Absorption of lactose and its digestion products in the normal and malnourished Ugandan

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EDITORIAL COMMENT There is a correlation between the rise in capillary blood glucose and the percentage absorption of lactose following an oral load in normal Ugandan subjects and without lactase deficiency. Lactase deficiency occurs in some children with kwashiorkor.

Primary or specific lactase deficiency is very common in healthy members of the Baganda and other Bantu tribes of Uganda after the first three or four years of life (Cook and Kajubi, 1966; Cook, 1967). Secondary lactase deficiency has been recorded in some cases of kwashiorkor (Bowie, Brinkman, and Hansen, 1965; Stanfield, Hutt, and Tunnicliffe, 1965), where there is mucosal damage in the proximal jejunum. The functional importance of the lactase deficiency in both of these situations is not yet clear.

The lactose-tolerance test has been widely used as a screening test for lactase deficiency and it is important to establish its true value. Several investigators have found a good correlation between the lactase level in specimens of jejunal mucosa and the maximum rise in blood glucose after lactose (Dunphy, Littman, Hammond, Forstner, Dahlqvist, and Crane, 1965; McMichael, Webb, and Dawson, 1965, 1966; Petrenel, 1965; Cook and Kajubi, 1966; Welsh, 1966). Others have not found such a good association (Friedland, 1965; Newcomer and McGill, 1966a).

The present study was undertaken to assess the relationship between the rise in capillary blood glucose and percentage absorption of lactose digestion products from the jejunum after oral lactose. By an intubation technique the amount of unab sorbed lactose in the jejunum was measured. In a few subjects jejunal disaccharidase levels were also estimated. Two groups were studied: a series of relatively normal Ugandan hospital patients, and a group of children with mild to moderately severe kwashiorkor.

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MATERIALS AND METHODS

Thirteen in-patients at Mulago Hospital without clinical evidence of malnutrition or gastrointestinal disease were investigated (Tables I and II).

Five children with kwashiorkor were studied at varying stages of their illness (Table IV). Their ages ranged from 14 to 26 (mean 18.8) months. All were Baganda. The mean body weight on admission to hospital was 8.5 (7.8-9.9) kg. Grading of the severity of kwashiorkor was based on the criteria of Dean (1960).

On the evening before each test, the subjects swallowed a polyvinyl tube3 of internal diameter 1·0 mm. in which six holes were cut at the distal end. The tube was threaded through the nose, out of the mouth, and a mercury weight attached before swallowing. The end of the tube was passed to a level which was calculated to be past the area in which most glucose and lactose absorption usually occurs (Hirsch, Ahrens, and Blankenhorn, 1956; Borgström, Dahlqvist, Lundh, and Sjövall, 1957), and its distance from the nasal septum was recorded (Tables I, II, and IV). A straight radiograph of the abdomen was taken before every test to ensure that the tube was not abnormally coiled, and was outlining the normal loops of small intestine. The tolerance tests were performed in the mornings after an eight-hour fast. In all subjects two tests were done on succeeding days, one with lactose and the other with glucose + galactose, both being given as 10% W/V solutions containing also polyethylene glycol (P.E.G.), molecular weight 4,000, at a concentration of 1% W/V. Lactose tolerance was estimated before glucose + galactose tolerance in all except subjects nos. 6 and 8-13. Capillary blood was taken at 0, 30, 60, 90, 120, and 150 minutes. Samples of jejunal contents were siphoned at 15-minute intervals, and 5 ml. normal saline solution was injected into the tube if intestinal contents did not readily appear. All specimens were immediately frozen solid. Percentage absorption of carbohydrate was calculated according to

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The method of Dahlqvist and Bergström (1961) and was taken to be the mean of the percentage absorption from all samples of jejunal aspirate containing reducing substance. In four normal subjects, mucosal biopsy was obtained under radiological control from the first or second jejunal loop with the biopsy capsule described by Choudhury, Nicholson, and Cooke (1964). The specimen was immediately divided; part was examined in the fresh state with the dissecting microscope; the remainder was frozen to -15°C., and disaccharidases—lactase, sucrase, maltase, and trehalase—were assayed by the method described by Cook and Kajubi (1966). In the kwashiorkor group, 24-hour stools during and after tolerance tests were immediately acidified with N sulphuric acid, weighed, and the lactic acid content was estimated (Long, 1946; Elsdon and Gibson, 1954), using a Markham micro-Kjeldahl apparatus.

