IGF-1 LEVELS IN DOWN SYNDROME*

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The authors examined IGF-1 plasma levels in 113 patients with Down Syndrome (DS) (68 males and 45 females, aged 1-23.66 years), all without severe congenital malformations, malnutrition, coeliac disease, alterations in thyroid or adrenal function, or taking any medication which might affect the study. The authors used a radioimmunoassay (RIA) method with plasma previously treated by acid-ethanol extraction and correlated the IGF-1 plasma levels with age and BMI (Body Mass Index) using multiple regression analysis.

The BMI was higher in Down syndrome compared with normal subjects (NS), both in males ($p=0.000006$) and females ($p=0.034829$). The two groups showed a positive correlation of IGF-1 levels with age. However, no correlation was found between IGF-1 levels and BMI.

Keywords: Down syndrome, Insulin-like growth factors, growth hormone

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foetal to the GH-dependent form of IgF-1. Barreca et al measured IgF-1 plasma levels in 39 children affected by Down syndrome, and found a positive correlation with age. In Down syndrome obesity is frequent (Cronk, 1985), probably due to eating behaviour disorders and/or reduced physical activity.

The aim of this study was to detect the IgF-1 plasma levels in subjects with Down syndrome, and to determine whether there is a correlation with age and BMI.

Materials and Methods

One hundred and thirteen subjects with Down syndrome (68 males and 45 females, aged 1-23.66 years and 1-23.8 years respectively), followed routinely at the Oasi Institute, were submitted to IgF-1 plasma assay. Body weight was measured on standard scales and height was measured using a Harpenden stadiometer. Pubertal stage was recorded, using Tanner’s method. There was no evidence of severe congenital malformations, malnutrition, coeliac disease, or alterations of thyroid or adrenal function. None of the patients were taking any medication which might have influenced the study. Blood samples were collected in test-tubes containing EDTA, and were immediately separated and frozen at -20º C until assay time.

Body Mass Index (BMI) was also calculated, (body weight divided by height squared (Kg/m²)).

IgF-1 radio-immuno assay (RIA) was performed (Nichols Institute diagnostic kit, san Juan Capistrano, USA). Plasma samples were matched beforehand with an ethanol-chloridric acid mix (ratio 87.5% ethanol 12.5% chloridric acid). The supernatant was neutralized by adding 0.855 M Tris-base, and was transferred into test-tubes containing 1.4 ml of buffer solution. Final dilution was 1:225.

IgF-1 standards, obtained with DNA-recombinant techniques, were necessary to produce a standard curve. This was calibrated against the WHO 1st International Reference Reagent 1998, Insulin-like Growth Factor-1 87/518. Values were expressed in ng/ml. The curve ranged from 0.3 to 4.9 ng/ml. The sensitivity of levels was calculated to 0.06 ng/ml. The IgF-1 antiserum does not show cross-reactivity with other peptide hormones.

This study was approved by the Ethical Committee of Oasi Institute. Informed consent was obtained from the parents of all the subjects enrolled.

Multiple regression analysis was utilized to correlate IgF-1 with age, and BMI.

Results

IgF-1 plasma levels were assayed for each pubertal stage in people with Down syndrome. Table 1 shows the standard deviation and the mean of IgF-1 levels, age and BMI in Down syndrome males and females respectively.

![Table 1. IgF-1 level, age and BMI found in Down syndrome](image)

<table>
<thead>
<tr>
<th></th>
<th>Down syndrome females (n=45)</th>
<th>Down syndrome males (n=68)</th>
</tr>
</thead>
<tbody>
<tr>
<td>IgF-1 level (ng/ml)</td>
<td>267.4 ± 148.12</td>
<td>251.7 ± 159.35</td>
</tr>
<tr>
<td>Age (years)</td>
<td>11.9 ± 5.13</td>
<td>12.3 ± 6.42</td>
</tr>
<tr>
<td>BMI</td>
<td>20.2 ± 5.89</td>
<td>20.8 ± 4.84</td>
</tr>
</tbody>
</table>

Table 2 shows the multiple regression analysis of IgF-1 levels against age, and IgF-1 levels against BMI in the two groups. The two groups show a significant positive correlation of IgF-1 levels with age. However, no correlation was found between IgF-1 levels and BMI.

![Table 2. Multiple regression analysis between IgF-1 level, age and BMI in Down syndrome](image)

<table>
<thead>
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<th>Down syndrome females (n=45)</th>
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</tr>
</thead>
<tbody>
<tr>
<td>Age</td>
<td>0.000006</td>
<td>0.000001</td>
</tr>
<tr>
<td>BMI</td>
<td>NS</td>
<td>NS</td>
</tr>
</tbody>
</table>

Discussion

We have confirmed a positive correlation between IgF-1 levels and age (Barreca, 1994). We believe that early reports (Anneren et al, 1990) suggesting an IgF-1 selective deficiency in Down syndrome similar to that in Pygmies must have been based on a different methodology.

The BMI figures were significantly higher in Down syndrome compared with normal subjects, and this is further evidence suggesting higher levels of obesity in this group (Cronck et al, 1985).

No correlation between IgF-1 and BMI was found. It follows that BMI remains a reliable tool for the diagnosis of overweight and/or obesity in Down syndrome. The same, however, cannot be said for IgF-1 levels, because, as shown above, they do not correlate with BMI.

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References


