The influence of the route of administration on the effect of hexoestrol on fattening wether lambs

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The administration of hexoestrol to livestock has been found by many workers to increase the rate of live-weight gain. It has been shown that hexoestrol treatment of lambs alters their metabolism in such a way that absorbed nutrients are diverted, in part, from fat anabolism to bone and muscle anabolism (Gee & Preston, 1957). Since fat has a much higher calorific value per unit weight than either bone or muscle, and is generally stored with relatively little water, it follows that a given intake of food will produce a greater live-weight gain in a lamb treated with hexoestrol than in a lamb that has not been treated.

An increase in efficiency of conversion of food to live weight as a result of hexoestrol treatment has been reported by many investigators. More detailed carcass studies (Gee & Preston, 1957) showed that treated lambs were 33% more efficient in converting food protein to muscle protein but that efficiency of energy conversion was not affected by treatment. If these changes are desirable the findings would appear to justify the use of hexoestrol treatment in commercial lamb and mutton production (Preston & Gee, 1957).

In our previous experiments treatment of lambs was mainly by subcutaneous implantation in the left ear of a single 15 mg pellet of hexoestrol. In poultry one of the factors governing the rate of absorption of the hormone is the surface area of the pellet (Lorenz, 1954). Division of the standard 15 mg pellet into three 5 mg portions would be a possible way of increasing absorption without increasing the total dose.

Synthetic oestrogen may also be given by mouth in the food. Burroughs, Culbertson, Cheng, Hale & Homeyer (1955) believe that objectionable side-effects, mainly slackening of the pelvic ligaments and mammary development, are less when cattle are fed stilboestrol than when the hormone is implanted subcutaneously.

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A comparison was therefore made of the relative effects of four methods of administering hexoestrol on the growth and certain carcass characteristics of wether lambs.

**EXPERIMENTAL**

**Treatment of lambs.** Fifty Suffolk × Greyface 5-month-old wether lambs were allocated to five groups. Group A were controls. Group B had implanted subcutaneously in the left ear a 15 mg pellet of hexoestrol. Group C received two 5 mg pellets of hexoestrol in the left ear and a third 5 mg pellet in the right ear. Lambs in group D were given in the ration 2 mg hexoestrol per head daily, the total dose in the trial period being 0.186 g. Lambs in group E received 4 mg per head daily, the total dose being 0.388 g.

Lambs were given daily 50 g hay, and concentrates to appetite. The concentrate mixture was: maize meal 30, flaked maize 20, bran 10, molassine meal* 10, barley meal 10 and decorticated groundnut meal 20%. The hexoestrol for groups D and E was incorporated in the diet at the rate of 1 g to 50 kg of concentrates, and 100 and 200 g of this mixture were fed to each lamb of groups D and E respectively. The remainder of the concentrate ration consisted of the unsupplemented mixture.

The lambs were housed individually indoors for about 90 days and were slaughtered when they reached a live weight of 50 kg.

**Slaughter technique.** Immediately after slaughter the warm carcass was weighed. The teats were cut off close to the udder, measured and weighed. The thyroid glands and pituitary body were dissected out and weighed. The carcasses were left to cool for 24 h, reweighed and measurements taken of the length of leg ($F$) and width of hindquarters ($G$) (Pálsson, 1939). The left leg was separated from the carcass by a transverse cut through the lateral muscles at the level of the articulation of the femur with the acetabulum of the pelvis. The leg was then disarticulated and cut along the ischial arch as close to the bone as possible. The joint was weighed and dissected into edible meat and bone. These tissues were weighed and the edible meat was minced to obtain a representative sample which was stored at $-20\,^\circ$. Each sample was freeze-dried, minced again and analysed for fat (Callow, 1944), residual moisture and ash. Protein was calculated as 90% of the dry fat-free residue (Callow, 1944).

The ears from the implanted animals were kept after slaughter and the remaining pieces of pellets were dissected out, dried and weighed.

**RESULTS**

Most of the lambs lost weight during the first 2 weeks of the experiment. This setback was probably due to a too rapid change-over from pasture to a high-concentrate diet.

