the prognosis by reducing the excess cardiovascular mortality in diabetic nephropathy.

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Local bone mineral response to brief exercise that stresses the skeleton

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Abstract

Objective-To compare grip strength and bone mineral content in the forearm in women and to test the effects on bone mineral content of short periods of exercise that stresses the skeleton.

Design-Assessment of both wrists in 69 volunteers and of the non-fractured wrist in 30 patients followed by an exercise regimen entailing squeezing a tennis ball as hard as possible for 30 seconds each day for six weeks.

Setting-Old people's homes and outpatient departments of Hammersmith and Northampton general hospitals.

Patients - 99 Women, of whom 69 were volunteers and 30 had a fractured forearm.

Main outcome measure-Grip strength and bone mineral content after six weeks and at six months after the exercises had stopped.

Results-The bone mineral content of the women's forearms was measured with a densitometer and the grip strength with a semi-inflated bag connected to an anaeroid barometer. Measurements before exercise showed that the two variables correlated closely, irrespective of age, and that there were significant differences in both between the dominant and non-dominant arms of the volunteers. After six weeks of exercise there was a mean increase in grip strength of 14.5% (95% confidence interval 9.9 to 19.2%) and in bone mineral content of 3.4% (1.4 to 5.3%) in the stressed forearms of the 77 women who attended for examination. After six months without exercise the improvements in the 33 women who attended for follow up had reversed. Women who had had a fractured forearm (n=13), however, had continued to gain grip strength and bone

mineral content in the arm that had not been injured

Conclusions-Grip strength in the forearm is a good indicator of bone mineral content. Both variables may be increased by brief periods of stressful exercise. If this principle can be applied to the whole skeleton it may provide a means of reversing osteoporosis.

Introduction

Physical activity is commonly accepted to have some role in preventing and treating osteoporosis,¹⁻³ but the role is unclear.⁴ Female marathon runners, for example, may become osteoporotic,5 and infantry recruits subjected to severe exercise developed either hypertrophy of the tibia or stress fractures.6 Pathological bone remodelling was shown by Wolff nearly a century ago,⁷ and Nilsson and Westlin showed that localised changes in bone density in athletes were related to the stresses put on their arms and legs during their particular sport.8 Rubin and Lanyon produced an osteogenic response to short periods of abnormal loading in the functionally isolated wing bone of a rooster.9

Patients with fractures of the arm or leg commonly develop a localised osteoporosis, which reverses as use returns. We examined the relation between bone mineral content in the forearm and the peak stress, equivalent to maximum grip strength, commonly experienced by the forearm and tried to modify grip strength in female volunteers and women with a fractured forearm.

Subjects and methods

We tested the grip strength of 99 women (69 volunteers and 30 patients with a fractured forearm)

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with a semi-inflated (at 40 mm Hg) bag connected to an anaeroid barometer with a scale of 0-600 mm Hg. The bone mineral density of their wrists was measured with a Molsgaard BDS1100A single photon absorption densitometer.¹⁰ This instrument measured absorption of photons of iodine-125 by the forearm in a water bath and was calibrated for decay of the isotope twice daily. The bone mineral content was given in arbitrary units. Data corrected for the amount of fat and bone width of the women's forearms were ignored as these were unlikely to vary during the brief experiment.

The exercise regimen required each woman to squeeze a tennis ball as hard as possible three times consecutively, morning and evening, every day for six weeks; this took less than 30 seconds each day. The 69 volunteers had a mean (SD) age of 62 (16) years. They were randomly allocated to stress one forearm (34 to stress the right forearm and 35 to stress the left) while the other arm remained unexercised as a control. Their grip strength and bone mineral content were measured at six weeks. The 30 patients had a mean age of 53 (21) years; they stressed their uninjured arm (15 stressed the right arm and 15 the left). Their grip strength and bone mineral content were measured at three and six weeks.

Six months after the exercise regimen had ended the women were asked to return at short notice for further measurements. Initial measurements were taken as 100%, and paired t tests of changes from these were used throughout, as in other studies.¹¹ Statistical analysis was carried out by a Minitab program.

