Soft tissue body composition differences in monozygotic twins discordant for spinal cord injury
Ann M. Spungen, Jack Wang, Richard N. Pierson, Jr. and William A. Bauman

You might find this additional info useful...

This article cites 35 articles, 11 of which can be accessed free at:
http://jap.physiology.org/content/88/4/1310.full.html#ref-list-1

This article has been cited by 8 other HighWire hosted articles, the first 5 are:
Measuring partial body potassium in the legs of patients with spinal cord injury: a new approach
L. Wielopolski, L. M. Ramirez, A. M. Spungen, S. Swaby, P. Asselin and W. A. Bauman
[Abstract] [Full Text] [PDF]

Effects of chronic spinal cord injury on body weight and body composition in rats fed a standard chow diet
Stefany D. Primeaux, Melissa Tong and Gregory M. Holmes
[Abstract] [Full Text] [PDF]

Relationship of fat mass and serum estradiol with lower extremity bone in persons with chronic spinal cord injury
William A. Bauman, Ann M. Spungen, Jack Wang, Richard N. Pierson, Jr. and Ernest Schwartz
[Abstract] [Full Text] [PDF]

Effect of Low-Dose Baclofen Administration on Plasma Insulin-like Growth Factor-I in Persons With Spinal Cord Injury
William A. Bauman, Steven C. Kirshblum, Nancy G. Morrison, Christopher M. Cirmigliaro, Run-Lin Zhang and Ann M. Spungen
[Abstract] [Full Text] [PDF]

Assessment of skeletal muscle mass in men with spinal cord injury using dual-energy X-ray absorptiometry and magnetic resonance imaging
[Abstract] [Full Text] [PDF]

Updated information and services including high resolution figures, can be found at:
http://jap.physiology.org/content/88/4/1310.full.html

Additional material and information about Journal of Applied Physiology can be found at:
http://www.the-aps.org/publications/jappl

This information is current as of May 29, 2012.
Soft tissue body composition differences in monozygotic twins discordant for spinal cord injury

ANN M. SPUNGEN,1,2 J ACK WANG, 3,4 RICHARD N. PIERSON, JR.,4 AND WILLIAM A. BAUMAN1–3

1The Spinal Cord Damage Research Center, 2Departments of Medicine and Rehabilitation Medicine, Mount Sinai Medical Center, New York 10029; 3Medical Service, Veterans Affairs Medical Center, Bronx 10468; and 4Body Composition Unit, Columbia University-St. Luke's Roosevelt Hospital Center, New York, New York 10025

Spungen, Ann M., Jack Wang, Richard N. Pierson, J r., and William A. Bauman. Soft tissue body composition differences in monozygotic twins discordant for spinal cord injury. J Appl Physiol 88: 1310–1315, 2000.—To determine the effect of paralysis on body composition, eight pairs of male monozygotic twins, one twin in each pair with paraplegia, were studied by dual-energy X-ray absorptiometry. Significant loss of total body lean tissue mass was found in the paralyzed twins compared with their able-bodied co-twins: 47.5 ± 6.7 vs. 60.1 ± 7.8 (SD) kg (P < 0.005). Regionally, arm lean tissue mass was not different between the twin pairs, whereas trunk and leg lean tissue masses were significantly lower in the paralyzed twins: −3.0 ± 3.3 kg (P < 0.05) and −10.1 ± 4.0 kg (P < 0.0005), respectively. Bone mineral content of the total body and legs was significantly related to lean tissue mass in the able-bodied twins (R = 0.88 and 0.98, respectively) but not in the paralyzed twins. However, the intrapair difference scores for bone and lean tissue mass were significantly related (R = 0.80 and 0.81, respectively). The paralyzed twins had significantly more total body fat mass and percent fat per unit body mass index than the able-bodied twins: 4.8 kg (P < 0.05) and 7 ± 2% (P < 0.01). In the paralyzed twins, total body lean tissue was significantly lost (mostly from the trunk and legs), independent of age, at a rate of 3.9 ± 0.2 kg per 5-yr period of paralysis (R = 0.87, P < 0.005). Extreme disuse from paralysis appears to contribute to a parallel loss of bone with loss of lean tissue in the legs. The continuous lean tissue lose may represent a form of sarcopenia that is progressive and accelerated compared with that in ambulatory individuals.

