SUDDEN DEATH OF ATHLETES:
IS IT DUE TO LONG-TERM CHANGES IN SERUM MAGNESIUM, LIPIDS AND BLOOD SUGAR?

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ABSTRACT

In young, apparently healthy, trained Israeli men, strenuous effort was reported to give rise to persistent magnesium (Mg) deficiency and a parallel long-term increase of cholesterol, triglycerides and blood sugar /1-3/.

The relationship of Mg deficiency to the pathogenesis of cardiovascular disease has been increasingly documented during the last decade. Several authors have highlighted the phenomenon of sudden deaths in sport and have suggested that it is associated with cardiovascular disease.

The association is discussed between Mg deficiency and increase of blood lipids and sugar, found as a sequel to strenuous effort, and cardiovascular morbidity and mortality risk reported in athletes.

It is postulated that sudden death of athletes and other intensely training individuals during exertion, is mediated by the deleterious cardiovascular effects of persistent magnesium deficiency and the resultant hyperlipaemia and hyperglycaemia, which, as we have documented, follows strenuous effort.

KEY WORDS

magnesium deficiency, exertion, athletes, sudden death

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INTRODUCTION

A number of studies showed that in young, trained, apparently healthy, Israeli men, as a sequel of strenuous effort, there was a persistent decrease of serum magnesium concentration (S-Mg) /1-4; Stendig-Lindberg, this issue/. A delayed 30% increase of triglycerides (p < 0.001) was observed in parallel, 3 days after the termination of strenuous effort (a 120 km brisk hike). This increase coincided with the peak decrease of S-Mg /1/. In addition, a significant increase of blood sugar (p < 0.001) was seen, immediately after the termination of the strenuous effort, which coincided with the first peak of the biphasic decrease of S-Mg.

The only previous report of an increase of triglycerides following effort was that of Sutherland et al. /5/ in 20-55 year-old men after four months of training culminating in a marathon run.

As regards the increase of blood sugar, Naven et al. /6/ reported a rise of blood sugar after strenuous intermittent running. However, it has been generally accepted that blood sugar and lipids decrease after effort. For this reason, further studies were undertaken to examine the long-term effect of strenuous effort on intermediary metabolism in young, trained, apparently healthy, Israeli men. The results are briefly summarized, to form an experimental basis for the proposed hypothesis. The relevant methodology is detailed in /1-3/.

In the first study, twenty men were examined (mean age 19.3, S.D. 1.4) following a 70 km brisk hike /2/. The results are shown in Figs. 1, 2 and 3. Serum triglycerides began to rise 24 h after termination of strenuous effort and were significantly raised 17 days later. Blood sugar, which decreased at 24 h after the hike, was significantly increased at day 18. Mean serum cholesterol, which was seen to decrease gradually, reaching the lower border of the reference value at 72 h after termination of the 70 km march, was also highly significantly raised on day 18. No bouts of strenuous effort took place within the 18 days after the termination of the march. Since the increase was delayed, it is possible that the generally accepted impression of a decrease of blood sugar and lipids occurring as a result of strenuous effort, may have been due to an insufficiently long follow-up.

In the second study, 15 volunteer soldiers (mean age 18.6, S.D. 1.3 years) were examined 6 and 10 months after a 120 km brisk march.
**Fig. 1:** Serum triglycerides before and after 70 km brisk hike. The horizontal upper line represents the upper border of the reference range. p values represent the probability in comparison with the baseline value. NS = Non-significant.

<table>
<thead>
<tr>
<th>Time</th>
<th>Serum triglycerides (mmol/L)</th>
<th>n</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>before</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>after 1 hr</td>
<td></td>
<td>20</td>
<td></td>
</tr>
<tr>
<td>after 24 h</td>
<td></td>
<td>16</td>
<td></td>
</tr>
<tr>
<td>after 72 h</td>
<td></td>
<td>19</td>
<td></td>
</tr>
<tr>
<td>after 18 days</td>
<td></td>
<td>20</td>
<td></td>
</tr>
<tr>
<td>70 Km march</td>
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<td></td>
<td></td>
</tr>
</tbody>
</table>

**Fig. 2:** Blood sugar before and after 70 km march. The shaded area represents the reference range. p values represent the probability in comparison with the baseline value. NS = Non-significant.