Blood glucose was estimated by a glucose-oxidase method (Marks, 1959) and total reducing substance in the jejunal aspirate and stools by the copper-reduction method of Asatoor and King (1954). Polyethylene glycol was estimated according to Hydén (1956) and Jacobson, Bondy, Broitman, and Fordtran (1963).

RESULTS

CARBOHYDRATE ABSorption in the normal group

Table I shows the results of lactose and glucose + galactose tolerance tests in 10 normal subjects. The mean fasting blood glucose level was 65.6 (48-83) mg. per 100 ml. Mean maximum blood glucose rise after lactose was 16.9 (1.35) mg. per 100 ml. in the Baganda and 33.0 (18.51) mg. per 100 ml. in the subjects from other tribes. The mean maximum rise after glucose + galactose was 43.4 (23.69) mg. per 100 ml. The mean percentage absorption was 50.8

(12.99) after lactose and 92.3 (78-100) after glucose + galactose. In a 1-year-old Baganda child and two adults from Hamitic tribes who have high lactase activity (Cook and Kajubi, 1966) the percentage absorption of lactose was at least 75%. The correlation between the maximum rise in blood glucose (mg. per 100 ml.) and percentage absorption of carbohydrate in the 10 subjects was statistically significant (Fig. 1). The results of duplicate lactose tolerance tests in three normal subjects are shown in

![Graph showing correlation between maximum rise in blood glucose and percentage absorption of carbohydrate after oral lactose and glucose + galactose in the normal Ugandan subjects.](http://gut.bmj.com/)

**Table I**

<table>
<thead>
<tr>
<th>Patient No.</th>
<th>Age (yr.)</th>
<th>Sex</th>
<th>Tribe</th>
<th>Diagnosis</th>
<th>Body Weight (kg.)</th>
<th>Distance of Sampling from Nasal Septum (cm.)</th>
<th>Maximum Glucose Rise (mg./100 ml.)</th>
<th>Percentage Absorption after Lactose (%)</th>
<th>Diarrhoea</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>1-5</td>
<td>F</td>
<td>Baganda</td>
<td>Pulmonary tuberculosis</td>
<td>10-4</td>
<td>100</td>
<td>35</td>
<td>25</td>
<td>99</td>
</tr>
<tr>
<td>2</td>
<td>3-5</td>
<td>M</td>
<td>Baganda</td>
<td>Normal</td>
<td>15-0</td>
<td>120</td>
<td>25</td>
<td>52</td>
<td>39</td>
</tr>
<tr>
<td>3</td>
<td>5-0</td>
<td>M</td>
<td>Baganda</td>
<td>Normal</td>
<td>19-5</td>
<td>120</td>
<td>1</td>
<td>69</td>
<td>55</td>
</tr>
<tr>
<td>4</td>
<td>7-10</td>
<td>M</td>
<td>Baganda</td>
<td>Normal</td>
<td>23-6</td>
<td>120</td>
<td>20</td>
<td>49</td>
<td>15</td>
</tr>
<tr>
<td>5</td>
<td>10-0</td>
<td>F</td>
<td>Baganda</td>
<td>Sickle cell disease</td>
<td>29-5</td>
<td>120</td>
<td>6</td>
<td>60</td>
<td>47</td>
</tr>
<tr>
<td>6</td>
<td>12-0</td>
<td>F</td>
<td>Baganda</td>
<td>Type I nephritis (convalescent)</td>
<td>43-2</td>
<td>130</td>
<td>1</td>
<td>30</td>
<td>30</td>
</tr>
<tr>
<td>7</td>
<td>12-0</td>
<td>M</td>
<td>Baganda</td>
<td>Burkitt's lymphoma in remission</td>
<td>28-6</td>
<td>150</td>
<td>30</td>
<td>53</td>
<td>12</td>
</tr>
<tr>
<td>8</td>
<td>13-0</td>
<td>F</td>
<td>Bahutu (Rwanda)</td>
<td>Idiopathic splenomegaly</td>
<td>38-6</td>
<td>135</td>
<td>30</td>
<td>39</td>
<td>46</td>
</tr>
<tr>
<td>9</td>
<td>23-0</td>
<td>M</td>
<td>Barutsi (Rwanda)</td>
<td>Tonsillitis (convalescent)</td>
<td>63-6</td>
<td>145</td>
<td>18</td>
<td>23</td>
<td>75</td>
</tr>
<tr>
<td>10</td>
<td>17-0</td>
<td>M</td>
<td>Bahima (Ankole)</td>
<td>Rheumatic arthritis</td>
<td>51-8</td>
<td>190</td>
<td>51</td>
<td>34</td>
<td>90</td>
</tr>
</tbody>
</table>

1 g/kg. body weight (maximum 50 g.)