paraplegia; lean tissue mass; percent fat

AN INDIVIDUAL WITH SPINAL cord injury (SCI) and the resultant extreme disuse has a predisposition to medical consequences and secondary disabilities, such as lipid abnormalities (5, 8, 12), carbohydrate intolerance (5, 15), and cardiovascular disease (3, 4, 6). One hypothesis for these secondary disabilities of extreme disuse consists of the adverse changes in body composition (33, 38, 39), which may contribute to insulin resistance and associated disorders. Previous investigators, by using cross-sectional designs comparing age- and height-matched reference populations, have demonstrated lean tissue loss and/or fat tissue gain in individuals with SCI (13, 23, 24, 38). Because of wide individual variability in body composition and the absence of premorbid body composition measurements, a study employing a cross-sectional design is inherently limited in accurately determining the amount of lean tissue lost and fat tissue gained that is attributable exclusively to paralysis. Longitudinal studies are relatively expensive and pose enormous logistical problems, including performing measurements before the body composition changes have occurred, immediately after SCI, a time when the patient is often medically unstable or reluctant to participate in a research study. The task of recruiting subjects to perform these measurements intermittently over ensuing decades is problematic. Thus a creative solution to defining these changes in body composition that is independent of genetic variability and aging is a monozygotic twin model in which one twin has paralysis from traumatic SCI.

METHODS

Subjects. Eight pairs of healthy identical twins, a paralyzed and an able-bodied twin, were studied. The subjects were recruited throughout the United States. To control for known variability within individuals with SCI, a group similar for gender and level and completeness of injury was selected to participate in the study. All subjects were men with motor complete paraplegia, levels T6–L1. None of the paralyzed twins was able to bear weight or had voluntary movement below the level of lesion. The duration of injury was 3–26 yr. All paralyzed twins used their arms daily for wheelchair mobility and were able to transfer independently. The able-bodied and paralyzed twins were in good health with no present known medical problems. The subjects were provided verbal and written information about the study, and written consent was obtained. Institutional Review Board approvals from the Veterans Affairs Medical Center and the Mount Sinai Medical Center were obtained for the study.

Genetic testing for zygosity was performed in all pairs at six independent, highly polymorphic loci (Lifecodes, Stamford, CT). All eight pairs were found to be strongly consistent with monozygosity, with a probability of being nonidentical of 1 in 4,096.

Body composition studies. Height was measured in the standing position for the able-bodied twins and in the supine
<table>
<thead>
<tr>
<th>Twin Pair No.</th>
<th>LOL (Para)</th>
<th>DOI (Para)</th>
<th>Age, yr</th>
<th>Height, m</th>
<th>Weight, kg</th>
<th>BMI, kg/m²</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>T₁₁–L₂</td>
<td>22</td>
<td>39</td>
<td>1.85</td>
<td>76.0</td>
<td>22.2</td>
</tr>
<tr>
<td>2</td>
<td>T₁₀–L₁₂</td>
<td>11</td>
<td>25</td>
<td>1.78</td>
<td>58.6</td>
<td>18.5</td>
</tr>
<tr>
<td>3</td>
<td>L₁</td>
<td>26</td>
<td>47</td>
<td>1.80</td>
<td>66.0</td>
<td>20.4</td>
</tr>
<tr>
<td>4</td>
<td>T₁₂–L₁</td>
<td>8</td>
<td>40</td>
<td>1.83</td>
<td>77.0</td>
<td>23.0</td>
</tr>
<tr>
<td>5</td>
<td>T₆</td>
<td>3</td>
<td>37</td>
<td>1.75</td>
<td>72.0</td>
<td>23.5</td>
</tr>
<tr>
<td>6</td>
<td>T₁₁–L₁₂</td>
<td>22</td>
<td>58</td>
<td>1.73</td>
<td>68.6</td>
<td>22.9</td>
</tr>
<tr>
<td>7</td>
<td>T₇–₈</td>
<td>11</td>
<td>34</td>
<td>1.75</td>
<td>80.5</td>
<td>26.3</td>
</tr>
<tr>
<td>8</td>
<td>T₅</td>
<td>25</td>
<td>43</td>
<td>1.83</td>
<td>72.7</td>
<td>21.7</td>
</tr>
</tbody>
</table>

**RESULTS**

A relatively young group of twins was studied (40 ± 10, range 25–58 yr). The mean duration of injury for the twins with paralysis was 16 ± 9 (range 3–26) yr. Height was similar for both sets of twins. Weight and BMI were significantly lower in the twins with paralysis than in their able-bodied co-twins: 71.4 vs. 81.5 kg (P < 0.05) and 22.3 vs. 25.4 kg/m² (P < 0.05), respectively (Table 1).

