiency was detected, in eight cases a low-grade mitral insufficiency was observed, and in five cases both tricuspid and mitral insufficiency were observed.

1. Group A vs. B: Left Ventricle Size Comparison
Analyzing the left ventricle size (expressed in ml/mq), no statistical difference was found between the GROUP A and the GROUP B (Figure 1A). A statistically significant difference was

| TABLE 1 | The principal biological parameters for each group. |
|---|---|---|
| Parameters | Group A | Group B |
| Mean age | 26.6 ± 3.1 | 18.3 ± 0.6 |
| Mean height (m) | 1.85 ± 0.08 | 1.83 ± 0.04 |
| Mean weight (Kg) | 78 ± 8.6 | 75.5 ± 5.2 |
| Mean BMI | 22.8 ± 1.13 | 22.5 ± 0.8 |
| Heart rate range | 37–58 bpm | 45–64 bpm |
| Mean blood pressure | 70/110 | 70/110 |
| % SpO₂ | 99% | 99% |
found between the two groups after the second year (58.4 vs. 66.7, \( p < 0.001 \)) and after the third year (60.2 vs. 70.4, \( p < 0.001 \)), Figures 1B,C.

2. Group A vs. B: Left Atrial Size Comparison

Atrial size indexed evaluation showed a statistically significant difference between the two groups after the second and the third year; no significant difference was found after first year (Figures 2A–C).

3. Group A vs. B: Interventricular Septum Size Comparison

The interventricular septum thickness was statistically significant lower in the Group B than in the Group A after the first (\( p = 0.002 \)), second (\( p < 0.001 \)) and third year (\( p < 0.001 \)), Figure 3.

4. Group A vs. B: Fraction of Ejection Comparison

There was no statistically significant difference between the A and the B group analyzing the ejection fraction (Figure 4).

5. Left ventricle size variation during follow-up

The left ventricle size of the athletes in the Group A during the 3 years follow-up showed a progressive statistically significant increase (\( p < 0.001 \)). A significant difference was also observed between the first and the second year (\( p = 0.017 \)), and between the first and the third year (\( p < 0.001 \)), as shown in the Figure 5.

6. Left atrial size variation during follow-up

The left atrium size of the athletes in the Group A increased significantly during the 3 years follow up (\( p < 0.001 \)). A statistically significant difference was also found between the first

---

**FIGURE 1** | The analysis of left ventricle size between Group A and Group B: statistically significant differences were found in the second and third year (A, First year; B, Second year; C, Third year).

**FIGURE 2** | The analysis of left atrial size between Group A and Group B: statistically significant differences were found in the second and third year (A, First year; B, Second year; C, Third year).
and the second year \( (p = 0.02) \), between the first and the third year \( (p < 0.001) \), and between the second and the third year \( (p = 0.003) \), as shown in the Figure 6.

7. Interventricular septum size variation during follow-up

The thickness of the interventricular septum in the Group A players showed a progressive statistically significant increase \( (p < 0.001) \), Figure 7.

8. Ejection fraction variation during follow up

No statistically significant ejection fraction difference was observed in the Group A athletes during the 3 year follow-up (Figure 8).

**DISCUSSION**

Recent guidelines suggested a number of investigations to be performed in order to properly evaluate cardiovascular function in the “Athlete’s Heart” (Salomone et al., 2014; Galderisi et al., 2015; Precenzano et al., 2016b).

Athletes should have unaltered diastolic and systolic functions. This is one of the main distinguishing factors from diastolic and systolic pathological degeneration (D’Andrea et al., 2009).
Recent studies showed peculiar echocardiography parameters changes in elite athletes (Vacchiano and Vyshka, 2012; D’Ascenzi et al., 2015).

Several papers focused on athletes’ echocardiography features, comparing the study group (athletes) with the control group (sedentary) (Galanti et al., 2016; Precenzano et al., 2016c).

This study focused on two different professional athletes groups: Group A, “Serie A” football team, compared to Group B, composed by the respective B team competing in the “Primavera” league. In both groups, clinical examination and color doppler echocardiography were performed. A number of echocardiography parameters were studied, including: left ventricle size, left atrium size, interventricular septum thickness and ejection fraction. Twelve players of Group A is followed for 3 years, recording echocardiographic parameters.

