# **ORIGINAL ARTICLE**

# IGFBP-3, a sensitive marker of physical training and overtraining

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**Objective:** To investigate the response of the somatotrope axis (insulin-like growth factor-1 (IGF-1), insulinlike growth factor binding protein-3 (IGFBP-3)) to intense exercise in relation to tiredness.

**Methods:** The study involved 11 rugby players who completed a questionnaire intended to evaluate fitness or, conversely, overtraining and who agreed to plasma samples being taken before and after an international rugby match.

**Results:** The main finding of our study is that we observed strong negative correlations between IGF-1 (r=0.652) and IGFBP-3 (r=0.824) levels and the overtraining state estimated using the French Society of Sport Medicine questionnaire. In particular, there was a fall (of up to 25%) in IGFBP-3 levels after the match in the more fatigued subjects compared to an increase (of up to 40%) in fit subjects.

**Conclusions:** A fall in IGFBP-3 in response to an intense bout of exercise may represent an index of tiredness in highly trained sportsmen, as indicated by the scores obtained from the overtraining questionnaire.

Physical exercise is a powerful stimulus for growth hormone (GH) secretion, certain actions of which take place by way of growth factors called somatomedins, the principal one being somatomedin-C or insulin-like growth factor-1 (IGF-1). IGF-1 plays a role in many physiological processes,1 particularly in organism anabolism, related to bone and muscular growth.<sup>2</sup> About 80% of IGF-1 occur in a ternary 150 kDa complex including insulin-like growth factor binding protein-3 (IGFBP-3) and a protein called acid-labile subunit; less than 1% of IGF-1 are free.3-5 The IGFBPs prolong the half life of the IGFs, serve as IGF carriers in the circulation, and act as autocrine and/or paracrine regulators of their biological actions.<sup>6</sup> <sup>7</sup> IGFBP-3, which is synthesised in many tissues, is considered the major IGF-1 carrier.<sup>7</sup> It is well documented that circulating levels of IGF-1 and IGFBP-3 are regulated by nutrition, age, pregnancy, chronic diseases, insulin, and GH.<sup>5 7 8</sup> A reduction in IGF-1 concentration is commonly associated with nutritional deficiency.8 Several studies have shown that intensive training stimulates both circulating IGF-1 and IGFBP-3, since positive significant correlations exist between these factors and physical fitness levels.9-13 In contrast, an associated fall in IGFBP-3 related to overtraining was observed.14 These correlations indicate that fitness in healthy subjects is related to increased activity of the IGF system, favouring an anabolic state;<sup>15</sup> thus, it can be confirmed that physical exercise affects tissue anabolism and may provide general health benefits. The evaluation of possible changes in concentrations of total IGF-1 and its binding proteins (mainly IGFBP-3) may be of interest because it is thought that they affect performance, and probably reflect the physical overload state of athletes.<sup>9</sup><sup>16</sup>

However, data reviewed in the literature concerning the variation of IGF-1 and IGFBP-3 after exercise, that is to say in a dynamic situation, present conflicting opinions, depending on the type of exercise and the degree of training.<sup>4 17-22</sup> To our knowledge, no study to date has investigated the responses of these factors to exercise in relation to overtraining.

In this investigation, we report the concentrations of total IGF-1 and total IGFBP-3 in rugby players during a rest day and after an international rugby competition. Rugby is a physical activity characterised by mixed physiological

demands, but essentially comprising of anaerobic lactic type physiological responses.<sup>23</sup> <sup>24</sup> At a high level, this sport requires intensive training (10 h per week), and during the championship season a match occurs typically every week. The duration of a rugby match (90 min including a 10 min half time) pushes players to exhaustion, particularly in an exercise situation of maximal activity levels as reflected during an international contest. Therefore, the aim of our study was to examine if changes in IGF-1 and IGFBP-3 were associated so that fitness or overtraining could be measured; overtraining was estimated by using the overtraining questionnaire of the French Society of Sport Medicine (SFMS).<sup>25</sup> An English translation of this questionnaire is given in the appendix.

### **METHODS**

This study was conducted in a realistic situation of international competition on 11 international rugby players who belonged to the 17 member national team of Tunisia. These 11 athletes gave their consent to participate in the study following an explanation of experimental procedures and purpose and agreed to be sampled on two occasions by sport physicians. The study was carried out in compliance with ethical rules currently applied in Tunisia. No subject was taking any drugs or medication or reported a history of endocrine disorders before or during any stage of the study. None of these 11 players was substituted during the match. The players, aged 26.6 (1.4) years, had all participated actively in rugby for 16.5 (1.3) years, and trained in their clubs 3–4 days a week (8–12 h/week) in addition to their weekly 1.5 h match.