<table>
<thead>
<tr>
<th>Time</th>
<th>Blood Sugar (mmol/L)</th>
<th>n</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>before</td>
<td></td>
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</tr>
<tr>
<td>70 Km march</td>
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<td>after 1 hr</td>
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</tr>
<tr>
<td>after 18 days</td>
<td></td>
<td>20</td>
<td></td>
</tr>
</tbody>
</table>
Fig. 3: Serum cholesterol before and after 70 km brisk hike. The shadowed area represents the reference range. $p$ values represent the probability in comparison with the baseline value. NS = Non-significant.

Fig. 4: Serum triglycerides before and after 120 km march. The upper horizontal line represents the upper border of the reference range. $p$ represents the probability in comparison with the baseline value. NS = Non-significant.
Figs. 4, 5 and 6 show a gradual, although non-significant, rise of blood sugar and lipids over time. In the cases of blood sugar and triglycerides, this was preceded by a decrease, a phenomenon which may have been the cause of the usually reported decrease of the latter following exertion.

Although the subjects carried out daily training (not exceeding 4000 kcal [16.7 mJ] energy expenditure), no bouts of comparable strenuous effort took place between the termination of the forced marches and the end of the observation period, in either study.

In the third and last study, 16 volunteer soldiers (mean age 18.7, S.D. 0.6 years) were followed for 11 months after a 120 km brisk march (with no further bouts of comparable strenuous effort in the interim period) /3/. The results are shown in Figs. 7-9. Serum triglycerides show a biphasic rise; at 11 months after the effort, the mean lies at the upper border of the reference range. Serum cholesterol rises, beginning 9 months after the effort, to become significantly raised 11 months afterwards, while blood sugar decreases after a significant rise at 9 months.

A rise in cholesterol and the HDL-cholesterol fraction has been reported in man in treadmill exercise until exhaustion /7/. This was, nevertheless, interpreted as a beneficial effect of effort on the lipid profile, due to the increase in the HDL-cholesterol fraction.

We did not carry out lipoprotein electrophoresis, but nonetheless the findings in the three studies show a consistent trend of increase in lipid levels. This suggests a deleterious effect of strenuous effort on the lipid profile which, together with the tendency to an increase in blood sugar, could pose a health hazard to large sections of populations engaged in strenuous effort (e.g. manual workers, soldiers). This hazard will be increased further by exertion at high temperatures (hot climate, factory conditions, etc.) due to the additional loss of magnesium in sweat /1, 2, 8, 9/.

THE CONTRIBUTION OF Mg DEFICIENCY TO LONG-TERM CHANGES IN INTERMEDIARY METABOLISM AFTER EXERTION

Mg is essential for intermediary metabolism /10-14/. In the glycolytic pathway, 7 out of 13 steps are Mg-dependent and in the tricarboxylic acid cycle (Krebs' cycle) 3 out of 9 steps are Mg-dependent, including the initial conversion of pyruvate to acetyl-CoA through activation by Mg\(^{2+}\) of pyruvate dehydrogenase /8/.
Fig. 5: Blood sugar before and after 120 km march. The horizontal lines represent the reference interval. p represents the probability in comparison with the baseline value. NS = Non-significant.

Fig. 6: Serum cholesterol before and after 120 km march. The horizontal lines represent the reference interval. p represents the probability in comparison with the baseline. NS = Non-significant.
Fig. 7: Serum triglycerides before and after 120 km march. The upper horizontal line represents the upper border of the reference range. p represents the probability in comparison with the baseline. NS = Non-significant.

Fig. 8: Blood sugar before and after 120 km march. The horizontal lines represent the reference interval. p represents the probability in comparison with the baseline. NS = Non-significant.
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Fig. 9: Serum cholesterol before and after 120 km march. The horizontal lines represent the reference interval. $p$ represents the probability in comparison with baseline. NS = Non-significant.

