



# Profile of Steroid Hormones and sex Hormone-Binding Globulin of Elite Soldiers

Thomaz Talarico Neto <sup>1,\*</sup>, Higino Carlos Hahns Júnior <sup>1</sup>, Maria Clara Santos Minelli <sup>1</sup>, José Mauricio Magraner Paixão dos Santos <sup>1</sup>, Marcos Corrêa Junior <sup>2</sup>, Yanesko Fernandes Bella <sup>3</sup>, Fernanda Galante <sup>4</sup>, Cesar Baraldi Gonella <sup>5</sup>, Leandro Ferreira <sup>6</sup>, Carlos Eduardo Martinelli Junior <sup>2</sup> and Hugo Tourinho Filho <sup>1</sup>

<sup>1</sup>School of Physical Education and Sport of Ribeirao Preto, University of Sao Paulo, Ribeirao Preto, Brazil

<sup>2</sup>Department of Childcare and Pediatrics, University of Sao Paulo, Ribeirao Preto, Brazil

<sup>3</sup>Mackenzie Presbyterian University, São Paulo, Brazil

<sup>4</sup>Institute of Biomedical Sciences, Integrated Colleges of São Paulo, São Paulo, Brazil

<sup>5</sup>University Center of Faculdades Metropolitanas Unidas, Piracicaba, Brazil

<sup>6</sup>Unifafibe University Center, Bebedouro, Brazil

\*Corresponding author: Master Student, School of Physical Education and Sport of Ribeirao Preto, University of Sao Paulo, Ribeirao Preto, Brazil. Email: thomaz.talarico.edf@gmail.com

Received 2020 October 16; Accepted 2020 October 28.

## Abstract

**Background:** Military police training should characterize the necessary abilities to conduct their activities with excellence. Biochemical parameters have a significant role in the performance of individuals and, therefore, are of crucial importance when trying to improve the teamwork of police officers, particularly for special operations groups. Hence, the use of laboratory analysis in activities that cause great physiological stress is of paramount importance.

**Objectives:** The present study aimed to evaluate the profile of steroid hormones and the sex hormone-binding-globulin of soldiers belonging to a special operations force.

**Methods:** In this cross-sectional study, blood samples of 75 volunteers collected after 12 hours of fasting are analyzed. Data were analyzed using descriptive statistics (i.e., mean and standard deviation) and compared to the reference values for each biomarker, referencing each individual in their age group.

**Results:** Most of the participants presented adequate concentrations for their respective ages in relation to the data analyzed. Even the inclusion of individuals from different age groups did not change the good results found in the group.

**Conclusions:** The positive results found in such a heterogeneous group with respect to age group can be attributed to an extremely rigorous selection process and a training program that elevates them to the rank of elite soldiers of the military police.

**Keywords:** Sex Hormone-Binding Globulin, Elite Soldiers, Steroid Hormones

## 1. Background

Military police training should characterize the necessary abilities to conduct their activities with excellence. They all must be physically and psychologically prepared to guarantee their own safety, the safety of others, and also of society regarding the fact that these professionals frequently endanger their lives (1).

That said, the training needed to achieve this excellence is very arduous, requiring exhaustive simulations to prepare the trainees for the realities that await them in the field (2). Since special operations groups (SOGs) require unique operational capabilities, this issue is even more evident for them. As physical, biochemical, and psychologi-

cal parameters heavily influence the performance of individuals, concerning both excellence in performing activities and longevity of the career, special attention should be paid to these parameters (2).

Battalions of Special Police Operations must deal with exceptional cases within the military police, acting in the fields of public defense, with a mission to solve problems that are beyond the aptitude of the operational units of the military police (3). Therefore, these groups should be made up of highly armed and specialized troops, working to combat heavily armed gangs on their missions in either urban or rural counter-guerrilla fields (3).

Therefore, laboratory analyses not only would provide valuable information for activities that cause great physi-

ological stress (4), but also are useful for monitoring the training and nutritional status as well as identifying possible complications caused by the practice of a particular modality and/or work activity, consequently, focusing on longevity and health (4).