#### Anthropometric measurements

Anthropometric measurements included height, body mass, body mass index (BMI: weight in kg/height in m<sup>2</sup>) and

**Abbreviations:** AFC, after the competition; BMI, body mass index; CV, coefficient of variability; G1, group 1; G2, group 2; GH, growth hormone; IGF-1, insulin-like growth factor-1; IGFBP-3, insulin-like growth factor binding protein-3; SD, standard deviation; SFMS, French Society of Sport Medicine

percent body fat, estimated from skin fold thickness at the biceps, triceps, and subcapsular and subiliac sampling sites.<sup>26</sup> Height was measured using a Harpenden anthropometer (Holtain, Crosswell, Pembrokeshire, UK) to the nearest 0.1 cm. Weight was recorded to the nearest 0.1 kg with a portable digital metric scale, which was calibrated prior to data collection using standard weights. Skin folds were measured on each subject with a Harpenden skin fold calliper (Holtain). All the data were collected by the authors who were experienced in taking skin fold thicknesses.

For the assessment of  $Vo_{2max}$ , subjects performed a continuous incremental test to exhaustion on an athletic track according to the protocol of Léger and Boucher.<sup>27</sup>

#### Food intake

To assess the adequacy of nutrient intake, a 7 day consecutive dietary record was completed. All players received a detailed verbal explanation and written instructions on data collection procedures. Subjects were asked to continue with their usual dietary habits during the period of diet recording, and to be as accurate as possible in recording the amount and type of food and fluid consumed. A list of common household measures, such as cups and tablespoons, and specific information about the quantity in each measurement (grams, etc) was given to each participant. Each individual's diet was calculated using the Bilnut 4 software package (SCDA Nutrisoft, Cerelles, France), a computerised database that calculates food intake and composition from French standard references.

#### **Psychometric assessment**

The subjects were asked to complete a standardised overtraining questionnaire proposed by the French consensus group on overtraining (French Society for Sports Medicine, SFMS). This questionnaire was self administered 48 h before the match. It allows the calculation of a score that helps to classify, on a clinical basis, the demands of sportsmen submitted to a heavy training program.<sup>25</sup> This questionnaire consists of a list of 54 items requiring an answered response of "Yes" or "No" by the subject and selected from the reported clinical manifestations of the syndrome. The total of positive items (quoted "Yes") is used as a score of overtraining (see appendix).

#### Blood sampling and analytical methods

Blood samples for each subject were collected from a forearm vein during a rest day, 48 h before the competition (Rest) and 5–15 min after the competition (AFC). Rest sampling was performed at 8 am after an overnight fast and 9 h of sleep. Sampling after the match occurred at 3 pm, 5 h after the last meal. Each subject was instructed to maintain a supine position on a clinical couch during blood collection. The blood was centrifuged and serum was stored immediately at  $-30^{\circ}$ C until analysis.

	Whole group	G1 (n=6)	G2 (n = 5)
Age, year	26.6 (1.4)	28.3 (2.0)	24.6 (1.6)
Weight, kg	91.1 (3.5)	93.5 (5.6)	88.2 (3.8)
Height, cm	180.9 (1.6)	179.7 (1.8)	182.4 (2.8)
BMI, kg/m <sup>2</sup>	27.56 (1.0)	28.42 (1.5)	26.5 (1.2)
% Fat mass	19.5 (1.0)	20.5 (1.6)	18.2 (1.2)
Lean mass, kg	72.4 (2.4)	72.8 (4.0)	72.0 (2.7)

#### IGF-1

Serum concentrations of total IGF-1 were estimated using the Non-Extraction Insulin-Like Growth Factor-I IRMA Kit (Diagnostic Systems Laboratories, Webster, TX, USA). The theoretical sensibility, or minimum detection limit, calculated by interpolation of the mean plus 2 SD of 20 replicates of the 0 ng/ml IGF-1 standard, is 2 ng/ml. The inter-assay coefficients of variability (CVs) were 7.4 and 4.2, respectively, for the concentrations 32.54 and 383.86 ng/ml. The intra-assay CVs were 7 and 3.9, respectively, for the mean concentrations 34.03 and 373.86 ng/ml.

#### IGFBP-3

Serum concentrations of total IGFBP-3 were estimated using the Non-Extraction Insulin-Like Growth Factor Binding Protein-3 (IGFBP-3) Immunoradiometric Assay Kit (Diagnostic Systems Laboratories). The minimum detection limit, calculated as 2 SD from the mean of 22 replicates of the 0 ng/ml IGFBP-3 standard, is approximately 0.5 ng/ml. The intra-assay CVs were 1.8 and 3.9, respectively, for the mean concentrations 82.72 and 7.35 ng/ml. The inter-assay CVs were 1.9 and 0.6, respectively, for the mean concentrations 76.9 and 8.03 ng/ml.