Athletes of the A group showed a higher left cardiac chambers size and a higher interventricular septum size comparing to the athletes of the B group. No significant differences were found regarding ejection fraction. Such results could be a consequence of a different professional training load.

Following the 12 Group A players, a significant increase in the heart size and interventricular septum size during the second and the third year was recorded, comparing to the data collected from the Group B players.

Such evidence suggests how professional athletes training programs lead to significant heart changes, improving heart function.

According to our data, different training load led to different cardiovascular and echocardiography parameters changes.

During follow-up, no significant differences were found regarding ejection fraction. These data are in accord with previous studies: Bangsbo (1994) reported only small increases in heart adaptations after a 5 week pre-season training period in professional soccer players. Other studies have described that the upper limit to which the ventricle wall thickens with training is 16 mm. These are the maximum heart adaptations and further training cannot influence these parameters (Pelliccia et al., 2010).

Our study confirms that physical activity has significant and relevant effects on the cardiovascular system, leading to structural and functional changes that characterize the “athlete’s heart.”

**CONCLUSIONS**

Our findings confirm the direct physiological effect of physical activity; it can be associated with the “Athlete’s heart” changes.
All players of both groups were undergone to clinical and instrumental evaluation; it was reported an increase in left cardiac chambers size and interventricular septum thickness. Such changes were much more relevant in long-term professional football players rather than in young football players, even though a considerable increase in ventricular mass and changes in left ventricle strength have been observed in the latter group. The improvement in diastolic function could possibly be due to the increase in ventricular size and performance. Previous studies conducted on diastolic function in the physiologically hypertrophic heart showed maximum increment in left ventricular size, and normal or higher than normal parietal thinning.

Improvements in diastolic function parameters are associated with increased ventricular size and performance. Mitral and tricuspid valves regurgitation are associated with an increase in cardiac chambers size and consequent enlargement of the valve annulus, similar to what occurs in dilated cardiomyopathy. By performing doppler color mapping, we observed that among 45 athletes, 17.6% had mitral insufficiency, 15.5% tricuspid insufficiency, and 11.9% both mitral and tricuspid insufficiencies. It is important to note that in the professional athletes a small amount of doppler regurgitation without any morphological valvular alterations and clinical symptoms, is considerably frequent and should not be considered as a pathological change, since they are due to a “physiological” volumetric increase of the size of the cardiac chambers induced by exercise.

Our work confirmed that professional players who underwent a more durable and intense physical training program had a much higher level of echocardiographic parameters than younger professional players. In addition, such data was always below the “Athlete’s heart” levels. No significant echocardiographic variation within the examined sample was observed for different roles (goalkeeper, defender, midfielder, or attacker) or skills of individual players.

**AUTHOR CONTRIBUTIONS**

Substantial contributions to the conception or design of the work; or the acquisition, analysis, or interpretation of data for the work: CF, FS, MS, IV, GiM, VM, MM, AM. Drafting the work or revising it critically for important intellectual content: CF, FS, LT, GA, FT, MR, GiM, GaM, MMP, GC, AV, CZ, AM. Final approval of the version to be published: CF, FS, GiM, VM, MM, AM. Agreement to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved: CF, MS, OC, GiM, AM. All authors read and approved the final manuscript.

**REFERENCES**

Atchley, A. E. Jr., and Douglas, P. S. (2007). Left ventricular hypertrophy in athletes: morphologic features and clinical correlates. *Cardiol. Clin. 25*, 371–382. doi: 10.1016/j.ccl.2007.06.009

Avola, R., Di Tullio, M. A., Fisichella, A., Tayebati, S. K., and Tosassoni, D. (2004). Glial fibrillary acidic protein and vimentin expression is regulated by glucocorticoids and neurotrophic factors in primary rat astroglial cultures. *Clin. Exp. Hypertens.* 26, 323–333. doi: 10.1080/10491960490456980

Bangsbo, J. (1994). The physiology of soccer - With special reference to intense intermittent exercise. *Acta Physiol. Scand. Suppl. 619*, 1–155.

