Nutritional support of malnourished lactose intolerant African patients

S J D O'KEEFE, J K ADAM, E CAKATA, AND S EPSTEIN

From the Gastrointestinal Unit, Department of Medicine, University of Natal/King Edward VIII Hospital, Durban, South Africa

SUMMARY The effectiveness of two commonly available liquid diets was assessed in 40 severely malnourished black African patients. All patients were shown to have normal xylose absorption. The diets were given according to the manufacturer's recommendations. One diet was lactose containing (LC diet) (150 g/d) and high protein (112 g/d), the other normal protein and lactose free (LF diet) (protein 67 g/d), total energy content being similar. Patients were randomly divided into two equal groups and allocated (blind) to one of the diets. Tolerance and nitrogen balance were assessed over two three day periods on half and then full strength formulations. Severe intolerant symptoms were observed in 50% of patients on half strength and 94% of patients on full strength lactose containing diet with evidence of malabsorption of fluid, nitrogen, and fat. Despite high stool nitrogen losses (3.75 ± 1.04 g/d), however, positive nitrogen balance was achieved in most patients receiving the full strength LC formulation. On the other hand, the full strength LF diet was generally well tolerated and was associated with significantly lower faecal losses and positive nitrogen balance. The results indicate that high density lactose containing liquid formulae are poorly tolerated by severely malnourished black African patients, while lactose free formulae containing approximately 10 g nitrogen/d are well tolerated and result in positive nitrogen balance.

A recent survey has shown a high incidence of malnutrition among our hospitalised medical patients: over 80% of male and 50% of female patients had significant losses of protein and fat stores.¹ As most of these patients have relatively normal gut absorption, nutritional support can be based on enteral regimens. There are at present only two such regimens locally available that are suitable for fine nasogastric infusion; one being milk based and therefore lactose containing, the other lactose free. A further study of ours has revealed the population to be predominantly lactose intolerant,² with 90% of our medical ward patients showing a flat blood glucose response to oral lactose. Consequently, we might have anticipated problems with the milk based diet and designed the study to assess the practical importance of this.

PATIENTS

Forty malnourished male African patients requiring nutritional support were studied. They were randomly divided into two groups, one given lactose containing (LC) and the other given lactose free (LF) diets. Informed consent was obtained and patients were free to withdraw from study at any time. All patients were shown to have functional gastrointestinal tracts by normal xylose absorption tests – that is, increase in blood xylose greater than 1.67 mmol/l two hours after intake of 25 g xylose by mouth. Breakdown of disease patterns within each group is shown in Table 1. Respiratory infection predominated in keeping with our previous observation of a strong association between malnutrition and respiratory infection, in particular pulmonary tuberculosis.¹

Anthropometric measurements showed both groups to be significantly malnourished as compared with matched healthy black African controls¹ (Table 2). There were no significant differences in nutritional status between the two groups (TSF

Address for correspondence: Dr S J D O'Keefe, Gastrointestinal Unit, Groote Schuur Hospital, Observatory, Cape Town. 7925. Republic of South Africa.
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Table 1 Disease patterns

	LC diet group	LF diet group
Respiratory infections	10	13
Pulmonary tuberculosis	5	8
Abdominal diseases (details see below)	5	4
Cardiovascular diseases	2	1
Renal diseases	2	—
Haematological diseases	1	3

Details of abdominal diseases: LC group: two cirrhotic patients, one amoebic liver abscess, one unexplained jaundice, one hepatomegaly. LF group: one cirrhosis, one hepatomegaly, one carcinoma stomach, one acute pancreatitis.

4.9±1.9 mm in LC diet group; 4.2±2.8 mm in LF group).

DESIGN OF STUDY

Patients were studied while in a specific side ward section of the main medical ward. One of the investigators personally supervised the administration of the diets, patient symptoms, faecal, and urine collections. All patients were initially given half strength (manufacturer's recommendation, to allow for 'adaptation') diet for three days. For the second and third day, 24 hour urine and faecal collections were made in order to measure nitrogen balance. Tolerance to the diet was observed throughout the period together with a specific tolerance test as described below. From days four to six, full strength diet was given, and the above routine repeated on days five and six. Completeness of urine collections were checked by the consistency of the consecutive 24 hour creatinine contents.

DIETS

Composition

The two diets were given in the form recommended by the manufacturers and therefore not balanced to provide equal quantities of protein. Table 3 compares the composition. Vitamin and mineral content in both full strength formulations was

Table 3 Dietary composition (full strength daily intake)

	Medifeed (LC)	Ensure (LF)
Volume (ml)	2000	2000
Protein (g)	112	67
Energy (Kcal)	2108	1911
Carbohydrate (g)	280	262
Lactose (g)	150	0
Nitrogen (g)	17.6	10.4
Fat (g)	60	67
Osmolarity (mOsm/l)	524	362

sufficient to cover daily requirements [for details refer to Roussel Laboratories, Sandton, Johannesburg, 'Medifeed' (LC) and Ross/Abbott Laboratories, Aeroton, Johannesburg 'Ensure' (LF)]. Both diets were made up in the identical containers and were of similar appearance. Consequently patients were 'blind' as to which liquid they were receiving.

