Intestinal transit in healthy Southern Indian subjects and in patients with tropical sprue

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SUMMARY Whole gut transit was measured in a group of 21 healthy volunteers and 21 patients with tropical sprue by radio-opaque marker technique, using mean transit time single (MTTS) and single stool transit (SST) method. Mean SST in controls was 25.8 (1.4) (SE) hours, which is considerably shorter than in controls in temperate zones. Mean SST (23.7 (0.6) h) correlated significantly with average MTTS (24.9 (1.6) h) ($r=0.88$; $p<0.001$) confirming that SST is a valid method to measure intestinal transit in the tropics. Patients with tropical sprue had a mean SST similar to controls (24.4 (1.1) h), in spite of significantly higher faecal weights (580 (41.2) g v 252 (17.2) g; $p<0.001$).

Variability in intestinal transit in different population groups may have considerable epidemiological significance because gut transit has been shown to be important in regulating intestinal function.^{1,2} Many of the published reports of intestinal transit are now considered unsatisfactory because of inaccuracies introduced by a high degree of variability in the methods used.³ Though the possibility of changes in gut transit in patients with tropical sprue has been suggested,^{4,5} the role of transit in the diarrhoea of malabsorption is not fully understood. Therefore we measured whole gut transit in a group of patients with tropical sprue and a matched group of asymptomatic controls, using two methods, mean transit time single (MTTS) and single stool transit (SST), which have been validated by a continuous steady state marker excretion technique.^{6,7}

Methods

PATIENTS AND SUBJECTS

Twenty one healthy, asymptomatic adults (10 men and 11 women) aged 16-60 years (median age 35 years) and 21 patients with tropical sprue (12 men and nine women) aged 17-58 years (median age 32 years) from villages around Vellore, were studied.

The study lasted 14 days during which time the subjects lived in a metabolic ward. Throughout the study, all subjects were fed a controlled diet similar to the diet in rural areas. The basic daily diet of about 10.9 MJ consisted of 60 g of protein, 50 g of fat and 450 g of carbohydrate with 30 g of dietary fibre. All dietary intakes except fibre were calculated from *Nutrition value of Indian foods* (NIN, ICMR, Hyderabad) 1985.⁸ Fibre content of diet was measured as neutral detergent fibre.⁹ During the first week (equilibration period) all subjects were investigated as described elsewhere⁴ to confirm the diagnosis of tropical sprue and to exclude other conditions. None of the healthy volunteers gave a history of diarrhoea for the preceding two months and all had normal faecal fat¹⁰ (mean (SE), 3.6 (0.23); range 1.8-5.6 g/d) and vitamin B₁₂ absorption¹¹ (0.87 (0.09); range 0.2-1.5% dose/l). Five of the controls had low urinary xylose excretion¹² (26.3 (2.3)%; range 8.5-41.8%) which is part of tropical enteropathy found in apparently healthy asymptomatic southern Indian villagers¹³ and plays no role in nutrient absorption.¹⁴

MEASUREMENT OF WHOLE GUT TRANSIT

Mean transit time single (MTTS) and single stool transit (STT) were estimated during the second week (study period) while the subjects were in the metabolic ward by the methods of Cummings *et al.*^{6,7} Briefly, 20 markers each of a specific shape were

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Fig. 1 Correlation between average MTT-S and mean SST ($n=22$, $y=2.38 \times -31.48$, $r=0.88$, $p<0.001$).

given on three successive days with breakfast. The radio-opaque markers, barium sulphate impregnated polyethylene pellets, used in the study were cubes (mean weight (MW) 7.8 mg, specific gravity (SG) 1.25), small circles (MW 8.0 mg, SG 1.63), and large circles (MW 16.0 mg, SG 1.63). Every stool passed after giving the first dose of marker was collected in a polyethylene bag, the time noted, the stool x-rayed and the number of each marker present counted. This was continued until all the markers given were recovered.

Single stool transit was calculated from the

Table Whole gut transit in southern Indian volunteer controls and patients with tropical sprue

Method	Whole gut transit time (h) mean (SE)		
	Southern Indian volunteer controls	Tropical sprue	Western controls (Cummings et al Gut 1976; 17: 219-23)
Single stool transit (SST)			
Day 2	21.95 (0.45)	22.03 (0.46)	41.87
Day 3	26.08 (1.08)	24.35 (1.51)	52.93
Day 4	30.0 (3.06)	28.33 (2.55)	60.3
Mean-SST	25.77 (1.38)	24.44 (1.06)	
Average MTT-S*	26.74 (2.52)	23.08 (1.81)	66.06
Mean-SST†	24.21 (1.03)	23.16 (0.55)	

*Average MTT-S was done only on 11 patients with sprue and 11 matching controls; †Mean-SST for 11 subjects in each group in whom average MTT-S was estimated.

first stool passed after rising on days 2, 3, and 4 separately.⁷ In the majority of subjects there were no markers present in the stools on days 5 and 6, indicating almost complete marker excretion by day 4. Four of the subjects studied did not excrete markers in the first stool passed after rising on day 2, one on day 3, and five on day 4. Therefore, mean single stool transit (mean-SST) was calculated as the average of the SST values on days 2, 3, and 4 in each subject.

Whole gut transit time was also estimated by the mean transit time single (MTTS) method for each marker separately⁷ in 11 controls and 11 patients with tropical sprue. The average of the MTTS values for the three markers was calculated as average MTTS.