#### Statistical analysis

Subjects were divided a posteriori into group G1 (n = 6) and group G2 (n = 5) following the response (positive or negative) of IGFBP-3 to the match.

Results are expressed as means (SE). The non-parametric Mann-Whitney test was used to test differences between variables within groups G1 and G2. Wilcoxon's test was used to test differences between variables within subjects (Rest  $\nu$  AFC). Simple correlations were performed to analyse relationships between biological, physical, and psychological variables. Analysis was performed using StatView software and the significance threshold was set at p<0.05.

#### RESULTS

All data are presented as results for the whole group (11 subjects) and for specific subjects presenting an increase (G1), or conversely, a decrease (G2), in IGFBP-3 levels.

The subjects' age and anthropometric characteristics are reported in table 1. Whatever the variable, there was no difference between G1 and G2.

Table 2 shows the data for  $Vo_{2max}$  results, overtraining score, and food intake. There was no difference in  $Vo_{2max}$  between G1 and G2. The mean overtraining score was 15.3 for the whole group. This score was higher (p<0.01) in G2 (21) than in G1 (10.5).

Nutritional assessment showed similar values for G1 and G2 for mean food intake. This intake is lower (-6%) than recommended (3373 kcal) for subjects with corresponding

	Whole group (n = 11)	G1 (n = 6)	G2 (n = 5)
Vo <sub>2max</sub> , ml/min per kg	58.4 (0.9)	56.9 (1.1)	60.1 (1.2)
Overtraining score Food intake	15.3 (2.3)	10.5 (0.6)	21.0 (3.5)**
kcal MJ	3148 (56) 13.16 (0.24)	3174 (66) 13.27 (0.28)	3116 (130) 13.03 (0.54)
Protein intake, g/kg per day	1.27 (0.04)	1.31 (0.03)	1.23 (0.09)



Figure 1 Concentration of serum total IGF-1 on the rest day (Rest) and just after competition (AFC). \*p<0.05, \*\*p<0.01, rest day v after competition values.

weights and activity levels.<sup>28</sup> The same is true for the protein intake.

The values of total serum IGF-1 (fig 1) during the rest day and just after the match are lower than the norms for adults of the same age. Compared to baseline levels, a significant increase of 42% in total IGF-1 serum concentrations was observed at the end of the match in all subjects (p<0.01).

In contrast, total serum IGFBP-3 values (fig 2) registered at the end of competition were not significantly different from the reference values when the whole group was considered. In fact, these values corresponded, in the first instance, to a significant increase in total serum IGFBP-3 levels recorded in six athletes (from 2911.66 (SE 170.9) ng/ml to 3540 (246.77) ng/ml, p<0.05), and in contrast, to a significant decrease in total serum IGFBP-3 levels observed in the other five athletes (from 3339 (641.35) ng/ml to 2932 (770.71) ng/ml, p<0.05). For this reason, we classified our subjects into the two subgroups outlined above.

The IGF-1/IGFBP-3 ratio increased significantly (p<0.05) when considering the whole group, but this variation resulted only from an increase in G2 (fig 3). The baseline ratio was also significantly lower in G2 than in G1.

Significant correlations (summarised in table 3) were found between the overtraining score and the resting IGF-1 levels, the IGFBP-3 levels measured after the match, and the  $\Delta$ IGFBP-3 (difference between resting and after match IGFBP-3 levels) (figs 4–6).

Considering figs 4–6, it appears that three subjects among the 11 presented a score >20. These three subjects had IGFBP-3 levels lower than 3000 ng/ml, while only one of them presented an IGF-1 level lower than 40 ng/ml. A nontired subject also had a similar low level of IGFBP-3. The three tired subjects only presented a >10% decrease in IGFBP-3 ( $\Delta$ IGFBP-3) between baseline and after competition values.