Results

In the 99 women examined both bone mineral content in the forearm and grip strength decreased with age (r=-0.746, p<0.001 and r=-0.645, p<0.001 respectively). Grip strength and bone mineral content before exercise correlated closely (r=0.659, p<0.001; fig 1). Multiple regression analysis of bone mineral content against both age and grip strength gave the equation: bone mineral content = $44.2 - (0.289 \times \text{ age}) + (0.0285 \times \text{ grip strength})$. The corresponding standard errors were 0.045 for age and 0.008 for grip strength. Grip strength, irrespective of age, is therefore a good predictor of bone mineral content in the forearm (p<0.001). Age and grip strength together accounted for 60.8% of the variance in bone mineral content.



FIG 1 — Correlation between bone mineral content (arbitrary units) and grip strength (mm Hg) before exercise in forearms of 99 women

Among the 69 volunteers there were significant differences between the dominant (65 right and four left) and non-dominant arms for both grip strength (11.9%, p<0.0003; 95% confidence interval 5.8 to 18.1%) and bone mineral content (2.7%, p<0.02; 0.5 to 4.9%).

The 77 women who were examined at six weeks had

a 14.5% (p<0.0001, 95% confidence interval 9.9 to 19.2%) improvement in grip strength and a 3.4% (p<0.001, 1.4 to 5.3%) improvement in bone mineral content in the exercised forearm. The 57 volunteers who reattended at six weeks had a mean gain of 12.5% (p<0.001, 6.6 to 18.4%) in grip strength and 3.1% (p<0.02, 0.6 to 5.6%) in bone mineral content in the exercised forearm (fig 2). There was also a rise in grip strength of 7.4% (p<0.002, 2.8 to 12.0%) that was associated with a non-significant rise in bone content of 1.9% (p<0.13, -0.6 to 4.3%) in the unexercised arm; several women, however, admitted to having stressed both forearms.



FIG 2—Percentage changes in grip strength and bone mineral content in exercised and unexercised forearms of 69 volunteers. Bars are 95% confidence intervals

Women with a fracture were seen at three and six weeks. The 27 who attended at three weeks had a $12\cdot3\%$ improvement in grip strength (p<0.001, 5.9 to $18\cdot8\%$) with a 4.8% rise in bone mineral content (p<0.0005, 2.4 to 7.1%). The 20 who also returned at six weeks had a 20.4% (p<0.0001, 14.0 to 26.8%) gain in grip strength and a 4.0% (p<0.005, 1.5 to 6.5%) increase in bone mineral content (fig 3).



FIG 3—Percentage changes in grip strength and bone mineral content in the uninjured forearm of 30 patients with a fractured forearm. Bars are 95% confidence intervals

Women with or without a fracture who had gained more than 50 mm Hg in grip strength after six weeks of exercise, (the "best exercisers") were analysed separately. Their 35.7% (p<0.0001, 29.1 to 42.3%) gain in strength was associated with a 4.7% (p<0.04, 0.2 to 9.3%) gain in bone mineral content (fig 4).

In all, 33 women attended a recall six months after the exercise ended (figs 2-4). The volunteers (n=20) had lost most of their gains (grip strength 5.1% (p<0.3, -4.7 to 14.9%) and bone mineral content -2.6% (p<0.1, -5.6 to 0.52%) of their initial measurements). Women who had had a fracture (n=13), however, had continued to gain strength (30.9%, p<0.003; 12.8 to 49.0%) and bone mineral



FIG 4—Percentage changes in grip strength and bone mineral content in the exercised arm for women who had a gain in grip strength of >50mm Hg at six weeks. Bars are 95% confidence intervals

content (13.3%, p<0.02; 2.6 to 24.0%) in their uninjured arm. Comparison of both forearms at six months showed that the exercised arm had 29.1% (p<0.004, 11.5 to 46.7%) greater grip strength and 4.9% (p<0.14, -1.9 to 11.7%) higher bone mineral content than the arm that had been injured. The best exercisers (n=10) conserved some of their increases in strength and bone density at six months, but neither was significant.

