Response of the gullet to gastric reflux in patients with hiatus hernia and oesophagitis

$\mathbf{D.} \mathbf{A.} \mathbf{K.} \mathbf{WOODWARD}^{1}$

Oesophageal Laboratory, Queen Elizabeth Hospital, Birmingham 15

The variability of reflux symptoms in patients with hiatus hernia is well known, and a poor correlation between the symptoms of oesophagitis and appearances at oesophagoscopy has been noted.

A study has been made of the relationship between four variables. These are the acid reflux occurring in the oesophagus, the peristaltic clearance, the oesophagitis present and the symptoms from which the patients are suffering.

Symptoms were found only in patients with oesophageal pH levels below 4. However, such acid levels could occur in the absence of significant symptoms and there was a poor correlation between symptoms and various levels of pH below this.

Poor motor function in the oesophagus was associated with poor clearance of acid from it. Patients with poor clearance show a greater tendency to have oesophagitis, but, despite this correlation, in the group as a whole, there was no clear relationship between the results of the tests and the severity of symptoms.

Evidence from three sources suggests that the symptoms of gastro-oesophageal regurgitation in patients with hiatus hernia may be related to prolonged oesophageal contact with gastric contents after an episode of reflux, because of a sluggish response to the refluxed material. Donner, Silbiger, Hookman, and Hendrix (1966) showed that in patients with hiatus hernia and symptoms of oesophagitis, delayed clearance of acid barium was associated with abnormal peristaltic activity in the gullet. Booth, Kemmerer, and Skinner (1968) demonstrated that symptomatic hiatus hernia patients required more voluntary swallows to clear a bolus of hydrochloric acid from the oesophagus than did normals or patients with asymptomatic hiatus hernia. Olsen and Schlegel (1965) have shown that, in severe inflammatory oesophagitis, abnormal oesophageal motor function is demonstrable manometrically. Since emptying of the oesophagus must depend in part on oesophageal motility, patients with hiatus hernia have been studied with the object of demonstrating a correlation between the removal of refluxed acid from the gullet, the patients' symptoms, and oesophageal motor function.

METHOD

Thirty-three patients with a sliding hiatus hernia were studied. There were 14 men and 19 women with ages ¹Present address: Walsgrave Hospital, Coventry, CV2 2DX between 38 and 81 years. Patients with radiological or endoscopic evidence of oesophageal stricture were excluded because it was felt that this condition might produce mechanical interference with the dynamics of acid reflux or clearance.

ASSESSMENT OF SYMPTOMS Although patients with oesophageal stricture were excluded, dysphagia occurred in a few patients with functional disturbance of the oesophagus. The severity of reflux symptoms was graded by giving a total score of 0 to 8 to each patient, calculated by adding the scores (0, 1 or 2) for the severity of heartburn after food and after postural activity, and of regurgitation after food and after postural activity.

OESOPHAGITIS The severity of oesophagitis was assessed visually at oesophagoscopy in 26 patients and graded as follows: 0, no oesophagitis; A, mild oesophagitis, a reddening of the mucosa; or B, marked oesophagitis, a red mucosa which bled easily on instrumentation (Tables I and II).

OESOPHAGEAL MOTILITY This was studied with the patient supine by pressure recordings from two polyvinyl open tipped catheters with openings 5 cm. apart continuously perfused with water at the rate of 17 μ l./sec. by a pump. For the purpose of the *p*H test the distance of the cardia from the nares was measured by recording the position of the lower oesophageal intrinsic sphincter, or by identifying the gastro-oesophageal mucosal junction with a third catheter monitoring transmucosal potential differences. The motor function of the lower 15 cm. of the oesophagus was studied by recording its response to voluntary swallows. Pressures produced were registered by externally situated pressure transducers, the output from which was amplified and fed to a multichannel recorder with an event marker operated manually, when the patient executed an order to swallow.

Tests of oesophageal motility were performed in 27 patients. Abnormal motor function of the body of the oesophagus was said to be present if (1) over 25% of the responses to voluntary swallows were non-peristaltic in nature; (2) repetitive non-peristaltic contractions occurred after swallows or spontaneously; or (3) there was evidence of motor failure with rises in oesophageal pressure in response to voluntary swallows remaining persistently below 20 mm. Hg.

OESOPHAGEAL ACIDITY Acidity in the gullet was monitored in all patients by placing a pH electrode (Cambridge, Intestinal) via the nares 5 cm. above the cardia and recording pH continuously for a period of 15 hours after the evening meal. The patient lay supine with two pillows and no restriction to movement during sleep. Symptoms of gastro-oesophageal reflux were noted by the patient if they occurred during the test.