Within this line of reasoning, hormones are chemical signals secreted by specialized glands or cells, which help to control various functions in the body. Hormones stimulate signals to both distant cells (endocrine action) and adjacent tissues (paracrine action). These stimulations always act selectively, because only the cell that has a certain hormone receptor can respond to the stimulus (5).

Since a certain type of hormone can influence another (either an amplifying or suppressing effect), they act in a synchronized manner, not in isolation. Therefore, joint harmony is necessary to demonstrate good functioning (5). Regarding the action pathways of hormones, they can be used as important metabolic biomarkers of training and nutritional status.

Testosterone is a steroid male hormone that circulates in the body by binding to Sex Hormone Binding Globulin (SHBG), having only a small free-flowing portion. However, based on the free hormone theory, this small free portion binds to androgen receptors (6). Testosterone is responsible for several important functions in the body, including having a fundamental role in protein and lipid metabolism (7). In humans, this hormone has a great influence on body composition and muscle mass (7). Several symptoms are associated with testosterone deficiency, including increased central adiposity, reduced insulin sensitivity, increased triglycerides, and reduced HDL. These factors, in turn, contribute to metabolic syndrome and diabetes mellitus, as well as the increased risk of cardiovascular problems (8). Therefore, to perform an adequate evaluation, it is necessary to measure both total and free testosterone as well as its main binding protein. Dehydroepiandrosterone sulfate (DHEA) is one of the most abundant steroid hormones in humans, which is produced by the adrenal glands. The DHEA is a steroid hormone precursor (such as testosterone). Moreover, it also has an important role in the immune system, which justifies the necessity for its adequate levels of production for the proper functioning of the body (9).

Historically, it was believed that testosterone solely plays several key activities. Nevertheless, now it's well proved that several other hormones (e.g., estradiol) also contribute to functions that were historically attributed to testosterone, mainly because testosterone can undergo peripheral aromatization and transform into estradiol (about half of male estradiol have this origin) (10). The other portion of this hormone is produced from testicular secretion. In men, estradiol exerts important actions

on bones, fat mass, and insulin resistance, which makes it an important hormone for analysis and one which is often overlooked in males (10).

When taking into account the importance of the balance of hormones and their binding proteins, presenting values within the normal range, according to age groups, are vital for health, contributing to the proper functioning of the metabolism of proteins, carbohydrates, and lipids.

If we believe that these hormonal values are of crucial importance for the normal population, when it comes to professionals who work in stressful conditions that need maximum efficiency of their organism, a higher value should be given to them. Concerning police officers working in SOGs, it should be noted that any disturbance can put their lives at unnecessary risk and may even cause them to leave work due to a health problem, a fact that can harm both the individual and society as a whole.

Therefore, hormonal monitoring is of great value since it provides a view of the current state of the subjects who are constantly exposed to extreme professional challenges.

## 2. Objectives

Within this line of reasoning, the present study aimed to evaluate the profile of steroid hormones and the sex hormone-binding protein of soldiers belonging to a special operations force.

## 3. Methods

### 3.1. Study Design and Sample Population

In this cross-sectional study, blood samples of 75 volunteers who were members of a SOG were collected after 12 hours of overnight fasting. Before collecting blood samples, the subjects were instructed to remain at rest and not perform vigorous activities before the analyses. The blood collection was performed at the battalion headquarters. Inclusion criteria were being an active member of a SOG, having medical clearance, and willingness to participate in the study.

### 3.2. Ethical Considerations

The present study is approved by the ethics and research committee of the Faculty of Physical Education and Sport of Ribeirão Preto EEFERP/USP (n. 3.230.057). Besides, written informed consent was obtained from all participants, and the researchers were available to answer any question regarding the objectives of the study.

### 3.3. Experimental Procedure

Blood samples were collected by puncture via access to the blood vessel through the anterior aspect of the forearm. Before collection, the forearm was properly cleaned with 70% ethanol. In total, 5 ml of blood were collected in tubes with separator gel to obtain the serum and immediately stored at 0 to 4°C. Afterward, samples were centrifuged at 3000rpm in a temperature ranging from zero to 4°C for 15 minutes to separate the serum. Then, serum was removed from the collection tube and stored in 1.5mL tubes for subsequent freezing at -80°C. The remaining contents of the collection tubes were disposed of appropriately.