Diets were administered in the form of 400 ml drinks every two hours between 8 00 am and 4 00 pm under direct supervision. Normal ward diet was withheld apart from water.

Specific tolerance tests

A protocol similar to that of a glucose or lactose tolerance test was followed, except the test diet consisted of 400 ml of either half or full strength diet as described above. In the full strength formulation, this would contain 56 g total carbohydrate of which 30 g was lactose in the LC diet, and 52 g of non-lactose carbohydrate (mainly sucrose) in the LF diet. Blood samples were taken at 0 (fasting), 30, 60, 90, and 120 minutes after ingestion. Intolerant symptoms – for example, bloating, abdominal cramps, flatus, diarrhoea – were documented during the two hour test and continued over the following 24 hour period. The results were compared to the blood glucose response to 50 g glucose solution obtained in a previous study on similar patients.²

Table 2 Details of nutritional status of the two groups of patients studied (mean values ± SD)

	Age (yr)	Weight (kg)	Height (cm)	Triceps skinfold (mm)	Arm circumference (cm)	Plasma albumin (g/l)	Plasma globulin (g/l)
LC diet	45.9±13.5	55.7±11.5*	173.7±5.12	4.9±1.9*	22.7±3.6*	36.1±7.4	44.5±11.3
LF diet	39.2±13.4	48.1±9.5*	163.5±10.2	4.2±2.8*	21.3±3.4*	33.7±7.5	38.0±8.7
Controls†	41±11	73±13	170±7	9±4	28±3	35–50	22–34

* $p < 0.001$ unpaired Student's t test.

† See ref 1.

SAMPLE ANALYSIS

Urine and faeces

Nitrogen content was measured by micro-kjeldahl technique, fat content by standard saponification/alkali titration method. Blood glucose was measured after collection in oxalate containers, plasma separation, and analysis by glucose oxidase method.

Results

DIETARY TOLERANCE

(a) Symptoms (Table 4)

The lactose free diet was well tolerated at both half and full strength formulations by all but one patient who complained of nausea. On the other hand, the lactose containing diet was poorly tolerated. Fifty per cent of patients developed intolerant symptoms to the half strength formula while all but one developed similar complaints on the full strength formulation. Symptoms (in particular abdominal cramps and diarrhoea) were so severe in three patients that they could only complete 24 hours on the full strength formula.

(b) Blood glucose responses

Figures 1 and 2 summarise the changes in fasting blood glucose in response to an oral intake of 400 ml half or full strength diet. The mean peak incremental rise on half strength lactose free diet was 1.56 mmol/l, and 1.90 mmol/l on full strength. The latter result was similar to the rise after a similar quantity of glucose – that is, 2.2 mmol/l – in a similar group of patients reported previously.² The peak response to the full strength lactose containing diet was significantly lower ($p<0.05$, unpaired t test).

(c) Stool analysis

The accepted faecal nitrogen output is taken to be less than 1.5 g/24 h.^{3,4} Consequently the mean

Fig. 1 Blood glucose response to bolus administration of 400 ml of either half (—) or full strength (---) lactose free diets. 0 hrs = fasting.

excretion was increased in all study groups, ranging from 1.93 ± 1.12 g/24 h on half strength lactose free diet to 3.75 ± 2.03 g/24 h on full strength lactose containing diet (Table 5). Expressed differentially, the quantity of dietary nitrogen not absorbed was 38 and 21% respectively, normal value being 8%.⁴

Stool nitrogen losses were significantly higher ($p<0.001$, Student's unpaired t test) in patients receiving the lactose containing diet. There was no direct relationship between dietary nitrogen intake and faecal loss – for example, a nitrogen intake of 9.0 g in half strength lactose containing diet was associated with a mean nitrogen loss of 2.96 g/d while an intake of 10.1 g in full strength lactose free diet was associated with significantly less faecal nitrogen 2.16 g/24 h ($p<0.01$). On full strength lactose containing diet 80% – that is, 16 patients – had losses in excess of 3.0 g/24 h, while only 25% (five patients) on the lactose free diet had such losses.

A summary of the full analysis of stool output on full strength diets is shown in Table 6. The major

Table 4 Patient symptoms

	LF diet	LC diet
Half strength		
No complaints	19	6
Diarrhoea and flatulence	0	10
Nausea and bloating	1	1
Loose stool	0	1
Constipation	0	2
Full strength		
No complaints	19	1
Diarrhoea and flatulence	0	17
Nausea and bloating	1	2
Loose stool	0	0
Constipation	0	0

Fig. 2 As for Fig. 1, but response to 400 ml of half and full strength lactose containing diets.

difference was in stool volume, those given the lactose containing diet excreting on average over a litre of fluid per day, in keeping with their symptoms shown in Table 4. In addition to incomplete nitrogen and fluid absorption, fat malabsorption – that is, >5 g/24 h – was apparent in 18 patients on full strength LC diet and six patients on full strength LF diet.