A 15-hour record of oesophageal pH was obtained for each patient and each record was analysed: (1) The total time for which the oesophageal pH was <5, <4, and <3 was measured and expressed as a percentage of the 15-hour period. (2) The pH trace was examined and the length of time measured for which oesophageal pH remained below basal levels after a drop of 2 pH units or more (signifying a reflux episode). The number of such episodes was recorded in three groups, of short, medium and long duration, lasting <15 minutes, 15-45 minutes and >45 minutes, respectively.

RESULTS

RELATION OF OESOPHAGEAL PH TO SYMPTOMS AND SEVERITY OF OESOPHAGITIS Tables I and II list the patients in the order of severity of acid reflux measured by the test. In nine (Table I), oesopha-

TABLE I REFLUX TO LESS THAN pH 5 AND GREATER THAN pH 4

Patient	Symptom Score	pН	% T	ime	Oesopha- goscopic Findings	Oesopha-	
		< 5	<4	< 3		geal Motility	
1	2	0.1	0	0	0	Normal	
2	2	0.3	Ó	Ó	0	Normal	
3	1	0.6	0	0	0	Normal	
4	3	1	0	0	-	Normal	
5	4	1.4	0	0	0	Normal	
6	2	2.0	0	0	B	Abnormal	
7	3	2.6	0	0	0	-	
8	2	7.6	0	0	0		
9	2	9.3	0	0	A	Normal	

 REFLUX TO LESS THAN pH 4 OR 3

 Patient
 Symptom Score
 pH % Time Oesophagoscopic Findings
 Oesophageal Motility

 10
 6
 2:7
 0.5
 0
 –
 Normal Abnorma

 11
 7
 3:2
 0:8
 0
 B
 Abnorma

 12
 8
 6:2:6
 1:1
 0
 0
 –
 Normal

TABLE II

10	6 7	2.7	0.2	0	-	Normal
11	7	3.2	0.8	0	В	Abnormal
12	8	6.2	1.1	0	0	
13	6	2.6	1.6	0	-	Normal
14	ă I	27	1.8	õ	Α	Normal
15	ŏ	20	2.5	ŏ	Ö	Abnormal
16	8 6 3 0 8 0 8 4 6 8 2 1	6	2.8	ŏ	-	Normal
17	ő	18	3.6	ŏ	_	
18	e e e e e e e e e e e e e e e e e e e	18	3.7	ŏ		Abnormal
19	0	19.6	9.6	ŏ	B	Abnormal
20	2	25	12	ŏ	D A	Abnormal
20	0	23	14		A 0	Abnormal
21	8	22	15	ò	U U	
22 23	2	9	1.4	1	0	Abnormal
23		14.4	4.4	1	0	Abnormal
24	6	12	5.2	1	0	Normal
25	3	18.5	9	1	В	Abnormal
26	2	14	7	1.5	-	Normal
27	4	5.6	3.3	1.8	0	Normal
28	6	12.5	8.3	1.8	Α	
29	6 3 2 4 6 6 2 0 8 6	6.4	4.4	2.8	0	Normal
30	2	21	12	3.2	Α	Normal
31	ō	38	19.3	4	0	Abnormal
32	Ř	38	23	4	Ā	
33	Ğ	66	30.6	5.3	B	Abnormal
					-	
	1	1				

geal pH did not fall below 4 during the test for a measurable period. Before the test these patients suffered only minor symptoms of gastro-oesophageal reflux (symptom score 1 to 4). Seven of the nine patients had absent or minor oesophagitis and only one had marked oesophagitis. In this patient it was felt that narrowing of the oesophagus produced by oesophagitis may have been preventing reflux.

Twenty-four patients exhibited acid reflux to levels less than pH 4, and 12 to less than pH 3 (Table II). The symptoms of patients refluxing to below pH 3 were no more severe than those refluxing only to below pH 4. In fact the severity

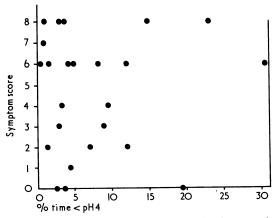


FIG. 1. There is not a constant relationship between the severity of symptoms and the degree of acid reflux to less than pH 4.

of reflux as assessed by the test showed a poor correlation with the severity of previously assessed symptoms of gastro-oesophageal reflux (Fig. 1). Similarly, severe reflux was not always associated with gross visual oesophagitis, one of the severest refluxers in the group (31) having no visual or histological oesophagitis.