The analyses of total testosterone, free testosterone, SHBG, S-DHEA, and estradiol were performed with the chemiluminescence method, using blood serum, and were dosed by an authorized laboratory in the city of Rio de Janeiro - CNPJ: 07.727.439 / 0001-60 - License: 1402.

Anthropometric assessment of weight and height was performed using a stadiometer with a 0.1 cm measurement scale and a Plenna digital scale model MEA-07400 with 100g accuracy, respectively.

### 3.4. Statistical Analysis

Data were analyzed using descriptive statistics (mean and standard deviation) and compared to the reference values for each biomarker, referencing each individual in their age range classification. SPSS version 22.0 was used for statistical analysis.

## 4. Results

The mean age of participants was  $37.23 \pm 6.08$  years. Also, their average weight and height were  $80.4 \pm 9.9$  kg and  $1.7 \pm 0.05$  m, respectively. An overall view of biochemical analyses of samples is provided in [Table 1](#).

### 4.1. Total Testosterone (TT)

Due to technical reasons, three samples were excluded from the analyses. Therefore, in total 72 blood samples were studied. The mean concentration of TT was  $602.43 \pm 247.82$  ng/dl. As the classification of this hormone varies according to age, each individual received an individual classification. Since the youngest and oldest participants were 26 and 54 years old, the following classification was applied as reference values: 21-49 years ( $240.24 - 870.68$  ng/dl);  $\geq 50$  years ( $220.91-715.81$  ng/dl). Using this framework, only nine individuals presented values outside the normal range; among these, one was below the normal range, and eight were above the recommended threshold.

### 4.2. Free testosterone (FT)

For the FT values, that is, the testosterone that is not bound to any protein, the group mean was  $439.40 \pm 253.37$ . The mean was calculated considering 71 individuals since four samples did not obtain a reading. The reference value for this variable is unique, with no difference between ages. Thus, a parameter range of 131.00 - 640.00 pmol/L was considered to be ideal. The group value was within the recommended. When analyzing the individual results, only four individuals presented values outside the normal range; among these, three were above the ideal and one below.

### 4.3. Sex Hormone-Binding Protein (SHBG)

In this part, five samples were removed due to not obtaining a reading. Hence, in total, 70 samples were analyzed. The mean SHBG of the whole group was  $35.78 \pm 12.38$ . The reference value is unique for this protein, with a normal range of 11.20-78.10 nmol/L. The overall mean was in the normal range. Furthermore, for individual participants, also the obtained values were in the normal range.

### 4.4. Dehydroepiandrosterone Sulfate (S-DHEA)

For the S-DHEA, the overall mean for all 75 samples was  $269.00 \pm 129.85$ . The normal value defined for the S-DHEA varies by age. In this sense, the following classification was used as reference values: 25-34 years ( $167.90-591.90$  ug/dl); 35-44 years ( $139.70-484.40$  ug/dl); and 45-54 years ( $136.20-447.60$  ug/dl). In this case, 17 individuals presented abnormal values, who 13 were below the recommended range, and only four were higher.

### 4.5. Estradiol

One sample was excluded due to not obtaining a valid reading. Hence, 74 samples were analyzed. The overall mean was  $26.55 \pm 10.87$  pg/ml. A single reference value is defined for the estradiol, covering all age groups, ranging from 11.00 to 44.00 pg/ml. The overall mean of the group fell in this range. Nevertheless, concerning individual samples, seven were out of the expected range (3 below and 4 above the recommended range).

## 5. Discussion

Older adults often are not efficient for performing activities that require high physical performance during working hours, mainly due to biological alterations caused by increased age, which led to declined functional capacity (1). The stability of hormonal factors indicates the biological preparedness of individuals, which is a very important factor for their training and overall performance (4).