#### DISCUSSION

This study was intended to investigate the response to a competition of the somatotrope axis (IGF-1, IGFBP-3) in relation to states of tiredness. The subjects involved in the study were high level rugby players in the Tunisian national team and had the anthropometric and energetic characteristics required to participate in this type of activity at this level.<sup>29</sup> The competition was an international contest which required the participants to produce maximal efforts. When confronted with a maximal intensity task incorporating both physiological and psychological demands, the organism presents a global physiological response corresponding to the stress. The pituitary adrenal axis response to stress has been well documented,<sup>30 31</sup> but other pituitary hormones may also respond to the imposed stress, in particular GH. This hormone is important in relation to metabolic supply when



Figure 2 Concentrations of serum total IGFBP-3 on the rest day (Rest) and just after competition (AFC). \*p<0.05, rest day v after competition values.

energetic demand is increased, as in sports performance. GH secretion is characterised by pulses and by a short half life  $(\frac{1}{2}$  h), so that a single dosage is of little significance. For that reason, it is better to assay IGF-1 which is a hepatic relay of GH action (agreed that it may originate from other places) having a linear production and a longer half life (6 h), representing an integration of the changes in GH levels. In plasma, IGF-1 is linked to carrier proteins (IGFBP-6) to IGFBP-6). The biologically active fraction is the free fraction; thus, the level of free plasmatic IGF-1 depends not only on GH production, but also on the degree of linking to its carrier proteins, particularly to IGFBP-3 which has been shown to demonstrate some changes in response to exercise.<sup>32</sup>

It was reported that stress related cortisol secretion is associated with low IGF-1<sup>33</sup> and that total IGF-1 does not present a circadian rhythm.<sup>34</sup> Therefore, we determined resting values from morning samples collected 48 h before the match in order to avoid the precompetitive stress due to cognitive anticipation leading to increased cortisol levels.<sup>31</sup>

Similarly, the SFMS questionnaire was self administered 48 h before the match, since it aims to evaluate a general state of tiredness and not reaction to the competition. The score of 20 is only suggestive and not absolutely indicative of overtraining, like many other indexes (comportmental or biological), when taken alone.<sup>25</sup>

Although not significantly different, G1 and G2 were not perfectly matched: G1 was a little bit older and heavier than G2; the difference in body weight was accounted for by a higher percentage of fat mass in G1. Effectively, the two groups had very similar lean body masses. As they also had very similar food intakes, this difference in fat mass may be related to a natural trend linked to age. Whatever the cause, this did not seem to introduce a bias in the general conclusions.



Figure 3 Ratio of total IGF-1 to IGFBP-3 in rest day (Rest) and just after competition (AFC). \*p<0.05, rest day v after competition values. p<0.05, G1 rest v G2 rest.



Figure 4 Relationships between the overtraining score and the resting serum total IGF-1 level.

The main finding of our study is that we observed strong correlations between IGF-1, IGFBP-3, and the overtraining state as estimated by the scores of the SFMS questionnaire completed at the same time, and more particularly, that the more tired subjects presented a fall in IGFBP-3 after the match compared to resting values.

Participation in regular physical exercise generally increases serum concentrations of IGF-1.<sup>11–13</sup> Conversely, it has been observed that high intensity training or prolonged endurance activities provoke a long term decrease in IGF-1 levels.<sup>9 10 17 18 35</sup> It has also been established that reduced energy intake, or, more generally, when the energetic cost exceeds energy consumption, a fall in free IGF-1 levels occurs.<sup>36 37</sup>

We observed similar results since, in all subjects, both at rest and after the match, the total IGF-1 levels were lower (range: 100–310 ng/ml) than adult norms. This may be related to the food intake of these sportsmen which was lower (6%) than recommended.<sup>28</sup> Similar observations are frequently reported in sportsmen who probably limit their food intake to avoid overweight.<sup>38</sup> More particularly, the mean protein intake (1.27 (SE 0.04) g/kg per day) was slightly lower than recommended (1.5–2 g/kg per day) which may partly account for the low IGF-1 level.<sup>39</sup>

We also found, using the SFMS overtraining questionnaire, that the mean overtraining score was rather high at 15.3 (range: 6–25). A score higher than 20 strongly suggests excessive exercise load or overtraining.<sup>25</sup> Below this value, the score is correlated with training volume and intensity.<sup>40</sup> This intensive physical training combined with low energy intake may explain the low IGF-1 levels. This is illustrated by the negative significant correlation found between the resting total IGF-1 and the overtraining score.

The total IGF-1 values recorded immediately after competition were significantly higher than baseline. This increase in serum total IGF-1 is in agreement with previous findings,<sup>4 17 19 22</sup> activation of the GH/IGF-1 axis being a potentially favourable hormonal response promoting anabolic actions.<sup>4</sup> The great intensity and duration of exercise (during



Figure 5 Relationships between the overtraining score and the after competition serum total IGFBP-3 level.



