develop subsequent pulmonary and hepatic complications. Phenotype \textit{P},\textit{MZ} will produce intermediate values of \(\alpha_1\)-antitrypsin (approximately 50\%) which may be associated with liver disease.\textsuperscript{4} The Z band is observed to migrate cathodally to the M band.

In these two patients there was an abnormal protein migrating similar to that of the Z band during the acute phase of the hepatitis. This was incompatible with the parents’ phenotypes and resolved during the recovery phase. This phenomenon has been seen once previously in neonatal hepatitis associated with cytomegalovirus infection.\textsuperscript{5} It may be caused by ‘switching on’ of the production of an abnormal protein during the acute phase of the illness, or to decreased post ribosomal sialylation of the normal M protein. This has yet to be clarified.

It is important to be aware of this transient abnormality, which may occur in neonatal hepatitis, and to check both parents’ \(\alpha_1\)-antitrypsin phenotypes. This may avoid erroneous counselling in respect of the prognosis and the genetic implications.

Oxandrolone in low dose for constitutional delay of growth and puberty in boys

R STANHOPE AND C G D BROOK

Department of Paediatrics, Middlesex Hospital, London

**Summary** Twenty four boys with delayed pubertal growth spurt were treated with oxandrolone (2.5 mg daily) for 0-21 to 0-65 years. The mean increment of growth velocity was 4-4 cm/year during treatment, which was maintained at 3-7 cm/year after treatment. There was no significant change in height for bone age standard deviation scores.

Constitutional delay of growth occurs in otherwise healthy adolescents with stature reduced for chronological age but generally appropriate for bone age and the stage of pubertal development, both of which are usually delayed. There is often considerable delay in the onset of the puberty growth spurt which may cause severe psychological disturbance.

Oxandrolone (17\(^{\text{b}}\)-hydroxy-17-methyl-2-oxa-5\(^{\text{b}}\)-androstan-3-one) is one of a group of anabolic steroids derived from testosterone which accelerate growth.\textsuperscript{1} Their use in the treatment of short stature has fallen into disrepute because of potential side effects. These include virilisation, suppression of the hypothalamic-pituitary axis, and hepatotoxicity but the most serious problem is the potential to advance bone age disproportionately to height gain, which compromises final adult height. The poor reputation that anabolic steroids have gained may partly be due to the large doses which were first used.\textsuperscript{1} Growth acceleration does not seem to be related to dose but advance in skeletal maturation is.\textsuperscript{1} We have attempted to define an effective treatment regimen, free of side effects, using oxandrolone.

**Patients and methods**

Twenty four, short, pubertal boys (testicular volumes more than 3 ml) were treated with oxandrolone because of severe psychological disturbance. Two had mild asthma and one had severe psoriasis which had been treated with topical fluorinated corticosteroids. The heights of all patients were less than the 3rd centile for chronological age, except for one, whose height was on the 10th centile. Fourteen had delayed puberty with testicular volumes less than the 10th centile for chronological age.\textsuperscript{2} Mean bone

\[\text{References}\]


\textsuperscript{3} Tobin MJ, Cook PJL, Hutchinson DCS. \(\alpha_1\)-antitrypsin deficiency: the clinical and physiological features of pulmonary emphysema in subjects homozygous for \(\text{P}\), Type \(\text{Z}\). \textit{Br J Dis Chest} 1983;77:14-27.


age retardation was 2-4 years. Oxandrolone was given in a dose of 2-5 mg daily for three to four months to 21 patients and for six months in three patients: four patients received a second course of treatment because height velocity was not maintained in the period after treatment on the first occasion. One child was lost to follow up.

Heights were measured using a stadiometer and height velocities calculated for each preceding inter-\p{v}

Table

<table>
<thead>
<tr>
<th>Pretreatment</th>
<th>Treatment</th>
<th>Post treatment</th>
</tr>
</thead>
<tbody>
<tr>
<td>No</td>
<td>24</td>
<td>24</td>
</tr>
<tr>
<td>No of courses</td>
<td>—</td>
<td>28</td>
</tr>
<tr>
<td>Chronological age at beginning of treatment (years).</td>
<td>Mean (range)</td>
<td>14±8 (12-7-16-7)</td>
</tr>
<tr>
<td>Duration of period (years).</td>
<td>Mean (range)</td>
<td>4-7 (5-8-13)</td>
</tr>
<tr>
<td>Height velocity (cm/yr).</td>
<td>Mean (range)</td>
<td>0-7 (0-8-5-5)</td>
</tr>
<tr>
<td>Change in velocity from pretreatment values.</td>
<td>Mean (SEM)</td>
<td>4-4 (0-37)</td>
</tr>
<tr>
<td>P</td>
<td>&lt;0-001</td>
<td>3-7 (0-47)</td>
</tr>
<tr>
<td>Bone age delay (years).</td>
<td>Mean (range)</td>
<td>-2-4 (-5-1 to -0-5)</td>
</tr>
<tr>
<td>Height SDS for bone age.</td>
<td>Mean (range)</td>
<td>-0-64 (-2-3 to +1-4)</td>
</tr>
<tr>
<td>Change in mean height SDS for bone age from pretreatment values.</td>
<td>Mean (SEM)</td>
<td>-0-26 (0-15)</td>
</tr>
<tr>
<td>P</td>
<td>&gt;0-1</td>
<td>10-5 (6-20)</td>
</tr>
<tr>
<td>Testicular volume at end of each period (ml).</td>
<td>Mean (range)</td>
<td>5-5 (5-15)</td>
</tr>
<tr>
<td>P</td>
<td>&gt;0-1</td>
<td>10-5 (6-20)</td>
</tr>
</tbody>
</table>