**Table 1.** An overall view of the results of the biochemical analyses performed on blood samples

Biomarker	No.	Mean $\pm$ SD	Min	Max
TT	72	602.43 $\pm$ 247.82	117.82	1944.21
FT	71	439.40 $\pm$ 253.37	83.11	2312.51
SHBG	70	35.78 $\pm$ 12.38	15.8	73.4
S-DHEA	75	269 $\pm$ 129.85	57.55	768.1
Estradiol	74	26.55 $\pm$ 10.87	10	77.27

Abbreviation: TT, total testosterone; FT, free testosterone; SHBG, sex hormone-binding protein; S-DHEA, dehydroepiandrosterone sulfate.

In the present study, participants were quite diverse concerning their age. So that out of the total 75 participants, 27 were in the age group of 26-34 years, 23 in 35-39 years, and 25 in 40-52 years. Nevertheless, age was not a confounding variable in the analyses since, when refining the results, it was found that the older participants and their younger counterparts were well-classified, which is most likely the result of an extremely rigorous selection applied in the present study.

Testosterone has a fundamental effect on the body. So that several population-based studies have reported that testosterone deficiency significantly increases the mortality from any cause and is also an independent risk factor for cardiovascular diseases (12-14). It also causes several other negative health consequences as follows: impaired insulin sensitivity, increased body fat (mainly central fat), dyslipidemia, and hypertension (8). Therefore, based on the laboratory results, most of the participants of the present study are immune against negative health consequences of testosterone deficiency.

Overall, estradiol values were almost entirely within the normal range; however, a thorough analysis pointed to two individuals who presented very higher estradiol values. These subjects also had high values of TT and FT, which demonstrates that the excess of testosterone is aromatized to estradiol in the periphery through the aromatase enzyme, which belongs to the cytochrome p450 enzymes family (15). Although they were isolated cases, but their blood samples were analyzed by a multidisciplinary team, which performed the entire procedure for necessary corrections and adjustments.

The good concentration of S-DHEA found in the present study guarantees an adequate contribution to the supply of testosterone concentrations because although testicles are the main producer of testosterone, S-DHEA also produces this hormone (9).

Regarding the SHBG, a study has demonstrated that during intense military training, individuals who attained an overreaching state (excessive accumulation of stress that causes a momentary decline in performance) already

had higher baseline SHBG values relative to before training. The authors argued that the enhanced concentration of testosterone results in its increased binding to the SHBG, which in turn decreases its availability, which may negatively influence the body responses during training (16). This issue highlights the importance of adequate levels of this binding protein.

Another study has shown that dietary restriction plus intense military training could increase SHBG and decrease the testosterone concentration, which in turn greatly reduced the availability of this hormone (17).

Continuous monitoring of hormonal concentrations and proteins that carry them is necessary to maintain the performance of highly active populations, as these analyses can provide valuable information regarding biomarkers that reveal biological changes of the body. This information can also be used to change behaviors to prevent negative consequences (18).

Therefore, when performing hormonal analyses, it is suggested to have a holistic approach, as investigating a sole hormone is unlikely to provide reliable information. Besides, applying a set of analyses creates an environment that favors a more reliable view of what is happening in the organism, which minimizes the possible misinterpretations that would have been occurred if an isolated hormonal fraction was followed.

In conclusion, it can be concluded that most of the elite soldiers investigated in the current study presented recommended values of steroid hormones and sex hormone-binding protein. Furthermore, including individuals of various ages didn't change the good results found in the group.

Attributing positive results found in the present study in such a heterogeneous group with a diverse age structure to the extremely rigorous selection process and the training program that has elevated them to the rank of elite soldiers of the military police seems to be reasonable.

## Footnotes

**Authors' Contribution:** Study concept and design: YB, and FG, and CBG; acquisition of data: JMMPs, and MCJ; analysis and interpretation of data: TTN, and HCHJ, and MCMS; drafting of the manuscript: TTN, and HCHJ, and MCMS; critical revision of the manuscript for important intellectual content: HTF, and CEMJ; statistical analysis: LF, and JMMPs; administrative, technical, and material support: MCJ, and YB, and FG, and CBG; study supervision: HTF, and CEMJ, and LF.