**Table 1. Characteristics of the twins**

Para, paralyzed twin with spinal cord injury; AB, able-bodied twin; LOL, level of spinal cord lesion; T, thoracic; L, lumbar; DOI, duration of injury; BMI, body mass index; IPD, intrapair difference score (paralyzed twin – able-bodied twin). *Paired tₙ₋₀ = −2.47, P < 0.05 for Para vs. AB for weight. †Paired t₁₉ = −2.70, P < 0.05 for Para vs. AB for BMI.

The power for total body and leg lean tissue differences between the twin groups was 0.91 and 0.99, respectively.

In the paralyzed twins, increasing duration of injury was significantly associated with total body lean tissue loss (R = 0.87, P < 0.005; Fig. 2). Multiple-regression analysis was performed to test the association of duration of injury with total body IPD lean tissue with control for age. Independent of age, duration of injury was significantly predictive of total body lean tissue loss (partial r = 0.51, P = 0.03). In the twins with paralysis, no relationships or trends were found for level of trunk musculature innervation (T₆–L₃) with total body or trunk lean tissue mass or IPD.

In the able-bodied twins, total body and leg BMC were highly correlated with total body and leg lean tissue mass, but no such relationship was found in the paralyzed twins (Fig. 3). However, in the paralyzed twins, the IPD scores for total body and leg BMC and total body and leg lean mass were significantly related (Fig. 4).

Total body and regional fat tissue. The percent fat in the legs was 17.5 ± 11.7% (P < 0.005) higher in the paralyzed than in the able-bodied twins (Fig. 1).
Fig. 1. Comparisons of total body and regional lean tissue (A) and percent fat (B) between paralyzed (open bars) and able-bodied (filled bars) twins. *P < 0.005; †P < 0.0005; ‡P < 0.05.

body, arm, and trunk fat mass or percent fat was not significantly different between the twin pairs (Fig. 1).

A significant linear relationship was found between BMI and fat mass for both groups (R = 0.75, P < 0.05 for paralyzed twins and R = 0.82, P < 0.01 for able-bodied twins). No significant difference was noted between the slopes of these relationships. With control for BMI by multiple regression, fat mass and percentage were significantly increased by an average of 4.8 ± 1.8 kg (partial r = 0.59, P = 0.02) and 7.2 ± 2.4% (partial r = 0.60, P = 0.01) per unit of BMI in subjects with paralysis compared with the co-twins (Fig. 5).

Table 2. Total body and regional lean tissue mass

<table>
<thead>
<tr>
<th>Twin Pair No.</th>
<th>Total Body Lean, kg</th>
<th>Arms Lean, kg</th>
<th>Trunk Lean, kg</th>
<th>Legs Lean, kg</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Para</td>
<td>AB</td>
<td>IPD</td>
<td>Para</td>
</tr>
<tr>
<td>1</td>
<td>57.9</td>
<td>74.4</td>
<td>-16.4</td>
<td>10.2</td>
</tr>
<tr>
<td>2</td>
<td>52.6</td>
<td>55.1</td>
<td>-2.5</td>
<td>9.8</td>
</tr>
<tr>
<td>3</td>
<td>42.9</td>
<td>64.3</td>
<td>-21.4</td>
<td>6.8</td>
</tr>
<tr>
<td>4</td>
<td>39.9</td>
<td>52.8</td>
<td>-12.8</td>
<td>7.8</td>
</tr>
<tr>
<td>5</td>
<td>49.0</td>
<td>51.2</td>
<td>-2.2</td>
<td>7.9</td>
</tr>
<tr>
<td>6</td>
<td>41.7</td>
<td>55.8</td>
<td>-14.0</td>
<td>7.9</td>
</tr>
<tr>
<td>7</td>
<td>53.7</td>
<td>62.2</td>
<td>-8.5</td>
<td>8.0</td>
</tr>
</tbody>
</table>

Mean ± SD 47.5 ± 6.7* 60.1 ± 7.8 −12.6 ± 7.9 8.3 ± 1.1 7.9 ± 1.3 0.4 ± 1.8 24.6 ± 4.0† 27.6 ± 3.7 −3.0 ± 3.3 10.9 ± 2.8‡ 20.9 ± 3.3 −10 ± 4.0