2 g/kg. body weight (maximum 50 g.)

3 Jejunul disaccharidases estimated.
Table II. In nine subjects the peak blood glucose level after lactose was at 30-60 minutes, in two at 90 minutes, and in two at 120 minutes. After glucose + galactose the maximum level was at 30 or 60 minutes in 11 and at 90 minutes in two. Reducing substance in the jejunal fluid was first detected between 30 and 45 minutes in nine subjects; in nos. 2, 6, and 12 it was first found at 15 minutes, and in no. 10 at 90 minutes. Four Baganda had diarrhoea after the lactose tolerance test.

JEJUNAL BIOPSIES IN NORMAL GROUP Table III shows the results of jejunal biopsies and disaccharidase levels. Well-formed leaves were present in all biopsy material. Lactase levels were consistent with the tribal pattern for Uganda described by Cook and Kajubi (1966). Other disaccharidase levels—sucrase, maltase, and trehalase—were within normal limits for a European population (McMichael et al., 1966).

CARBOHYDRATE ABSORPTION IN THE KWSASHIKOR GROUP Table IV shows details of tolerance tests in the five children. The mean fasting blood glucose was 46-3 (23-68) mg. per 100 ml. and the mean maximum rise in blood glucose was 20-4 (0-50) mg. per 100 ml. after lactose, and 39-3 (8-90) mg. per

![Graph](http://gut.bmj.com/)

**FIG. 2.** Correlation between maximum rise in blood glucose (mg. per 100 ml.) and percentage absorption of carbohydrate after oral lactose and glucose + galactose in the children with kwashiorkor. The line represents the calculated regression (percentage absorption = 44.57 + 2.955 (maximum rise in blood glucose — 29.83); r = +0.592; P < 0.001). 

100 ml. after glucose + galactose. The mean percentage absorption was 34-7 (5-90) after lactose, and

## TABLE II

<table>
<thead>
<tr>
<th>Patient No.</th>
<th>Age (yr.)</th>
<th>Sex</th>
<th>Tribe</th>
<th>Diagnosis</th>
<th>Body Weight (kg.)</th>
<th>Distance of Sampling Tube from Nasal Septum (cm.)</th>
<th>Maximum Glucose Rise after Lactose (mg/100 ml.)</th>
<th>Percentage Absorption after Lactose</th>
<th>Diarrhoea after Lactose</th>
</tr>
</thead>
<tbody>
<tr>
<td>11</td>
<td>1-7</td>
<td>F</td>
<td>Baganda</td>
<td>Pulmonary tuberculosis</td>
<td>8-6</td>
<td>100</td>
<td>33</td>
<td>12</td>
<td>60</td>
</tr>
<tr>
<td>12</td>
<td>12-0</td>
<td>M</td>
<td>Baganda</td>
<td>Chronic rheumatic cardiac disease</td>
<td>42-3</td>
<td>120</td>
<td>0</td>
<td>6</td>
<td>25</td>
</tr>
<tr>
<td>13</td>
<td>10-0</td>
<td>F</td>
<td>Bahutu (Rwanda)</td>
<td>Nephrotic syndrome in remission</td>
<td>38-2</td>
<td>120</td>
<td>18</td>
<td>0</td>
<td>47</td>
</tr>
</tbody>
</table>

1 2 g./kg. body weight (maximum 50 g.).
2 Jejunal disaccharidases estimated.