OESOPHAGEAL MOTILITY AND SEVERITY OF REFLUX Of the 27 patients with motility studies, by the criteria defined 15 patients had normal and 12 abnormal oesophageal motility (Table III). Abnormal oesophageal motility was not associated with more severe symptoms of reflux though the three patients with dysphagia all occurred in this group.

TABLE III

RESULTS OF OESOPHAGEAL MOTILITY TESTING

	Pressure in Lower					
Patient	Oesopha- geal Sphincter (mm. Hg)	Total	No Res- ponse	Peri- stal- tic	Non Peri- stal- tic	Max. Pressure (mm. Hg)
Normal 1 2 3 4 5 9 10 13 14 16 24 26 27 29 30	0 4 9 11 12 7 20 6 0 3 4 18 7 2 13	15 21 33 10 21 24 15 18 25 17 18 20 20 32 13	0 3 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	12 18 33 10 16 24 11 16 25 17 14 20 15 19 13	300050300040580	50 40 40 30 35 25 60 40 40 30 40 25 45 30 40
Abnormal 6 11 15 18 19 20 211 22 ² 23 25 ¹ 33	0 7 0 0 0 6 3 4 3 2	14 33 14 25 5 16 16 16 16 14 26 11 18	4 12 12 0 0 0 0 0 0 0 0 0 0	0 5 0 25 0 0 8 7 13 0 16	10 16 2 0 5 16 8 7 13 11 2	$ \begin{array}{r} 14\\10\\15\\15\\25\\>10\\8\\35\\>100\\45\\30\end{array} $

¹ Also showed repetitive and spontaneous contractions suggestive of spasm.

^a Abnormal motility detected radiologically ^a Picture of diffuse spasm shown radiologically

As judged by the percentage time pH in the gullet was less than 5 or 4, patients with abnormal motility were exposed to acid for longer than patients with normal motility, though the differences were not statistically significant (Table IV).

The mean sphincter strength was significantly greater in patients with normal motility than in those with abnormal motility. Nevertheless, the former group were as prone to reflux episodes as the latter (Table IV).

	Т	Α	В	L	Ε	Ι	V	
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SEVERITY OF REFLUX AND OESOPHAGEAL MOTILITY

			Oesophag	geal Motility	Significance
			Normal Abnormal		of Difference
No. of patients			15	12	
Oesophageal acidity % time pH < 5					
Mean . SD		::	10·3 13·3	20·1 17·1	0.05 < P < 0.1
6D	4 		4·3 3·3	7·8 5·9	0.02 < P < 0.1
Reflux episodes Sphincter strengt	h.	••	171	90	—
(mm. Hg) Mean . SD	•	::	7·7 5·9	3·3 3·9	P < 0.025 (t test)

The patients with abnormal motility were exposed to acid for longer than patients with normal motility, though the differences are not statistically significant.

DURATION OF REFLUX EPISODES Table V relates oesophageal motor function to short, medium, or long duration reflux episodes in each patient. Short duration episodes, with oesophageal pHremaining low for periods lasting less than 15 minutes, occurred equally in patients with abnormal as in those with normal oesophageal

 TABLE V

 DURATION OF REFLUX EPISODES AND OESOPHAGEAL

 MOTILITY

	Oesophag	a :		
	Normal	Abnormal	Significance of Difference	
No. of patients	15	12	_	
No. of patients with reflux episodes lasting <15 min 15-45 min >45 min	15 6 1	10 9 11	$\chi^{a} 2.042;$ P > 0.3 $\chi^{a} 16.217;$ P < 0.001	

Patients with abnormal motility showed a significant increase in prolonged reflux episodes.

motility. Slower clearance of acid in times of between 15 and 45 minutes occurred in a higher proportion of patients with abnormal than with normal oesophageal motility tests, though the difference was not significant. However, all but one of the 12 patients with abnormal motility demonstrated episodes when oesophageal pH remained low for longer than 45 minutes, whereas this occurred in only one of the 15 patients with normal motility, a statistically significant difference.

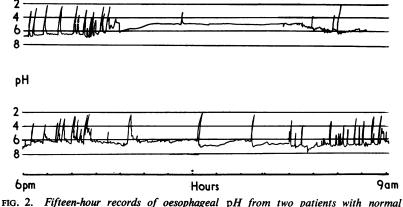


FIG. 2. Fiftee-nour records of oesophageal pH from two patients with normal oesophageal motility. Frequent reflux episodes are shown with rapid return of pH to neutrality on each occasion. In the nocturnal period, reflux episodes are much less frequent in both patients. In the upper record the slow fall in pH of < 2 units occurred when the patient went to sleep, and the level of acidity reached was not sufficient for pepsin activation.