(d) Nitrogen balance

Table 6 summarises the nitrogen balance results for half and full strength diets. It is evident that both half strength formulae were generally unable to maintain nitrogen equilibrium, unlike the full strength diets. Despite the high faecal nitrogen losses on the full strength lactose containing diet,

Table 6 24 hour stool analysis: on full strength formulae

	LC diet	LF diet
Weight	1118.5±711.2 g*	280.3±204.7
pH	5.5±0.5	6.0±0.6
Nitrogen	3.75±2.03 g*	2.3±2.1
Fat	24±9.6 g*	6.76±6.81

Mean values $\pm$ SD, and where * = $p < 0.05$ unpaired Student's *t* test.

and relatively high urinary excretion, nitrogen intake was sufficiently high to cover such losses, resulting in an estimated mean positive daily balance of 4.9 g nitrogen.

Discussion

The most important finding of the present study was that the lactose containing diet was poorly tolerated. Despite the fact that nitrogen balance could be achieved on the full strength formulation, symptoms were severe enough to preclude its general prescription to severely malnourished hospital patients.

The reason for poor tolerance to the lactose containing diet was most likely to be because of the high lactose content as almost identical symptoms were detected in a previous study when a similarly malnourished group of black patients were given oral lactose tolerance tests: 90% developed intolerant symptoms.² This was in keeping with previous observations of high incidence rates – that is, $>90\%$ – of primary lactose intolerance in the Bantu races.^{5,6} It was noteworthy that Bowie *et al*⁷ were able to show a dramatic drop in stool weight when milk diet was changed to a carbohydrate free diet. The quantity of lactose given in the lactose containing liquid diet in bolus form was 15 g (half strength) and 30 g (full strength formulation). Extrapolating from the literature, one would expect about 50% of normal individuals with intestinal lactase deficiency to develop symptoms from ingestion of half strength,⁸ and 90% to develop

Table 5 Nitrogen balance data for both groups of patients on half and full strength formulations (mean values $\pm$ SD) (based on 48 hour collections)

	LC diet		LF diet	
	Half strength	Full strength	Half strength	Full strength
Nitrogen in diet (g/d)	9.0	18.0	5.04	10.08
Nitrogen in urine (g/d)	6.1±1.8	9.35±3.93	5.3±2.2	5.4±2.3
Nitrogen in stool (g/d)	2.96±2.44	3.75±2.03	1.93±1.12	2.16±1.69
Nitrogen balance (g/d)	-0.06	+4.9	-2.19	+2.52

symptoms to the full strength lactose containing diet.⁹ These figures correspond very closely to our actual observations. Another explanation could be the higher osmolarity of the lactose containing diet. This is an unlikely contender, however, as the half strength lactose containing solution (osmolarity 262 mOsm/l) was associated with more intolerance than the higher osmolarity (362 mOsm/l) full strength lactose free solution.

It is of interest that Silk and coworkers noticed no significant intolerance to a similar lactose containing solution when it was given as a constant nasogastric infusion.¹⁰ As tolerance is related to the load of lactose given,¹¹ it remains possible that slow continuous infusion will be better tolerated. This has been attempted on our wards with poor results. Zulu patients strongly resist the insertion of nasogastric tubes as they believe it indicates the end is near. The origin of this misconception presumably arises from their observation of the tube feeding of terminal unconscious patients. On the other hand, they are very happy to take regular liquid drinks in the manner followed in our present study.

Subclinical intestinal malabsorption is well recognised in the tropics.¹²⁻¹³ Our observation of mildly impaired nitrogen and fat absorption in some patients is consistent with this hypothesis. The explanation is probably complex as it has been related to malnutrition *per se*,¹²⁻¹⁴⁻¹⁵ abnormal absorptive mucosa¹⁶⁻¹⁸ and subclinical tropical sprue,¹⁹ and also bacterial and parasitic overgrowth.²⁰⁻²³ Although detailed work up of intestinal absorption was not done (apart from xylose absorption tests) in the present study, we have previously reported a high incidence of small intestinal mucosal abnormalities in a similarly malnourished group of black Africans.²⁴ In the present study we noted a high incidence of fluid, nitrogen, and fat malabsorption in the group of patients given the lactose formula. Debongie *et al*²⁵ showed reduced protein, fat, calcium, magnesium, and phosphorus absorption by the ileum in lactose intolerant normal volunteers. Consequently it seems reasonable to ascribe the malabsorption seen in our lactose dietary group to the added insult of lactose maldigestion on an already compromised gastrointestinal tract. This is in keeping with previous reports of fat malabsorption in lactose intolerant patients after gastric surgery.²⁷⁻²⁸

Our main conclusion is that lactose free liquid formula diets containing approximately 10 g nitrogen and 2000 Kcal will provide adequate nutritional support for most of our malnourished adult African patients. On the other hand, high protein lactose containing regimens should be

avoided, even at half strength dilution, not so much for reasons of malabsorption, but more for patient comfort. It must be stressed that we cannot extrapolate our results to the normal population. Indeed milk products are part of the normal diet and remain very popular despite the high incidence of primary lactose intolerance.

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