## TABLE III

JEJUNAL DISACCHARIDASE ESTIMATIONS IN THE NORMAL SUBJECTS

<table>
<thead>
<tr>
<th>Patient No.</th>
<th>Tribe</th>
<th>Dissecting Microscope Appearance of Jejunal Biopsy Specimen</th>
<th>Lactase</th>
<th>Sucrase</th>
<th>Maltase</th>
<th>Trehalase</th>
</tr>
</thead>
<tbody>
<tr>
<td>6</td>
<td>Baganda</td>
<td>Thin leaves; no finger villi</td>
<td>0-7</td>
<td>15-9</td>
<td>50-8</td>
<td>8-6</td>
</tr>
<tr>
<td>8</td>
<td>Bahutu (Rwanda)</td>
<td>Thick leaves; no finger villi</td>
<td>4-3</td>
<td>10-8</td>
<td>38-1</td>
<td>3-9</td>
</tr>
<tr>
<td>10</td>
<td>Bahima (Ankole)</td>
<td>Thin leaves; no finger villi</td>
<td>0-5</td>
<td>12-1</td>
<td>37-6</td>
<td>6-2</td>
</tr>
</tbody>
</table>

1 All units expressed as μM disaccharide hydrolysed per g. tissue wet weight per minute at 37°C.
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TABLE IV
RESULTS OF LACTOSE AND GLUCOSE + GALACTOSE TOLERANCE TESTS IN FIVE CHILDREN WITH KWASHIORKOR

<table>
<thead>
<tr>
<th>Child No.</th>
<th>Age (months)</th>
<th>Sex</th>
<th>Severity of Kwashiorkor</th>
<th>Initial Plasma Protein (g./100 ml.)</th>
<th>Day after Admission</th>
<th>Distance of Sampling from Nasal Septum (cm.)</th>
<th>Maximum Glucose Rise (mg./100 ml.)</th>
<th>Percentage Absorption</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>19</td>
<td>F</td>
<td>Moderately severe</td>
<td>4·5</td>
<td>1 &amp; 2</td>
<td>45</td>
<td>0</td>
<td>29</td>
</tr>
<tr>
<td>2</td>
<td>14</td>
<td>F</td>
<td>Mild</td>
<td>3·8</td>
<td>1 &amp; 2</td>
<td>45</td>
<td>20</td>
<td>11</td>
</tr>
<tr>
<td>3</td>
<td>26</td>
<td>F</td>
<td>Moderately severe</td>
<td>5·5</td>
<td>2 &amp; 3</td>
<td>45</td>
<td>19</td>
<td>28</td>
</tr>
<tr>
<td>4</td>
<td>17</td>
<td>F</td>
<td>Mild</td>
<td>3·6</td>
<td>1 &amp; 2</td>
<td>45</td>
<td>12</td>
<td>46</td>
</tr>
<tr>
<td>5</td>
<td>18</td>
<td>F</td>
<td>Moderately severe</td>
<td>5·1</td>
<td>2 &amp; 3</td>
<td>45</td>
<td>38</td>
<td>46</td>
</tr>
</tbody>
</table>

12 g. lactose/kg. body weight.  
1 g. glucose + 1 g. galactose/kg. body weight.

54·5 (6-100) after glucose + galactose. The correlation between the maximum rise in blood glucose (mg. per 100 ml.) and percentage absorption of carbohydrate after all tests in the five children was statistically significant (Fig. 2). The peak glucose level was late (at 120 or 150 minutes) in five tests after lactose and in three after glucose + galactose. In 20 tests reducing substance was present in the jejunal fluid in the 15-30-minute sample, in six at 30 to 45 minutes, in three at 45 to 60 minutes, and in one at 90 minutes. Moderately severe diarrhoea was present on admission in children nos. 1 and 5. It was not worsened by lactose. In all children, loose stools were recorded as frequently after glucose + galactose as lactose.

STOOL WEIGHT, LACTIC ACID, AND TOTAL REDUCING SUBSTANCE IN KWASHIORKOR GROUP Table V summarizes the results. The maximum lactic acid excretion in the 24 hours after lactose was 305 mg. (equivalent to approximately 3% of the oral load).

The mean lactic acid excretion after glucose + galactose was slightly greater than that after lactose. The correlation between the 24-hour stool weight and lactic acid excretion was significant (r = +0·698; P<0·001). The correlation between stool weight and total reducing substance was also significant (r = +0·803; P<0·001). No significant alteration in any of these measurements took place during treatment.