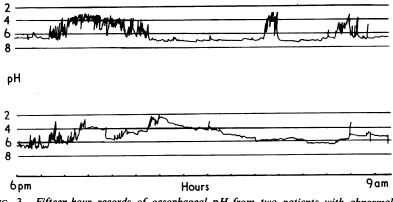


FIG. 3. Fifteen-hour records of oesophageal pH from two patients with abnormal oesophageal motility.

Upper: a patient with severe visual oesophagitis and motor failure.

Lower: No visual oesophagitis present. Manometric and radiological evidence of diffuse spasm.

In both patients prolonged periods of low pH are maintained, and there is no regular pattern of the clearance of acid from the gullet.

Thus the patients with normal motility tended to show a rapid return of pH to neutrality after reflux, whereas though this feature was also present in patients with abnormal motility, they demonstrated consistently occasions when a low pH was maintained for a long period. The mean duration of these prolonged reflux episodes was 126 minutes with a range of 45 to 300 minutes. These features are demonstrated in Figures 2 and 3. OESOPHAGEAL MOTILITY AND SEVERITY OF OESOPHA-GITIS Of the 12 patients with abnormal oesophageal motility, 11 underwent oesophagoscopy. All 5 patients with marked inflammatory oesophagitis showed abnormal oesophageal motility. However, 6 patients with abnormal motility showed little or no evidence of oesophagitis.

Oesophagoscopy was performed on 11 of the 15 patients with normal oesophageal motility and oesophagitis was grade 0 in 8 and grade A in 3.

DISCUSSION

Many workers have used the pH electrode as a diagnostic and qualitative measure of the reflux of acid into the gullet in patients with hiatus hernia. With the electrode at diaphragmatic level Weber and Gregg (1959) found that normal patients demonstrated increased acidity in full expiration. For this reason in this study the electrode has been placed 5 cm. above the cardia. With an electrode at this level Tuttle, Rufin, and Bettarello (1961) defined reflux as occurring when the pH fell below 4 and, recording for a period of between 30 and 60 minutes, they found a good correlation between reflux and the sensation of heartburn. pH levels are only an index of acid concentration, but in this study, by recording over a long period and measuring the length of time under certain pH levels, we have obtained a guantitative index of gastro-oesophageal reflux for comparison with the patients' continuing symptoms.

In our patients with minor symptoms of reflux. pH did not fall below 4 for a significant period. Patients, however, with reflux to less than pH 4had variable symptoms, on the whole poorly related to the quantitative estimation of acid reflux. Symptoms at the time of the test were not a marked feature and not always associated with a reflux episode. The variability of reflux symptoms in patients with hiatus hernia obviously depends on whether or not gastric juice is entering the gullet, but our findings suggest that even when this occurs the patient's symptom response may be very variable. This unpredictable behaviour of patients with hiatus hernia is emphasized by Palmer (1968), who found that only 8.9% of 1,011 patients with hiatus hernia followed up for 20 years had classic symptoms of gastro-oesophageal reflux. Furthermore, only a quarter of his patients with visual oesophagitis had significant complaints. Skinner (1966) also reported a poor correlation between patients' symptoms and oesophagoscopic appearances. Our findings suggest that when electrode monitored reflux is slight, oesophagitis is usually mild or absent, but that when it is severe the degree of oesophagitis can be very variable.

In this study acid reflux has been recognized when a sudden drop of pH to 2 or more units below the basal level occurred, and the duration of a reflux episode as the time taken for pH to return to this level. The lowest pH reached depends on the concentration of hydrogen ions

in the refluxed material, and the subsequent pHchange is related to alteration in strength of the acid solution around the electrode produced by the buffering action of oesophageal secretion and swallowed saliva. Since mixing around the electrode is slow, the duration of a reflux episode probably depends primarily on bulk emptying of the oesophagus produced by contractile activity. It is likely that the reflux of small quantities of acid around the electrode produces a sharp fall in pH followed by an equally abrupt rise due solely to the buffering effect of oesophageal resting secretions. On the records this type of reflux episode has the appearance of a sharp spike without any width (Fig. 2). Examination of the results of Tuttle et al. (1961) supports this suggestion because rapid fluctuations in pH occur without any simultaneous manometric evidence of oesophageal contractions.