Informed consent was obtained from each subject and the protocol was approved by the Research Committee of the Institution. Statistical analysis was done by Student's *t* test and correlation and linear regression were done using the Epistat package for microcomputers.

Results

Single stool transit calculated from day 2, 3, and 4 stool correlated significantly with average MTTS ($p<0.001$ in all cases). Mean SST (23.7 hours (SE) 0.6, range 20.0-31.7 h) correlated significantly with average MTTS (24.9 h (1.6), range 15.5-46.88 h) in the 22 subjects (11 controls and 11 tropical sprue) in whom transit was measured by both methods ($r=0.88$; $p<0.001$) (Fig. 1). The correlation was still significant when each group of subjects were considered separately.

The transit time (mean SST) in controls (25.8 (SE) 1.4, range 20.9-45.8 h) was longer than in patients with tropical sprue (24.4 (1.1) h; range 20.0-42.7 h). But the differences were not significant for any of the calculated values, SST on day 2, 3, and 4, mean SST or average MTTS (Table).

Mean daily faecal wet weight in patients with tropical sprue (580 (41.2) g; range 348-898 g) was significantly higher than in controls (252 (17.2) g; range 142-454 g) ($p<0.001$). There was no correlation between faecal wet weight and transit times both in controls and patients with sprue (Fig. 2).

Discussion

The correlation between average MTTS and mean SST in the present study confirms that SST is a valid method to measure gut transit. When SST was calculated for each day, SST of day 4 correlated best with MTTS. Five of the 22 controls excreted no markers on day 4, however, and if there were no other data, transit could not have been calculated.

Fig. 2 Faecal output and mean SST in Southern Indian controls and patients with tropical sprue ($r=0.19$, $p=0.227$).

This is presumably related to the considerably shortened transit time in Indians compared with subjects in the West (Table). The results indicate that when SST method is used to measure gut transit in populations with a shorter transit than in temperate climates it is necessary to estimate mean SST.

This study as well as other published reports from India¹⁵⁻¹⁷ confirm that intestinal transit is considerably shorter in Indians compared with residents of temperate zones. Shortened transit time is primarily attributed to increased faecal bulk and high consumption of dietary fibre.^{1,18} The mean faecal weight of control subjects in the present study is high¹⁹ and may contribute to the shortened transit time. The dietary fibre intake in the present study (30 g/day estimated as neutral detergent fibre) is higher than in the United Kingdom,²⁰ and may also have contributed to the fast transit.

Although fibre has been shown to play an important role in shortening transit,¹⁸ its role in the tropics has been questioned.¹⁷ The role of other dietary factors like spices and environmental factors are not known. Poorly understood biological factors may also influence transit as it has been shown that Japanese living in Honolulu and eating a Western type of diet have the same short intestinal transit as Japanese living in Japan and consuming a high fibre diet.²¹ The differences in transit time between the studies from India¹⁵⁻¹⁷ may be because of dietary or biological differences between populations.

The control subjects in the present study have similar whole gut transit in spite of the wide range of faecal weights (142–454 g/day). Faecal bulk is considered to be an important factor in regulating gut transit.¹ Burkitt *et al*¹⁸ have reported that intestinal transit decreases rapidly when faecal weights increase from 20–30 g/day to about 120–200 g/day and a further increase causes little or no change in transit time. The lack of correlation between faecal weights

and transit time in controls may be because the faecal weights fall in the asymptotic part of the curve correlating faecal weights with transit times. It is usually believed that transit is the result rather than the cause of colonic events. Cummings *et al*,¹ however, have shown interesting evidence to support the view that transit itself influences events in the human large intestine and may independently modify the effect of dietary factors on the gut. If this view is taken, it could be assumed that transit time may be one of the major determinants of faecal weight. The study by Burkitt *et al*¹⁸ may then be interpreted to mean that the cause for the sharp increase in faecal weight from 30 to 200 g/day is the decrease in transit time from 120 h to 40 hours. At transit times below 40 hours, however, there is a wide variation in faecal weights (200–500 g/day) without a significant change in transit time. The mechanism by which the latter occurs is not clear. The relation between transit times and faecal weights of the control group in the present study is similar to the findings of Burkitt *et al*¹⁸ when they fed rural African adults a high fibre diet.

Whole gut transit was measured in patients with tropical sprue to investigate the role of transit in the pathogenesis of diarrhoea in these patients. Previous studies^{4,5} have shown that small intestinal transit is prolonged in patients with tropical sprue, probably secondary to stimulation of ileal brake mechanism by malabsorbed fats and proteins.²²

Although there is a striking increase in stool weights in tropical sprue, there is no change in whole gut transit time as compared with controls. This could be because of a combination of prolonged small intestinal transit in tropical sprue^{4,5} together with acceleration of colonic transit. This may not be the sole explanation, however, as small gut transit contributes to less than 15% of whole gut transit.²³ The study by Burkitt *et al*¹⁸ and our own data suggest, that at transit times below 40 hours, there may be a dilatation of the colon with accommodation of faeces giving rise to a wide variability of faecal weights with similar transit times. It is possible that the accommodation limit of the colon is not exceeded in the tropical sprue patients studied, resulting in transit times similar to controls. Further, the prolonged small intestinal transit in tropical sprue may play a complimentary role reducing the volume of material entering the colon per unit time. It may be difficult to determine the precise role of transit in the diarrhoea of tropical sprue as morphological and functional abnormalities of the colon have also been shown in this disease.²⁴

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