Discussion

There was a good correlation in the normal group and the group of children with mild to moderately severe kwashiorkor between the maximum rise in glucose and percentage absorption of lactose and its constituent monosaccharides. Reproducibility of the method was reasonably good (Tables II and IV). Using a technique similar to that used in the present study, Borgström et al. (1957) showed that approximately 80% of glucose and lactose had been absorbed in the normal European adult at 150 cm. from the nasal septum. Maximum levels of lactate have been thought to occur in the proximal jejunum (Dahlqvist and Borgström, 1961; Dahlqvist, 1962), although recent evidence (Newcomer and McGill, 1966b) suggests that a high level normally exists in the distal jejunum and ileum also.

In most of the normal subjects there was complete absorption of glucose + galactose. The distal end of the tube was therefore far enough down to be below an area of high monosaccharide absorption.
In most investigations in the kwashiorkor group, the end of the tube was positioned at 45 cm. from the nasal septum, and in most of these, less than 50% of the glucose + galactose load was absorbed. When the tube was placed at 80 cm. there was complete absorption. Approximately 50% of the absorption of monosaccharides in the children with kwashiorkor occurred therefore between 45 and 80 cm. from the nasal septum.

A high incidence of primary lactase deficiency and lactose intolerance in the Baganda tribe (Cook and Kajubi, 1966) has been confirmed. Most of the children with mild or moderately severe kwashiorkor studied in this investigation did not have severely impaired lactose absorption. This was confirmed by the finding of only small amounts of lactic acid and total reducing substance in the stools after the tests. The lactic acid levels in the stools after lactose in the present study were very much lower than in children with established disaccharide intolerance (Weijers, van de Kamer, Dickie, and Ijsseling, 1961). Lactose absorption during the period of treatment was markedly increased in only two children with kwashiorkor.

Four normal Baganda subjects in the present study had clinical lactose intolerance (Anderson, Burke, Messer, and Kerry, 1966), which is a dose-dependent phenomenon, after the lactose load. Diarrhoea after oral lactose in kwashiorkor is well recognized (Dean, 1952), and high stool lactic acid levels after lactose have been reported (Bowie et al., 1965); this is probably, however, a reflection of the severity of the kwashiorkor. It seems likely that those children who get lactose intolerance either have severe kwashiorkor and a grossly damaged jejunal mucosa with severe secondary disaccharide deficiency, or an early onset of primary (specific) lactase deficiency which has been shown to occur in Baganda infants (Cook, 1967). Low jejunal lactase activity is very common in children who have recovered from mild to very severe kwashiorkor (Cook and Lee, 1966); it has been demonstrated in both Baganda and Bahutu (Rwanda) children after kwashiorkor. In Baganda children this is likely to be a result of the fall in lactase activity which normally occurs in the first three to four years of life and is probably genetically determined (Cook, 1967). Some, but not all, members of the Bahutu (Rwanda) tribe have lactase deficiency in adult life (Cook and Kajubi, 1966); the reason why children from that tribe all have low lactase levels after recovery from kwashiorkor of varying grades of severity is not clear. In view of the present findings it does not seem to be a result of mucosal damage during kwashiorkor, because the children studied in this investigation with mild to moderately severe kwashiorkor did not have a high incidence of lactase deficiency. It seems more likely that individuals, from either tribe, who will show a fall in lactase activity during infancy (Cook, 1967) are in some way predisposed to kwashiorkor. The mechanism of this is not clear.

SUMMARY

A good correlation has been found between the maximum rise in capillary blood glucose levels and the percentage absorption of carbohydrate (measured by estimating carbohydrate levels in the jejunum) during oral lactose and glucose + galactose tolerance tests.

Lactase deficiency and defective lactose absorption have been confirmed in normal Baganda subjects. Five children with mild to moderately severe kwashiorkor had less severely impaired lactose absorption.

Lactic acid and total reducing substance were estimated in the stools of the children with kwashiorkor during and after the tolerance tests; only small amounts were found.

We are indebted to the late Professor R. F. A. Dean for suggesting the investigation in the children with kwashiorkor. We are grateful to the nursing staff of Mulago Hospital and the M.R.C. Malnutrition Unit for help with the patients; to Mr. P. G. Ward for technical help with the stool analyses; and to Professor R. A. McCance, F.R.S. for reading the manuscript and for helpful suggestions.

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