The clearance of larger amounts of gastric juice probably depends on secondary peristaltic activity induced by the refluxed bolus, as demonstrated by Creamer (1955). However, it is possible that this motor response is related to the volume refluxed because Nagler and Spiro (1961) found that secondary peristalsis occurred only in patients in whom relatively large volumes could be aspirated from the gullet after reflux induced by abdominal compression. This is important because it suggests that peristaltic activity might not remove small amounts of refluxed gastric juice from the oesophagus. Our records, however, fail to demonstrate slow falls of pH which might be expected to occur if small amounts of gastric juice oozed past an incompetent cardia at night, as suggested by Aylwin (1953). In several patients a slow fall of less than 2 pH units occurred during sleep, but the range of peptic activity was not reached. It was felt that this slight fall of pHduring sleep was due to the decreased amount of swallowed saliva rather than to reflux of acid gastric juice (Fig. 2).

Creamer (1955) suggested, in patients with the symptoms of oesophagitis, that the secondary peristaltic response following reflux was sometimes inco-ordinated, and Donner and his associates (1966), studying similar patients radiologically, found that swallowed acid barium was cleared abnormally slowly from the gullet because of the presence of motor abnormalities in the lower oesophagus.

The clearance of refluxed acid must also depend on primary peristaltic activity produced by a swallow. Tuttle (Tuttle and Grossman, 1958; Tuttle, Bettarello, and Grossman, 1960) states that dry swallows have little effect on an acid pHexisting in the lower oesophagus, and even wet swallows have only a transitory effect lasting 5 to 30 seconds. However, since their patients were examined in the supine position, prolonged low pH levels may have been due to reflux following relaxation in the lower oesophageal intrinsic sphincter produced by swallowing. The work of Booth et al. (1968) more specifically suggests that the efficiency of the primary peristaltic wave may be at fault in hiatus hernia patients with the symptoms of oesophagitis because they required more voluntary swallows to raise oesophageal pH to normal following the experimental instillation of hydrochloric acid than did asymptomatic hiatus hernia patients or normals.

Thus altered primary and secondary peristaltic activity may be responsible for delayed clearance of acid in symptomatic patients. Our results suggest that delayed clearance of acid is much commoner in patients with abnormal oesophageal motility. The relationship of delayed clearance of acid to symptoms is, however, less clear since 2 of our 12 patients with abnormal motility and slow acid clearance had no symptoms and 4 only minor symptoms of gastro-oesophageal reflux. In the patients with abnormal motility the severity of symptoms did not appear to have any relationship to the degree of oesophagitis, though the numbers are too small to be of any significance.

It has been suggested (Ingelfinger, 1963; Tuttle *et al.*, 1961) that for the symptoms of gastro-oesophageal reflux to occur a 'fertile soil' must be prepared. The absence of symptoms in many of our patients with demonstrable acid reflux, and particularly the presence of prolonged reflux in patients with abnormal motility suggests that the latter may allow sufficient exposure of the gullet to refluxed acid to cause damage to the mucosa and the subsequent development of symptoms.

All of our patients with severe visual oesophagitis displayed abnormal oesophageal motor function, and this finding agrees with those of Olsen and Schlegel (1965). However, their suggestion that abnormal oesophageal motility is secondary to the oesophagitis is not completely supported by our findings, because over half of our patients with abnormal motility had absent or minor oesophagitis. This finding would support the suggestion that abnormal motor function in some patients may be the initiating factor allowing refluxed acid to remain in the oesophagus long enough to produce inflammatory oesophagitis.

CONCLUSIONS

A history of significant symptoms of reflux was usually associated with the occurrence, for a measurable time, of oesophageal acid levels of less than pH 4. However, pH levels of less than 4 could occur in the absence of a history of symptoms of acid reflux, and there was a poor correlation between symptoms and measured acid levels at less than pH 4.

Normal oesophageal motility was associated with rapid clearance of refluxed acid from the gullet. Long periods of low pH after reflux occurred in patients with abnormal oesophageal motility, suggesting that bulk removal of refluxed acid was impaired in this group.

Such delayed acid clearance was associated with the presence of oesophagitis, but, although there was a correlation between these two, they were not together as a group related to the symptomatic state of the patients.

It is suggested that in some patients with hiatus hernia and reflux abnormal motor function of the oesophagus may predispose to the development of inflammatory oesophagitis because of delayed clearance of refluxed gastric juice.

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