Lean, soft tissue lean body mass, excluding bone mineral content; see Table 1 footnote for other abbreviations. *Paired t_{(7)} = -4.6, P < 0.005 for total body lean for Para vs. AB. †Paired t_{(7)} = -2.57, P < 0.05 for trunk lean for Para vs. AB. ‡Paired t_{(7)} = -7.10, P < 0.0005 for legs lean for Para vs. AB.
in the size of the quadriceps in the elderly group. In another study by Rice and co-workers (33) using computerized tomography, leg muscle size was reduced by 28–36% in subjects >65 yr of age. Our subjects with paralysis appear to exhibit an exaggerated form of sarcopenia, which is strongly associated with duration of injury and trends with age. The twins with paralysis at a mean age of 40 yr had already lost 20% of their total body mass and 48% of their leg lean tissue mass. This is twice the amount of leg muscle loss and 40 yr earlier than has been previously described in the general population (16, 27, 34).

Several studies have shown that physical activity delays loss of muscle mass with increasing age (9, 18, 19, 28). Our twins with SCI had an abrupt change in level of activity coincident with paralysis. Thus the identical-twin model has provided the design necessary to address whether adverse changes in lean tissue mass as a result of paralysis reach a new plateau or continue to show an age effect. Our findings suggest that chronic inactivity is associated with accelerated and progressive sarcopenia.

A strong correlation between change in body weight and change in total body lean mass has been well established (20, 21). Forbes et al. (22) reported a significant linear relationship between intrapair variation for lean body mass with intrapair variation for weight in monozygotic and dizygotic twins. In our study, total body weight was 10 kg less in the
paralyzed twins than in the able-bodied co-twins. The difference in weight was predominantly from loss of lean tissue in the leg. A direct association was found between intrapair body weight differences and loss of lean and fat tissues in the leg.

Total body percent fat was not significantly different between the twin pairs. Regional percent fat in the legs was significantly greater by an average of 18% in paralyzed twins because of the combination of leg lean tissue loss and a slight increase in leg fat tissue. The able-bodied twins had an average of 1.3 kg less leg fat than the paralyzed twins. Bouchard et al. (10) reported in able-bodied persons that truncal-visceral fat increases with age, particularly in men. Neither total body fat nor regional fat mass was significantly related to age in the paralyzed or the able-bodied twin group, which was probably due to their relatively young ages.

In several studies of monozygotic twins reared apart, adult BMI and weights were reported to have heritability estimates of 0.70 (1, 17, 30a, 39). In the general population, the relationship of fat mass with BMI is strong enough to use BMI as a surrogate measure for obesity (30a). Further studies with a larger population base need to be performed to determine the validity of using BMI as a surrogate measure for obesity in those with SCI. Although the absolute values for lean tissue and bone did not correlate in the twins with paralysis, the IPD scores for total body and leg lean and bone tissues for the able-bodied twins but not for the paralyzed twins. This was probably due to the extreme loss in both parameters (lean and bone) in all twins with SCI. Although the absolute values for lean tissue and bone did not correlate in the twins with paralysis, the IPD scores for total body and leg lean tissue and BMC were well correlated, demonstrating a close relationship between bone and lean tissue loss.

In conclusion, in persons with paralysis from SCI, there is an accelerated form of sarcopenia, similar to that seen in the elderly during the sixth through the eighth decades of life. SCI appears to cause a chronic catabolic state, whereby total body lean tissue, albeit depleted at a rate of 8 kg/decade, about twice the rate of matched controls. Loss of lean tissue is linearly related to loss of BMC. Without the use of an identical-twin model discordant for paralysis, the progression and severity of lean tissue loss throughout the chronic phase of immobilization could not have been identified with this degree of accuracy. In individuals with chronic immobilizing conditions, further studies with identical twins may identify other potential body composition changes, as well as hormonal and metabolic alterations, that may be associated with these body composition changes.

This research was supported by grants from the Spinal Cord Research Foundation (Washington, DC), the Eastern Paralyzed Veterans Association (Jackson Heights, NY), the Veterans Affairs Medical Center (Bronx, NY), and The Mount Sinai Medical Center (New York, NY).

Address for reprint requests and other correspondence: A. M. Spungen, Spinal Cord Damage Research Center, Veterans Affairs Medical Center,
REFERENCES