The results at six weeks for the 33 women who were seen at six months showed a 20.6% (p<0.0001, 13.5 to 27.7%) improvement in grip and a 3.4% (p<0.036, 0.2 to 6.5%) improvement in bone mineral content in the exercised arm; their initial grip strength was 261 mm Hg (231 to 290 mm Hg) and bone mineral content 36.9units (33.9 to 39.9 units). The initial grip strength in the exercised arm of the 66 women who did not reach the end of the study was 275 mm Hg (249 to 301 mm Hg) and their initial bone mineral content 33.9 units (31.4 to 36.4 units). Those who did not attend at six months were older (mean age 60.5 years) than those who did (mean age 56.8 years).

Discussion

As expected, our results showed that both grip strength and bone mineral content in the forearm fall with age, but we also found a close correlation between the two variables irrespective of age. In addition, the small but significant initial difference in both variables between the dominant and non-dominant arms of the 69 volunteers supports our hypothesis that a forearm's bone mineral content is related closely to the physical demands made on it by activities that require grip strength. We had assumed that a grip of maximum power stresses the skeleton of the forearm as much as any other activity that is performed commonly. We had not expected the change in grip strength in the unexercised arm, which was associated with a nonsignificant increase in bone mineral content. There was, however, insufficient emphasis in our instructions to prevent volunteers exercising both forearms, and several admitted to doing this.

We had also assumed that after a Colles' fracture the uninjured forearm is used more. This may be wrong: a manual worker might avoid physical stress when off work, whereas an independent retired person might be obliged to undertake more stressful activity with the intact arm. Women with fractures achieved a significant increase in bone mineral density by three weeks. Several of those with minor fractures were out of plaster before six weeks and were unlikely to attend hospital again, which may account for the non-linear results over three to six weeks. Similarly, frail volunteers were less likely to attend follow up clinics, and it was not feasible to retest some of those in old people's homes at six months. Those who attended six months after the exercise regimen ended were therefore likely to be the more motivated women who had originally done well with the exercises.

As numbers of women fell with succeeding visits the mean strength and bone mineral content rose. Measure-

ments at six months showed a loss of the previous gains in grip strength and bone mineral content in all of the women except those who had had a fracture. It may be that after a fracture patients with apparently normal function continue to protect the injured arm from peak stresses by using the uninjured wrist for activities that require a powerful grip. They may thereby continue to increase both grip strength and bone mineral content in the non-fractured arm.

Changes in grip strength were mirrored by changes in bone mineral content at all stages, and this supports our contention that mineral content reflects quickly the physical demands made on bone. Muscle may be trained for great endurance without bulk by prolonged exercise, as in marathon runners, or for strength and bulk by isometric exercise, as in weight lifters. A similar analogy might apply to bone, in which short periods of skeletal stress may trigger osteogenesis more effectively than hours of gentler exercise.

Several questions remain—for example, Can this principle be applied to the entire skeleton? and What is the minimum daily requirement of skeletal stress? A few seconds of grip exercise each day may stress the forearm skeleton sufficiently to stimulate a local gain in bone mineral content. The best method of regularly stressing the whole skeleton of aging and reluctant women, however, remains to be shown.

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Retraction in the light of new data Altered calcitonin gene in a young patient with

osteoporosis

From M Alevizaki and S Legon: We recently reported the sequence of the calcitonin gene of a young male osteoporotic patient in whom no calcitonin could be detected (M Alevizaki, J C Stevenson, S I Girgis, I MacIntyre, S Legon. Br Med J 1989;298:1215-6 (6 May)). We noted a difference of a single base between this sequence and the previously published "normal" sequence in a region which may be important for the maturation of the mRNA, and we speculated that this might have caused his osteoporosis. However, using the polymerase chain reaction to screen normal subjects we now find this sequence to be widespread in the general population. We therefore conclude that this is a neutral polymorphism and is not responsible for the patient's condition. The cause of his calcitonin deficiency thus remains to be established.