Figure 6 Relationships between the overtraining score and IGFBP-3 level changes from baseline to after competition.

the rugby match) performed by the subjects in this study may explain the significant rise in serum total IGF-1. Physical exertion stimulates GH release into the circulation,<sup>17 41</sup> which induces short term increases in tissue IGF-1 production and leads to increased serum levels.<sup>17</sup>

In contrast to the findings of our study, other authors have shown unaltered<sup>4</sup><sup>21</sup> or decreased levels<sup>18</sup><sup>42</sup> of total serum IGF-1. This variability in exercise induced IGF-1 changes may be explained by the very different exercise protocols examined and by variation in individual subject profiles. Results from Nguyen *et al*<sup>20</sup> further illustrate the variability in IGF-1 response after exercise, as they reported a 12% increase, a 15% decrease, and no change in an incremental ergometer cycling exercise, in a long distance Nordic ski race, and in a treadmill simulated soccer game, respectively.

Free IGF-1 may be affected by the decrease in IGFBP-3 level, which is a compensatory mechanism allowing an increase in free IGF-1. It has been proposed that exercise induced IGFBP-3 proteolysis contributes significantly to the anabolic effects of exercise.<sup>15</sup> It has been documented that there was no change in IGFBP-3 after short (4–5 min) intense exercise,<sup>4</sup> but a decrease in IGFBP-3 was found after longer exercise.<sup>15 22</sup>

These low levels of IGFBP-3 seem to indicate excess training, suggesting a risk of overtraining. Some studies found a decrease in IGFBP-3 after 1 year of gymnastic training9 10 14 and negative correlations were found between IGFBP-3 and overtraining states in high level soccer and volleyball players, suggesting that a lowered IGFBP-3 level may reflect a reversal of the neuroendocrine adaptation to training.16 This is in agreement with the results from this study since we also found a strong negative correlation between the overtraining score and the serum levels of IGFBP-3 recorded just after the match. These low levels of IGFBP-3 may exert a protective mechanism against catabolism through an increase in the free IGF-1 fraction. Effectively, it was suggested that a high IGF-1 to IGFBP-3 ratio, as was the case in G2 after the competition, leads to a greater disponibility of IGF-1.43

Table 3      Relationships between the resting IGF-1 level, after competition IGFBP-3 level, IGFBP-3 changes, and overtraining score					
	Equation	r	р		
Rest level of IGF-1 v OS	y = 45.52 - 0.5088x; $r^2 = 0.425$	r=0.652	<0.05		
AFC level of	y = 43.05 - 0.0085x	r=0.824	< 0.01		

r = 0.767

A FC	attor compatition OS	overtraining score

r<sup>2</sup>=0.679 y=16.88-0.2708x;

 $^{2} = 0.593$ 

IGFBP-3 v OS

∆IGFBP-3 v OS

< 0.01

# What is already known on this topic

Several studies have shown that intensive training stimulates both circulating IGF-1 and IGFBP-3. However, an associated fall in IGFBP-3 related to overtraining has been observed.

## What this study adds

A strong negative correlations was found between IGF-1 and IGFBP-3 levels and overtraining as estimated by the SFMS questionnaire. In particular, levels of IGFBP-3 after exercise decreased in the fatigued subjects but were increased in fit subjects.

The more salient finding of the present study was that the IGFBP-3 levels increased in six rugby players and decreased in the five others after the match. This led us to classify our subjects into two subgroups. The first group (G1) included the six subjects who presented both elevated total serum IGF-1 and elevated IGFBP-3; these subjects exhibited low scores of overtraining (mean 10.5). The second group (G2) included the subjects whose IGFBP-3 levels decreased and who presented high scores (mean 21). Another approach was to consider the subjects whose score was >20 (overtrained subjects according to the SFMS guideline) and to examine the corresponding values of IGF-1, IGFBP-3, and ΔIGFBP-3 from baseline for after competition values. From this, it follows that  $\Delta$ IGFBP-3 is a very reliable index of overtraining. However, considering that the overtrained subgroup is rather small (only three subjects ), it is not possible to calculate a reliable sensitivity. In spite of this, we strongly suggest the use of this index in further surveys to confirm our finding.

Thus, the results of our study suggest that low levels of IGFBP-3 are linked to a state of tiredness. Moreover, in a dynamic situation such as a rugby match, IGFBP-3 level decreases in the more fatigued subjects. Therefore, we conclude that a fall of IGFBP-3 in response to an intense bout of exercise may represent an index of tiredness in highly trained sportsmen, as indicated by the scores obtained from the overtraining questionnaire.