**Conflict of Interests:** The authors declare no conflict of interest.

**Ethical Approval:** The current research was approved by the ethics and research committee of the Faculty of Physical Education and Sport of Ribeirão Preto EEFERP/USP (n. 3.230.057).

**Funding/Support:** The current study did not receive any funding.

**Informed Consent:** Written informed consent was obtained from all participants and the researchers were available to answer any question regarding the objectives of the study.

## References

- Zanini MTF, Santos MCCC, Lima DDFP. A influência do estilo de liderança consultivo nas relações de confiança e comprometimento no Batalhão de Operações Policiais Especiais do Rio de Janeiro. *Rev Adm*. 2015;**50**(1):105-20. portuguese. doi: [10.5700/rausp1187](https://doi.org/10.5700/rausp1187).
- Gomez-Merino D, Chennaoui M, Drogou C, Guezennec CY. Influence of energy deficiency on the insulin-like growth factor I axis in a military training program. *Horm Metab Res*. 2004;**36**(7):506-11. doi: [10.1055/s-2004-825730](https://doi.org/10.1055/s-2004-825730). [PubMed: [15305236](https://pubmed.ncbi.nlm.nih.gov/15305236/)].
- dos Santos MR. Estudo do perfil dermatoglífico, somatotípico e das qualidades físicas dos policiais do batalhão de operações especiais (PMERJ) do ano de 2005. *Fitness & Performance Journal*. 2007;**6**(2):98-104. doi: [10.3900/fpj.6.2.98.p](https://doi.org/10.3900/fpj.6.2.98.p).
- Nunes LAS. *Exames laboratoriais no esporte: Guia prático para interpretação dos exames laboratoriais de atletas e praticantes de atividade física*. 2th. São Paulo: PoloBooks; 2019. Portuguese.
- Kraemer WJ, Rogol AD. *The endocrine system in sports and exercise*. 2005. doi: [10.1002/9780470757826](https://doi.org/10.1002/9780470757826).
- Keevil BG, Adaway J. Assessment of free testosterone concentration. *J Steroid Biochem Mol Biol*. 2019;**190**:207-11. doi: [10.1016/j.jsbmb.2019.04.008](https://doi.org/10.1016/j.jsbmb.2019.04.008). [PubMed: [30970279](https://pubmed.ncbi.nlm.nih.gov/30970279/)].
- Kelly DM, Jones TH. Testosterone: a metabolic hormone in health and disease. *J Endocrinol*. 2013;**217**(3):R25-45. doi: [10.1530/JOE-12-0455](https://doi.org/10.1530/JOE-12-0455). [PubMed: [23378050](https://pubmed.ncbi.nlm.nih.gov/23378050/)].
- Wang C, Jackson G, Jones TH, Matsumoto AM, Nehra A, Perelman MA, et al. Low testosterone associated with obesity and the metabolic syndrome contributes to sexual dysfunction and cardiovascular disease risk in men with type 2 diabetes. *Diabetes Care*. 2011;**34**(7):1669-75. doi: [10.2337/dc10-2339](https://doi.org/10.2337/dc10-2339). [PubMed: [21709300](https://pubmed.ncbi.nlm.nih.gov/21709300/)]. [PubMed Central: [PMC3120209](https://pubmed.ncbi.nlm.nih.gov/PMC3120209/)].
- Cormier C, Souberbielle J, Kahan A. DHEA in bone and joint diseases. *Joint Bone Spine*. 2001;**68**(6):588-94. doi: [10.1016/s1297-319x\(01\)00327-x](https://doi.org/10.1016/s1297-319x(01)00327-x).
- Russell N, Grossmann M. Mechanisms in endocrinology: Estradiol as a male hormone. *Eur J Endocrinol*. 2019;**181**(1):R23-43. doi: [10.1530/EJE-18-1000](https://doi.org/10.1530/EJE-18-1000). [PubMed: [31096185](https://pubmed.ncbi.nlm.nih.gov/31096185/)].