Table  Clinical data on pubertal boys treated with oxandrolone (pretreatment, treatment and post treatment periods were in continuity)

No side effects associated with oxandrolone were experienced during this study.

Discussion

Growth delay in adolescent boys is a common problem. When associated with delayed puberty it may be particularly distressing and cause deviant behaviour and severe psychological problems. These may interfere with education at a time when academic achievement is of life long importance and the deviant behaviour may be of such severity that it causes problems with the law (shoplifting, vandalism etc). Most of these boys do not require extensive endocrinological investigation but they do urgently need a treatment for their delay in growth that is free of side effects. All our patients had an improvement of their psychological disturbance following the oxandrolone induced growth acceleration.

Oxandrolone does not seem to advance skeletal maturity excessively, even in dosages of 2-5 to 10 mg/day for periods of up to two years. In pubertal boys we have found that as little as 2-5 mg of oxandrolone daily for a period of three months has produced a highly significant growth acceleration, which persisted without a disproportionate advance in bone age. Even lower doses may be effective in this situation but this requires further study.

The mechanism by which oxandrolone brings its effect is probably by increasing growth hormone secretion. Many boys with a delayed growth spurt have a suppression of growth hormone secretion to the usual provocative stimuli which recovers when
they enter puberty. Administration of testosterone before tests of growth hormone secretion are performed may separate those with temporary growth hormone deficiency from boys with true growth hormone deficiency, who are in any case much shorter than the patients we are treating. We have not, at present, investigated the mechanism of action of oxandrolone but it is clear that we are observing a definite effect of treatment. Boys with testicular volumes of less than 12 ml do not suddenly double their growth velocity over a three month period. The triggering of a sustained growth spurt in individual cases is most impressive.

Oxandrolone has been available for the last two decades but it does not have a product licence in the United Kingdom. We submit that for the purposes we have defined, it would be an important addition to the therapeutic armamentarium.

Candida in mouth or on dummy?

D J MANNING, R P COUGHLIN, AND E M E POSKITT

Alder Hey Children's Hospital, Liverpool; Royal Liverpool Children's Hospital; and Institute of Child Health, University of Liverpool

SUMMARY

Mouth and dummy swabs for Candida spp. were obtained from 100 children under 18 months old admitted with acute medical conditions. Dummies were sampled using swabs moistened with sterile saline. Mouth and dummy swabs were spread on Sabouraud's medium and incubated at 37°C for 24 to 48 hours. Colonies were identified as Candida albicans by Gram stain and a positive germ tube reaction. Germ tube negative strains were identified using the API-Zym system (API Products). Children were allowed to continue sucking their dummies if this was comforting. Further mouth and dummy swabs were obtained twice weekly during each child's admission.

Results

Sixty six children sucked a dummy regularly and 34 did not. There was no significant difference in age, sex, gestational age at birth, or method of feeding between those who sucked dummies and those who did not. Similarly, there was no social class difference between the two groups. Medical admissions to this hospital come predominantly from a deprived urban area with a high rate of unemployment.

Seven children had clinical oral thrush (all C albicans). Six of these children sucked dummies and positive dummy cultures were obtained from all but one of this group.

Thirty six children who sucked dummies had medical conditions. Dummies were sampled using swabs moistened with sterile saline. Mouth and dummy swabs were spread on Sabouraud's medium and incubated at 37°C for 24 to 48 hours. Colonies were identified as Candida albicans by Gram stain and a positive germ tube reaction. Germ tube negative strains were identified using the API-Zym system (API Products). Children were allowed to continue sucking their dummies if this was comforting. Further mouth and dummy swabs were obtained twice weekly during each child's admission.

References


Correspondence to Dr C G D Brook, Department of Paediatrics, The Middlesex Hospital, Mortimer Street, London, W1N 8AA.

Received 2 November 1984