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#### REFERENCES

- Stewart CEH, Rotwein P. Growth, differentiation, and survival: multiple physiological functions for insulin-like growth factors. Physiol Rev 1996;**76**:1005–26.
- Chevenne D. Les somatidines. Ann Biol Clin (Paris) 1991;49:69-91.
- Baxter RC. Insulin-like growth factor binding proteins in the human circulation: a review. Horm Res 1994;42:140–4. 3
- 4 Dall R, Lange KHW, Kjær M, et al. No evidence of insulin-like growth factor-binding protein 3 proteolysis during a maximal exercise test in elite athletes. J Clin Endocrinol Metab 2001;86:669–74.
- 5 Rosenfeld RG, Pham H, Cohen P. Insulin-like growth factor binding proteins and their regulation. Acta Paediatr 1994;83(suppl 399):61-7.
- 6 Cohen K, Nissley SP. The serum half-life of somatomedin activity: evidence for growth hormone-dependence. Acta Endocrinol 1976;83:243-58.
- Koistinen H, Koistinen R, Selenius L, et al. Effect of marathon run on serum IGF-I and IGF-binding protein 1 and 3 levels. J Appl Physiol 1996;80(3):760-4.

- 8 Thissen JP, Ketelslegers JM, Underwood LE. Nutritional regulation of the insulin-like growth factors. Endocr Rev 1994;15(1):80-101
- 9 Bouix O, Brun JF, Fédou C, et al. Exploration de gymnastes adolescents de classe sportive: quel suivi médical pour la croissance et la puberté? Sci Sport 1997;12:51-65
- 10 Filaire E, Jouanel P, Colombier M, et al. Effects of 16 weeks of training prior to a major competition on hormonal and biochemical parameters in young elite mnasts. J Pediatr Endocrinol Metab 2003;16(5):741-50.
- 11 Kelley PJ, Eisman JA, Stuart MC, et al. Somatomedin-c, physical fitness, and bone density. J Clin Endocrinol Metab 1990;70:718-23
- Koziris LP, Hickson RC, Groseth RT, et al. Serum levels of total and free IGF-I and IGFBP-3 are increased and maintained in long-term training. J Appl Physiol 1999;86:1436-42
- Poehlman ET, Copeland KC. Influence of physical activity on insulin-like 13 growth factor-I in healthy younger and older men. J Clin Endocrinol Metab 1990;**71**:1468–73.
- 14 Peyreigne C, Brun JF, Monnier JF, et al. Interactions entre la fonction somatotrope et l'activité musculaire. Sci Sport 1997;12:4-18.
- 15 Rosendal L, Lanberg H, Flyvbjerg A, et al. Physical capacity influences the response of insulin-like growth factor and its binding proteins to training. J Appl Physiol 2002;**93**(5):1669–75
- 16 Aïssa Benhaddad A, Bouix D, Khaled S, et al. Early hemorheologic aspects of overtraining in elite athletes. Clin Hemorheol Microcirc 1999;**20**:117–25.
- Chicharro J, Lopez A, Hoyos J, *et al.* Effects of an endurance cycling competition on resting serum insulin-like growth factor I (IGF-I) and its binding proteins IGFBP-1 and IGFBP-3. Br J Sports Med 2001;**35**:303–7. 17
- 18 Eliakim A, Brasel JA, Mohan S, et al. Increased physical activity and the growth hormone-insulin-like factor-I axis in adolescent males. Am J Physiol 1998;**275**:R308–14.
- Kraemer WJ, Aguilera BA, Terada M, et al. Responses of IGF-I to endogenous increases in growth hormones after heavy-resistance exercise. J Appl Physiol 1995;**79**:1310–5.
- 20 Nguyen UN, Mougin F, Simon-Rigaud ML, et al. Influence of exercise duration on serum insulin-like growth factor and its binding proteins in athletes. *Eur J Appl Physiol* 1998;**78**:533–7.
- Nindl BC, Kraemer WJ, Marx JO, et al. Overnight responses of the circulating IGF-I system after acute, heavy-resistance exercise. J Appl Physiol 2001:90:1319-26