Four lambs, one from each of the groups A, B, D and E had difficulty in urination and oedema of the anal area: they were slaughtered before the end of the experiment. In each lamb urinary calculi, mainly calcium phosphate, were present in the penile region of the urethra.

* Contains 75% molasses and 25% sphagnum moss (Molassine Co. Ltd, Greenwich, London, S.E. 10).
Live-weight gains, food-conversion rates, carcass weights and measurements, and
dressing out percentages (see Gee & Preston, 1957) for the five groups of lambs are
set out in Table 1. Although overall differences in daily live-weight gain between
treatments were not significant, the difference between the control and the other four
treatments taken together was in itself significant at the 5% level and showed the rate
of gain to be higher for the treated than for the untreated lambs. Treated lambs
required less food per unit of live-weight increase than untreated lambs, which is in
agreement with our earlier report (Gee & Preston, 1957). There were no significant
differences between treatments for dressing-out percentage or the ratio 100 $G:F$.

Table 1. Growth, food-conversion rate and carcass measurements
of control and hexoestrol-treated lambs

<table>
<thead>
<tr>
<th>Group</th>
<th>Group B (15 mg subcutaneously)</th>
<th>Group C (3 x 5 mg subcutaneously)</th>
<th>Group D (2 mg daily in ration)</th>
<th>Group E (4 mg daily in ration)</th>
<th>Standard error of difference between any two treatments</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. of lambs</td>
<td>10</td>
<td>9</td>
<td>10</td>
<td>10</td>
<td>—</td>
</tr>
<tr>
<td>Initial weight (kg)</td>
<td>32.1</td>
<td>32.9</td>
<td>31.6</td>
<td>32.2</td>
<td>31.4</td>
</tr>
<tr>
<td>Final weight (kg)</td>
<td>49.9</td>
<td>51.9</td>
<td>50.8</td>
<td>50.2</td>
<td>50.5</td>
</tr>
<tr>
<td>Daily gain (kg)</td>
<td>0.174*</td>
<td>0.195</td>
<td>0.208</td>
<td>0.193</td>
<td>0.197 ± 0.013</td>
</tr>
<tr>
<td>Food-conversion rate (kg gain/kg food)</td>
<td>0.168</td>
<td>0.195</td>
<td>0.210</td>
<td>0.192</td>
<td>0.194 ± 0.014</td>
</tr>
<tr>
<td>Carcass weight (kg)</td>
<td>24.4</td>
<td>24.4</td>
<td>24.2</td>
<td>24.5</td>
<td>24.3</td>
</tr>
<tr>
<td>Dressing-out percentage</td>
<td>48.9</td>
<td>47.9</td>
<td>47.7</td>
<td>48.9</td>
<td>48.3 ± 0.8</td>
</tr>
<tr>
<td>Ratio, 100 $G:F$</td>
<td>99.1</td>
<td>97.9</td>
<td>97.8</td>
<td>102.0</td>
<td>101.1 ± 2.7</td>
</tr>
</tbody>
</table>

* Control group gained significantly less ($P < 0.05$) than groups B–E taken together.

There were significant differences between treatments in the weight and length of
rudimentary teats and in the weights of the thyroid gland and pituitary body (Table 2).
All the hexoestrol-treated lambs showed greater development of the teats than the
d controls. Weights of the thyroid glands and pituitary bodies were also heavier in the
treated animals, with the exception of the thyroid weight for group B. For each
measurement the highest value was recorded for lambs in group E, which were given
by mouth 4 mg hexoestrol daily.

The weights of the leg joints, expressed as percentages of carcass weights, and the
chemical composition of the leg joints for the five groups of lambs are presented in
Table 3.

The leg joints of all groups of treated lambs contained significantly less fat and more
muscle than those from control lambs. The percentage of bone was also increased by
hexoestrol treatment although the differences between groups were not significant.

Group C lambs, which were treated with three 5 mg pellets, absorbed 87% of the
original implants while group B lambs which received one 15 mg pellet absorbed only
69%. This difference was significant at the 1% level.