Bangsbo, J., Mohr, M., and Krstrup, P. (2006). Physical and metabolic demands of training and match-play in the elite football player. Nutrition and Football: The FIFA/FMARC consensus on sports nutrition. *J. Sports Sci. 24*, 665–674. doi: 10.1080/0266611050048329

Brancati, F., Tosassoni, D., Grasso, S., Bronzi, D., Napoli, M., Campisi, A., et al. (2012). Cholinergic precursors modulate the expression of heme oxygenase-1, p21 during astroglial cell proliferation and differentiation in culture. *Neurochem. Res. 37*, 2795–2804. doi: 10.1007/s11064-012-0873-3

Braschi, A., Abbrignani, M. G., Francavilla, V. C., and Francavilla, G. (2011). Novel electrocardiographic parameters of altered repolarization in uncomplicated overweight and obesity. *Obesity 19*, 875–881. doi: 10.1038/oby.2010.252

Braschi, A., Francavilla, V. C., Abbrignani, M. G., Lombardo, R., Colivicchi, F., and Francavilla, G. (2009). Usefulness of QT dispersion in clinical practice [Utilità della dispersione del QT nella pratica clinica]. *Monaldi Arch. Chest Disease Card. Series. 72*, 118–129.

Braschi, A., Francavilla, V. C., Abbrignani, M. G., Todaro, L., and Francavilla, G. (2012). Behavior of repolarization variables during exercise test in the athlete’s heart. *Ann. Noninvasive Electrocardiol. 17*, 95–100. doi: 10.1111/j.1542-474X.2012.00495.x

Chieffi, S., Messina, G., Villano, I., Messina, A., Esposito, M., Monda, V., et al. (2017). Exercise influence on hippocampal function: possible involvement of orexin-A. *Front. Physiol. 8*:85. doi: 10.3389/fphys.2017.00085

Cianci, P., Tartaglia, N., Altamura, A., Fersini, A., Vovola, F., Sanguedolce, F., et al. (2016). A recurrent epidermoid cyst of the spleen: report of a case and literature review. *World J. Surg. Oncol. 14*:98. doi: 10.1186/s12957-016-0857-x

D’Andrea, A., Galderisi, M., Scieri, S., Nistri, S., Agricola, E., Ballo, P., et al. (2009). Echocardiographic evaluation of the athlete’s heart: from morphological adaptations to myocardial function [Lo studio ecocardiografico del cuore d’atleta: dagli adattamenti morfologici alla funzione miocardica]. *G. Ital. Cardiol.* 10, 533–544.

D’Ascenzi, F., Polliccia, A., Cameli, M., Lisi, M., Natali, B. M., Focardi, M., et al. (2015). Dynamic changes in left ventricular mass and in fat-free mass in top-level athletes during the competitive season. *Eur. J. Prev. Cardiol.* 22, 127–134. doi: 10.1177/2047487315058820

Fagard, R. (2003). Athlete’s heart. *Heart* 89, 1455–1461.

Forlano, I., Fersini, A., Tartaglia, N., Ambrosi, A., and Neri, V. (2011). Bilary papillomatosis: case report. *Ann. Ital. Chir.* 82, 405–408.

Francavilla, G., Abbrignani, M. G., Braschi, A., Sciaccia, R., Francavilla, V. C., Caracchiolo, M. M., et al. (2007). Physical exercise and sport activities in patients with and without coronary heart disease. *Monaldi Arch. Chest Dis. 68*, 87–98.

Francavilla, V. C., Braschi, A., Ciacono, A., Di Pietro, V., Rongiovanni, T., and Francavilla, G. (2016). A new prevention strategy for amateur athletes: spatial QT dispersion [Una nuova strategia di prevenzione per gli sportivi amatoriali: la dispersione spaziale del QT]. *Med. Dello Sport 69*, 249–253.