- Schwarz AJ, Brasel JA, Hintz RL, et al. Acute effect of brief low- and high intensity exercise on circulating insulin-like growth factor (IGF) I, II, and IGF-binding protein-3 and its proteolysis in young healthy men. J Clin Endocrinol Metab 1996;**81**:3492–7.
- Doutreloux JP, Tepe P, Demont M, et al. Exigences énergétiques estimées selon les postes de jeu en rugby. Sci Sports 2002;17:189–97.
- 24 McLean DA. Analysis of the physical demands of international rugby union. J Sports Sci 1992;10:285–96.
- **Brun JF**. The overtraining: to a system of evaluation usable by routine examination. *Sci Sport* 2003;**18**:282–6. 25
- Durnin JVGA, Rahaman MM. The assessment of the amount of fat in the human body from measurement of skinfold thickness. Br J Nutr 26 1967;**21**:681-9
- 27 Léger LA, Boucher R. An indirect continuous running multistage field test: the University of Montreal track test. Can J Appl Sport Sci 1980;5(2):77–84.
- Martin A. Apports nutritionnels conseillés pour la population française. Paris: Editions Tec & Doc, 2001:27–36. 28
- 29
- Editions Tec & Doc, 2001:27–30. Maso F, Cazorla G, Godemet M, et al. Physiological features of rugby players of the French team. Sci Sports 2002;17:297–301. Elloumi M, Maso F, Michaux O, et al. Behaviour of saliva cortisol (C), testosterone (T) and the T/C ratio during a rugby match and during the post-competition recovery days. Eur J Appl Physiol 2003;90:23–8. Passelergue P, Robert A, Lac G. Salivary cortisol and testosterone variations during an official and a simulated unight lifting acmedition. Int J Sport Mad 30
- 31 during an official and a simulated weight-lifting competition. Int J Sport Med 1995;16:298-303
- Manetta J, Brun JF, Maimoun L, *et al.* The effects of intensive training on insulin-like growth factor I (IGF-I) and IGF binding proteins 1 and 3 in 32 competitive cyclists: relationships with glucose disposal. J Sports Sci 2003;21(3):147-54.
- Rosmond R, Dallman MF, Bjorntorp P. Stress-related cortisol secretion in men: relationships with abdominal obesity and endocrine, metabolic and hemodynamic abnormalities. J Clin Endocrinol Metab 1998;83(6):1853-9.
- Heuck C, Skjaerbaek C, Orskov H, et al. Circadian variation in serum free 34 ultrafiltrable insulin-like growth factor I concentrations in healthy children. *Pediatr Res* 1999;**45**:733–76.
- Tigranian RA, Kalita NF, Davydova NA. Observations on the Soviet/ Canadian transpolar ski trek: status of selected hormones and biologically 35 active compounds. Med Sci Sports Exerc 1992;33:106-38.
- Suikkari AM, Sane T, Seppala M, et al. Prolonged exercise increases serum insulin-like growth factor-binding protein concentrations. J Clin Endocrinol Metab 1989;**68**:141–4.
- Nemet D, Connolly PH, Pontello-Pescatello AM, et al. Negative energy 37 balance plays a major role in the IGF-I response to exercise training. J Appl Physiol 2004;**96**(1):276–82.
- Ebine N, Rafamantanantsoa HH, Nayuki Y, *et al.* Measurement of total energy expenditure by the doubly labelled water method in professional soccer 38 players. J Sports Sci 2002;20(5):391-7.
- Smith WJ, Underwood LE, Clemmons DR. Effects of caloric or protein restriction on insulin-like-growth-factor (IGF-1) and IGF-binding proteins in Children and adults. J. Clin Endocrinol Metab 1995;80(2):443–9.
   Maso F, Lac G, Michaux O, Robert A. Le questionnaire de surentraînement:
- 40 corrélations avec le cortisol et la testostérone dans le cadre du suivi d'une équipe de haut niveau en rugby. Sci Sports 2003;18:169-71.