Histological examination of the ears of lambs implanted with hexoestrol showed a
localized area of fibrosis with an accumulation of pigmented phagocytic cells at the
site of the injection.
### Table 2. Development of teats, thyroid glands and pituitary bodies of control and hexoestrol-treated lambs

<table>
<thead>
<tr>
<th></th>
<th>Group A (control)</th>
<th>Group B (15 mg subcutaneously)</th>
<th>Group C (3 × 5 mg subcutaneously)</th>
<th>Group D (2 mg daily in ration)</th>
<th>Group E (4 mg daily in ration)</th>
<th>Standard error of difference between any two treatments</th>
<th>Overall significance of treatment differences</th>
</tr>
</thead>
<tbody>
<tr>
<td>No. of lambs</td>
<td>10</td>
<td>9</td>
<td>10</td>
<td>10</td>
<td>10</td>
<td>-</td>
<td>-</td>
</tr>
<tr>
<td>Mean length of teats (cm)</td>
<td>1.1</td>
<td>2.2</td>
<td>2.0</td>
<td>2.0</td>
<td>2.2</td>
<td>±0.12</td>
<td>P &lt; 0.001</td>
</tr>
<tr>
<td>Mean weight of teats (g)</td>
<td>0.47</td>
<td>1.85</td>
<td>1.46</td>
<td>1.70</td>
<td>1.95</td>
<td>±0.20</td>
<td>P &lt; 0.001</td>
</tr>
<tr>
<td>Weight of thyroid (g)</td>
<td>1.95</td>
<td>1.65</td>
<td>2.36</td>
<td>2.16</td>
<td>2.64</td>
<td>±0.32</td>
<td>P &lt; 0.05</td>
</tr>
<tr>
<td>Weight of pituitary (g)</td>
<td>0.57</td>
<td>0.70</td>
<td>0.69</td>
<td>0.60</td>
<td>0.82</td>
<td>±0.07</td>
<td>P &lt; 0.01</td>
</tr>
</tbody>
</table>

### Table 3. Composition of leg joints of control and hexoestrol-treated lambs

<table>
<thead>
<tr>
<th>Leg joint</th>
<th>Group A (control) (10)</th>
<th>Group B (15 mg subcutaneously) (9)</th>
<th>Group C (3 × 5 mg subcutaneously) (10)</th>
<th>Group D (2 mg daily in ration) (10)</th>
<th>Group E (4 mg daily in ration) (9)</th>
<th>Standard error of difference between any two treatments</th>
<th>Overall significance of treatment differences</th>
</tr>
</thead>
<tbody>
<tr>
<td>Weight (as percentage of carcass weight)</td>
<td>11.5</td>
<td>11.1</td>
<td>11.0</td>
<td>11.4</td>
<td>11.2</td>
<td>-</td>
<td>-</td>
</tr>
<tr>
<td>Content of edible meat (as percentage of weight of joint):*</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>‘Fat’</td>
<td>18.7</td>
<td>14.6</td>
<td>13.5</td>
<td>15.6</td>
<td>14.0</td>
<td>±1.52</td>
<td>P &lt; 0.05</td>
</tr>
<tr>
<td>Protein</td>
<td>13.1</td>
<td>13.9</td>
<td>14.2</td>
<td>13.4</td>
<td>13.9</td>
<td>±0.45</td>
<td>-</td>
</tr>
<tr>
<td>Moisture</td>
<td>53.0</td>
<td>55.6</td>
<td>56.0</td>
<td>54.8</td>
<td>56.3</td>
<td>±1.24</td>
<td>-</td>
</tr>
<tr>
<td>Muscle†</td>
<td>66.0</td>
<td>69.5</td>
<td>70.2</td>
<td>68.2</td>
<td>70.1</td>
<td>±1.43</td>
<td>P &lt; 0.05</td>
</tr>
<tr>
<td>Bone</td>
<td>13.9</td>
<td>14.6</td>
<td>14.8</td>
<td>14.7</td>
<td>14.1</td>
<td>±0.50</td>
<td>-</td>
</tr>
</tbody>
</table>

* Figures in parentheses are the numbers of lambs in the group.