Galanti, G., Stefani, L., Mascherini, G., Di Tante, V., and Toncelli, L. (2016). Left ventricular remodeling and the athlete's heart, irrespective of quality load training. *Cardiovasc. Ultrasound* 14:86. doi: 10.1186/s12974-016-0088-x

Galderisi, M., Cardim, N., D’Andrea, A., Bruder, O., Cosyns, B., Davin, L., et al. (2015). The multi-modality cardiac imaging approach to the athlete’s heart: an expert consensus of the european association of cardiovascular imaging. *Eur. Heart J. Cardiovasc. Imaging* 16:353. doi: 10.1093/ehjci/jeu332

Ganau, A., Devereux, R. B., Roman, M. J., de Simone, G., Pickering, T. G., Saba, P. S., et al. (1991). Patterns of left ventricular hypertrophy and geometric remodeling in essential hypertension. *J. Am. Coll. Cardiol. 19*, 1550–1558. doi: 10.1016/0735-1097(92)90617-V

Lamotte, M., Fleury, F., Pirard, M., Jamon, A., and van de Borne, P. (2010). Acute cardiovascular response to resistance training during cardiac rehabilitation: effect of repetition speed and rest periods. *Eur. J. Prev. Cardiol.* 17, 329–336. doi: 10.1177/1078181309353266

Lavie, C. J., Milani, R. V., Marks, P., and de Gruiter, H. (2001). Exercise and the heart: risks, benefits, and recommendations for providing exercise prescriptions. *Oxidizer J. 3*, 207–213.
Maté-Muñoz, J. L., Lougado, J. H., Barba, M., García-Fernández, P., Garnacho-Castaño, M. V., and Dominguez, R. (2017). Muscular fatigue in response to different modalities of CrossFit sessions. *PLoS ONE* 12:e0181855. doi: 10.1371/journal.pone.0181855

Messina, G., Di Bernardo, G., Viggiano, A., De Luca, V., Monda, V., Messina, A., et al. (2016). Exercise increases the level of plasma orexin A in humans. *J. Basic Clin. Physiol. Pharmacol.* 27, 611–616. doi: 10.1515/jbcpp-2015-0133

Mielpo-Ayuso, J., Maroto-Sánchez, B., Luzardo-Socorro, R., Palacios, G., Palacios-Gil-Antuñano, N. and González-Gross, M. (2015). Evaluation of nutritional status and energy expenditure in athletes. *Nutr. Hosp.* 31, 227–236. doi: 10.3305/nh.2015.31.sup3.8770

Mitchell, J. H., Haskell, W., Snell, P., and Van Camp, S. P. (2005). Task force 8: classification of sports. *J. Am. Coll. Cardiol.* 45, 1364–1367. doi: 10.1016/j.jacc.2005.02.015

Morgans, R., Orme, P., Anderson, L., and Drust, B. (2014). Principles and practices of training for soccer. *J. Sport Health Sci.* 3, 251–257. doi: 10.1016/j.jshs.2014.07.002

Naylor, L. H., George, K., O’Driscoll, G., and Green, D. J. (2008). The athlete’s heart: a contemporary appraisal of the ‘Morganroth hypothesis’. *Sports Med.* 38, 69–90. doi: 10.2165/00007256-200838010-00006

Neri, V., Ambrosi, A., Fersini, A., Tartaglia, N., Lapolla, F., and Forlano, I. (2013). Severe acute pancreaticitis: clinical forms of different gravity. *Ann. Ital. Chir.* 84, 47–53.

Neri, V., Fersini, A., Ambrosi, A., Tartaglia, N., and Valentino, T. P. (2009a). Diagnostic evaluation prior to cholecystectomy in mild-moderate acute biliary pancreatitis. *Ann. Ital. Chir.* 80, 363–367.

Neri, V., Fersini, A., Ambrosi, A., Tartaglia, N., and Valentino, T. P. (2009b). Milder-moderate acute biliary pancreatitis: role of magnetic resonance cholangiopancreatography in preparation of cholecystectomy. *Pancreas* 38, 6, 717.