- Milanovic Z, Pantelic S, Trajkovic N, Sporis G, Kostic R, James N. Age-related decrease in physical activity and functional fitness among elderly men and women. *Clin Interv Aging*. 2013;**8**:549-56. doi: [10.2147/CIA.S44112](https://doi.org/10.2147/CIA.S44112). [PubMed: [23723694](https://pubmed.ncbi.nlm.nih.gov/23723694/)]. [PubMed Central: [PMC3665513](https://pubmed.ncbi.nlm.nih.gov/PMC3665513/)].
- Araujo AB, Dixon JM, Suarez EA, Murad MH, Guey LT, Wittert GA. Clinical review: Endogenous testosterone and mortality in men: a systematic review and meta-analysis. *J Clin Endocrinol Metab*. 2011;**96**(10):3007-19. doi: [10.1210/jc.2011-1137](https://doi.org/10.1210/jc.2011-1137). [PubMed: [21816776](https://pubmed.ncbi.nlm.nih.gov/21816776/)]. [PubMed Central: [PMC3200249](https://pubmed.ncbi.nlm.nih.gov/PMC3200249/)].
- Vikan T, Schirmer H, Njolstad I, Svartberg J. Endogenous sex hormones and the prospective association with cardiovascular disease and mortality in men: the Tromso Study. *Eur J Endocrinol*. 2009;**161**(3):435-42. doi: [10.1530/EJE-09-0284](https://doi.org/10.1530/EJE-09-0284). [PubMed: [19542243](https://pubmed.ncbi.nlm.nih.gov/19542243/)].
- Khaw KT, Dowsett M, Folkard E, Bingham S, Wareham N, Luben R, et al. Endogenous testosterone and mortality due to all causes, cardiovascular disease, and cancer in men: European prospective investigation into cancer in Norfolk (EPIC-Norfolk) Prospective Population Study. *Circulation*. 2007;**116**(23):2694-701. doi: [10.1161/CIRCULATIONAHA.107.719005](https://doi.org/10.1161/CIRCULATIONAHA.107.719005). [PubMed: [18040028](https://pubmed.ncbi.nlm.nih.gov/18040028/)].
- Ghosh D, Griswold J, Erman M, Pangborn W. Structural basis for androgen specificity and oestrogen synthesis in human aromatase. *Nature*. 2009;**457**(7226):219-23. doi: [10.1038/nature07614](https://doi.org/10.1038/nature07614). [PubMed: [19129847](https://pubmed.ncbi.nlm.nih.gov/19129847/)]. [PubMed Central: [PMC2820300](https://pubmed.ncbi.nlm.nih.gov/PMC2820300/)].
- Tanskanen MM, Kyrolainen H, Uusitalo AL, Huovinen J, Nissila J, Kinunen H, et al. Serum sex hormone-binding globulin and cortisol concentrations are associated with overreaching during strenuous military training. *J Strength Cond Res*. 2011;**25**(3):787-97. doi: [10.1519/JSC.0b013e3181c1fa5d](https://doi.org/10.1519/JSC.0b013e3181c1fa5d). [PubMed: [20543745](https://pubmed.ncbi.nlm.nih.gov/20543745/)].
- Friedl KE, Moore RJ, Hoyt RW, Marchitelli LJ, Martinez-Lopez LE, Askew EW. Endocrine markers of semistarvation in healthy lean men in a multistressor environment. *J Appl Physiol* (1985). 2000;**88**(5):1820-30. doi: [10.1152/jappl.2000.88.5.1820](https://doi.org/10.1152/jappl.2000.88.5.1820). [PubMed: [10797147](https://pubmed.ncbi.nlm.nih.gov/10797147/)].
- Nindl BC, Alemany JA, Kellogg MD, Rood J, Allison SA, Young AJ, et al. Utility of circulating IGF-I as a biomarker for assessing body composition changes in men during periods of high physical activity superimposed upon energy and sleep restriction. *J Appl Physiol* (1985). 2007;**103**(1):340-6. doi: [10.1152/japplphysiol.01321.2006](https://doi.org/10.1152/japplphysiol.01321.2006). [PubMed: [17412783](https://pubmed.ncbi.nlm.nih.gov/17412783/)].