- The values for ‘fat’, protein, moisture and bone do not add to 100% because of the assumption that 90% of dry fat-free tissue is muscle protein (see p. 159). The difference represents mainly carbohydrate and inorganic constituents.

† Protein plus moisture.
DISCUSSION

In several American experiments (Jordan, 1953; Bell, Smith & Erhart, 1954; Henne- 
man, Rust & Meites, 1957) there have been reports of urinary calculi in lambs treated
with stilboestrol. In our experiment the presence of urinary calculi in a control lamb
indicated that factors other than hexoestrol treatment were involved. The lambs were
confined in small pens, and the lack of exercise combined with the high-concentrate
diet may have been predisposing factors. In this connexion Elam, Schneider & Ham
(1956) reported a high incidence of calculi in wether lambs taken off a diet of lucerne
hay and fed on rations containing over 50% of concentrates.

The increase in daily live-weight gain with hexoestrol treatment is in agreement
with the findings of most other workers. Expressed as a percentage of the daily weight
gain made by control lambs, the weight gains made by groups B, C, D and E were
112, 119, 111 and 113% respectively. The significant increase in hexoestrol absorption
and the slightly greater daily live-weight gain recorded for lambs implanted with three
5 mg pellets compared with those that received a single 15 mg implant were probably
due to the larger surface area of the smaller pellets, a suggestion that is in agreement
with findings for poultry (Lorenz, 1954). During the early stages of the experiment
implanted lambs grew faster than those given hexoestrol in the diet (Fig. 1). In the
remaining weeks of the experiment the position was reversed and the greatest gains
were made by lambs in groups D and E, which would suggest that absorption from
an implant is high in the first weeks after implantation but subsequently declines with
diminishing surface area of the pellet and as fibrous tissue forms round the pellet.
On the other hand, when hexoestrol is given in the ration one would expect the rate
of absorption to be constant throughout the treatment period. If mammary develop-
ment in wether lambs is an index of the amount of oestrogen being absorbed, then this

Fig. 1. Mean weekly live-weight gains of lambs given hexoestrol in the ration (groups D and E, black
columns) and lambs implanted with hexoestrol (groups B and C, white columns).

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hypothesis is supported since animals in group E showed greatest development of the rudimentary teats at the end of the experiment. Group E lambs also had the heaviest thyroid and pituitary glands which are other reported effects of oestrogen treatment.

The most marked effect of hexoestrol treatment was shown in the composition of the leg joints. The increase in bone and muscle and accompanying decrease in fat in all hexoestrol-treated lambs, compared with the controls, is in accordance with our earlier findings (Gee & Preston, 1957). That fat content should be least and bone and muscle content greatest in the group gaining weight most rapidly and requiring least food per unit of live-weight increase supports the suggestion that hexoestrol stimulates bone and muscle anabolism at the expense of fat anabolism. This increase in the proportion of tissues of low calorific value, tissues, moreover, associated with more water than is fat, accounts for the greater increase in live weight of treated animals.

SUMMARY

1. Fifty 5-month-old wether lambs were allocated to five groups. Group A received no hexoestrol; lambs of group B were implanted with a 15 mg pellet and those of group C with three 5 mg pellets of hexoestrol. Group D were given 2 mg hexoestrol per head daily in the ration and group E 4 mg.

2. Four lambs, one from each of groups A, B, D and E, developed urinary calculi and were slaughtered before the end of the experiment.

3. Live-weight gains of groups B, C, D and E expressed as percentages of that of the untreated group A were 112, 119, 111 and 113% respectively. The differences between these figures were not statistically significant, but the mean gain for groups B, C, D and E considered together was greater than the mean gain in group A.

4. Treated animals showed greater development of thyroid, pituitary and mammary glands.

5. The leg joints of treated lambs contained less fat and more muscle and bone than those of control lambs.

6. Lambs in group C absorbed significantly more of the implanted hexoestrol than those in group B.

We are indebted to Mr W. A. Boyne for carrying out the statistical analyses and to Mr B. F. Fell for histological examination of the ears from implanted lambs.

REFERENCES