### TABLE 1
Type of activity during which sudden death was reported in 145 athletes /19, 20/

<table>
<thead>
<tr>
<th>Type of sport activity</th>
<th>No.</th>
<th>%</th>
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<tbody>
<tr>
<td>Marathon running</td>
<td>55</td>
<td>38%</td>
</tr>
<tr>
<td>Football</td>
<td>34</td>
<td>23%</td>
</tr>
<tr>
<td>Squash</td>
<td>30</td>
<td>21%</td>
</tr>
<tr>
<td>Golf, sailing, mountain climbing</td>
<td>13</td>
<td>9%</td>
</tr>
<tr>
<td>Basketball</td>
<td>9</td>
<td>6%</td>
</tr>
<tr>
<td>Tennis</td>
<td>4</td>
<td>3%</td>
</tr>
</tbody>
</table>

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Animal experiments show that the plasma activity of lecithin-cholesterol acetyltransferase (LCAT), which is needed to esterify cholesterol, is decreased in magnesium deficiency /15/. This decrease may cause the lowering of cholesterol esters, which bind to the HDL-cholesterol fraction. The resultant decrease would impair the breakdown of triglycerides. In addition, lipoprotein lipase activity may be impaired in magnesium deficiency causing hypercholesterolaemia /16/. Although research on the effect of Mg deficiency on glucose and lipid metabolism has been scanty hitherto in man, evidence derived from animal experiments allows us to extrapolate that Mg deficiency due to strenuous effort would impair the breakdown of blood glucose and lipids. Hormonal changes seen during exercise, including a rise of catecholamines, thyroid stimulating hormone, glucagon, and corticosteroids and a decrease of plasma insulin, which favour mobilization of fatty acids and hepatic glucose /17/, may enhance the demand for Mg during strenuous effort, resulting in a vicious circle. This is even more likely to occur when dietary Mg intake is inadequate, as, e.g., found in Israel /18/.

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Northcote and Ballantyne analysed the type of activity during which death occurred in 145 reported cases of sudden death in athletes /19, 20/; see Table 1.

According to the literature, the most common cause of sudden death in sport is cardiac arrest. According to Northcote and Ballantyne /19/, 14% of cases are due to structural abnormalities such as hypertrophic myopathy or congenital anomaly, 7% are due to myocarditis, 2% to fatal arrhythmias and 73% to coronary artery disease. The latter is the most frequent cause of sudden death in sport. In the study of Thompson et al. /22/, all deaths while jogging in Rhode Island during 1975-1980 were due to coronary artery disease.

Mg DEFICIENCY, HYPERGLYCAEMIA AND HYPERLIPIAEMIA IN STRENUOUS EFFORT AND SUDDEN DEATH IN ATHLETES

Mg regulates smooth muscle contraction /23/ and its deficiency favours vasoconstriction. The latter, occurring in the coronary vessels, predisposes to coronary disease /24/. The relationship between
Mg, mineral content of drinking water and cardiovascular morbidity was first reviewed by Marier et al. [25]. Since then, an increasing body of literature has dealt with the role of magnesium in the pathogenesis of ischaemic heart disease.

Coronary disease was reported to be the most frequent cause of sudden death in sports [21]. In 33% of the cases of sudden death in sport, the prodromal complaints were chest pain of anginal type and fatigue [21, 26]. Fatal ventricular fibrillation was reported as one of the causes of death in strenuous exercise [21]. The association of Mg deficiency and cardiac arrhythmia was first documented in man by Chadda et al. [27], and experimental animal research has shown that Mg facilitates defibrillation [28]. Finally, in experimental animals, Mg deficiency has recently been shown to be implicated in the pathogenesis of arteriosclerosis [29].

In subjects engaged in strenuous effort, therefore, the deleterious effect of Mg deficiency will compound the already extensively documented negative effects of hyperlipaemia and hyperglycaemia on the cardiovascular system; together, they will enhance the risk of cardiovascular morbidity and potential mortality.

The increased cardiovascular morbidity of exertion, mediated by Mg deficiency and raised blood lipids and sugar, may give rise both to the “burn out” of athletes, as well as to an increased risk of occurrence of coronary disease, myocardial infarction and cardiac arrhythmias, which, in turn, predispose to the risk of sudden death [21]. The dramatic decrease of S-Mg found to follow strenuous exertion [1, 2], together with a pre-existing Mg deficiency and the accompanying metabolic changes [1, 2, 3], may at one point in time serve as a trigger for cardiovascular sudden death in exposed individuals, e.g. athletes or elite soldiers during strenuous effort.

REFERENCES