## IGFBP-3, a sensitive marker of physical training and overtraining

- Felsing NE, Brasel JA, Cooper DM. Effect of low- and high-intensity exercise on circulating growth hormone in men. J Clin Endocrinol Metab 1992;75:157–63.
  Jahreis G, Kauf E, Frohner G, et al. Influence of intensive exercise on insulin-like growth factor I, thyroid and steroid hormones in female gymnasts. Growth Regul 1991;1:95–9.
- 43 Juul A, Dalgaard P, Blum WF, et al. Serum levels of insulin-like growth factor (IGF)-binding protein-3 (IGFBP-3) in healthy infants, children, and adolescents: the relation to IGF-I, IGF-II, IGFBP-1, IGFBP-2, age, sex, body mass index, and pubertal maturation. J Clin Endocrinol Metab 1995;80(8):2534–42.

# **APPENDIX**

ex	Μ	F	
ate of birth			
Vhat is your profession?	V	NL.	
you are a student, are you doing examinations?	řes	INO	
In a many hours do you practice per week?	4_9	8_10	Mara than 10
wou play other sports or games write them:	0-0.	0-10	More man to
his month has there been any significant event which may have disturbed	Yes	No	
our private or professional life?	103	140	
his month			
1–My level of sports performance/my general form has decreased	Yes	No	
2–I am not as attentive as before	Yes	No	
3–My close friends think that my behaviour has changed	Yes	No	
4–I have a sensation of pressure in my chest	Yes	No	
5 –My heart seems to beat faster	Yes	No	
6–I have a lump in my throat	Yes	No	
7–I have less appetite than before	Yes	No	
8-l eat more	Yes	No	
9–1 do not sleep as well as before	Yes	No	
10–I drowse and yawn in the daytime	Yes	No	
11-The time between training sessions seems to me too short	Yes	No	
12–My sexual libido has decreased	Yes	No	
13–My performances are poor	Yes	No	
14-I trequently catch a cold	Yes	No	
15-I have put on weight	Yes	No	
16-I have memory problems	Yes	No	
I/-I offen feel fired	Yes	No	
18-I underestimate myself	Yes	No	
19–1 offen have cramps, muscular pain	Yes	No	
20-I suffer from headaches more frequently	Yes	INO N I	
21 - I do not reel filt	tes	INO N.I.	
22-I sometimes reel dizzy, on the point or rainting	res	INO Na	
23-I do not contide in others so easily	res	INO Na	
25 I have a care threat more after	Yee	NO Na	
25-I have a sore mirodi more ofien	Vec	No	
27-I do not boar training so well	Vos	No	
28-At rest my heart rate is faster than before	Yes	No	
29-During exercise my heart rate is faster than before	Yes	No	
30-Loften feel rotten	Yes	No	
31-I get tired more easily	Yes	No	
32-I often have digestive disorders	Yes	No	
33–I feel like staving in bed	Yes	No	
34-I am not so confident in myself	Yes	No	
35-I get injured more easily	Yes	No	
36–I have more difficulties in organising my thoughts	Yes	No	
37–I have more difficulties in concentrating in my sports activity	Yes	No	
38–My sporting gestures are less precise, less skilful	Yes	No	
39–I have lost force and aggressiveness	Yes	No	
40-I feel as if I have no one to talk to	Yes	No	
41–I sleep longer	Yes	No	
42–I cough more often	Yes	No	
43–I do not enjoy practising my sports as much	Yes	No	
44–I do not enjoy my leisure activities as much	Yes	No	
45–I get irritated more easily	Yes	No	
46–I am less efficient in my school or professional activity	Yes	No	
47–People around me think that I have become less pleasant	Yes	No	
48–Training seems harder and harder	Yes	No	
49-It is my fault if my results are worse	Yes	No	
50–My legs teel heavy	Yes	No	
51–I lose my personal things more easily (wallet, keys, etc)	Yes	No	
52–I am pessimistic, I have the blues	Yes	No	
53–I have lost weight	Yes	No	
54–My motivation, will and tenacity are weaker	Yes	No	
ut a cross to range between these two opposite states			
My physical level			
Great tormBad form			
I teel tatigued			

Continued			
I recover from my state of tiredness			
More quicklyMore slowly			
Very relaxedVery anxious			
I feel that my muscular strength has			
IncreasedDecreased			
I feel that my endurance has			
IncreasedDecreased			
Have you had any difficulties in understanding some of the questions?	Yes	No	
If Yes, which questions did you find difficult to understand (write the numbers)			

Each "Yes" answer receives one point, giving a score of between 0 and 54. According to the French multicentric study (F Maso, in press) scores >20 are highly suggestive of overtraining and are constantly found in overtly overtrained individuals.

## Sports Physician required for LTA Academies

Doctors with experience in Sport and Exercise Medicine (minimum 5 years) and a passion for tennis are required for one session per week at the LTA academies. Two positions are available (Bath and Loughborough), starting in September 2005.

In addition to providing medical support for the LTA Academy players (aged 13–16 years), the appointed doctor will be required to act as a tournament medical officer (Nottingham, Birmingham, Eastbourne, Bournemouth). This would involve no more than 1 week—twice per annum—for each doctor.

The appointed doctors will be involved in coach education and be required to attend monthly meetings at the LTA National Centre (Queens Club or Roehampton). They will also be expected to ensure continued professional development in the field of "tennis medicine" (e.g. attendance at seminars, workshops, and conferences focussing on injury prevention, racket sports, etc).

Remuneration and expenses would be at rates no less than those currently paid by the British Olympic Association, UKSport, and the EIS.

Each applicant should submit the following documentation: complete Curriculum Vitae, a copy of the current GMC Registration Certificate, a certificate of medical malpractice indemnity, and a Child Protection Enhance Disclosure Certificate.

All correspondence should be sent to: Dr Michael Turner Chief Medical Adviser The Lawn Tennis Association Queens Club London W14 9EG Email: michael.turner@LTA.org.uk