Pelliccia, A., Culasso, F., Di Paolo, F. M., and Moron, B. J. (1999). Physiologic left ventricular cavity dilatation in elite athletes. *Ann. Intern. Med.* 130, 23–31. doi: 10.7326/0003-4819-130-1-199901050-00005

Pelliccia, A., Di Paolo, F. M., De Blassis, E., Quattrini, F. M., Piccigdio, C., Guerra, E., et al. (2010). Prevalence and clinical significance of aortic root dilation in highly trained competitive athletes. *Circulation* 122, 698–706. doi: 10.1161/CIRCULATIONAHA.109.901074

Pelliccia, A., Moron, B. J., Spataro, A., Proschan, M. A., and Spirito, P. (1991). The upper limit of physiologic cardiac hypertrophy in highly trained elite athletes. *N. Engl. J. Med.* 324, 295–301. doi: 10.1056/NEJM199101313240504

Precenzano, F., Lombardi, P., Ruberto, M., Parisi, L., Salerno, M., Maltese, A., et al. (2016a). Internalizing symptoms in children affected by childhood absence epilepsy: a preliminary study. *Acta Med. Mediterr.* 32, 1749–1753. doi: 10.19193/0393-6384_2016_6_158

Precenzano, F., Ruberto, M., Parisi, L., Salerno, M., Maltese, A., D’Alessandro, I., et al. (2016b). Borderline intellectual functioning and parental stress: an Italian case-control study. *Acta Med. Mediterr.* 32, 1762–1765. doi: 10.19193/0393-6384_2016_6_160

Precenzano, F., Ruberto, M., Parisi, L., Salerno, M., Maltese, A., D’alessandro, I., et al. (2016c). Adhd-like symptoms in children affected by obstructive sleep apnea syndrome: a case-control study. *Acta Med. Mediterr.* 32, 1756–1759. doi: 10.19193/0393-6384_2016_6_159

Salomone, F., Li Volti, G., Vitaglione, P., Morisco, F., Fogliano, V., Zappalà, A., et al. (2014). Coffee enhances the expression of chaperones and antioxidant proteins in rats with nonalcoholic fatty liver disease. *Transl. Res.* 163, 593–602. doi: 10.1016/j.trsl.2013.12.001

Sessa, F., Chetta, M., Petito, A., Franzetti, M., Bafunno, V., Pisanelli, D., et al. (2011). Gene polymorphisms and sport attitude in Italian athletes. *Genet. Test. Mol. Biomarkers* 15, 285–290. doi: 10.1089/gtmb.2010.0179

Spirito, P., Pelliccia, A., Proschan, M. A., Granata, M., Spataro, A., Bellone, P., et al. (1994). Morphology of the ‘athlete’s heart’ assessed by echocardiography in 947 elite athletes representing 27 sports. *Am. J. Cardiol.* 74, 802–806. doi: 10.1016/0003-9497(94)00439-1

Turtizlaz, E., Vacciano, G., Luna-Maldonado, A., Neri, M., Pomara, C., Rabboczi, R., et al. (2010). Tryptase, CD15 and IL-15 as reliable markers for the determination of soft and hard ligature marks vitality. *Histol. Histopathol.* 25, 1539–1546. doi: 10.14670/HH-25.1539

Vacciano, G., and Vyshka, G. (2012). Balkans syndrome: a potential link between multiple sclerosis and hypervaccination. *J. R. Army Med. Corps* 158, 338–340. doi: 10.1136/rarmc-158-04-15

Vyshka, G., and Vacciano, G. (2014). Severe flaccid paraparesis following spinal anaesthesia: a sine materia occurrence. *BMJ Case Rep.* 2014:bcr2013202071. doi: 10.1136/bcr-2013-202071

Zeppilli, P. (2008). “Adattamenti cardiocircolatori nelle diverse discipline sportive,” in *Cardiologia dello Sport*, ed P. Zeppilli (Roma: CESI), 41–83.

Zeppilli, P., D’Andrea, A., Gaddi, O., Romano, S., and Penco, M. (2008). “L’ecocardiogramma nell’atleta,” in *Cardiologia dello Sport*, ed P. Zeppilli (Roma: CESI), 197–269.

Zurutuza, U., Castellano, J., Echezarra, I., and Casamichana, D. (2017). Absolute and relative training load and its relation to fatigue in football. *Front. Psychol.* 8:878. doi: 10.3389/fpsyg.2017.00878

Conflict of Interest Statement: The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

The handling Editor declared a shared affiliation, though no other collaboration, with one of the authors OC.

Copyright © 2018 Francavilla, Sessa, Salerno, Albano, Villano, Messina, Triolo, Todaro, Ruberto, Marsala, Cascio, Mollica, Monda, Ciulli, Valenzano, Zammit, Monda and Messina. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.