ARTIGO ORIGINAL / ORIGINAL ARTICLE

INTRODUCTION

Chagas’ disease is the consequence of infection by the flagellate protozoan Trypanosoma cruzi that causes mainly alterations in the esophagus, colon and heart(15, 22). There is degeneration and reduction of the neurons of the myenteric plexus(15, 16, 22) causing esophageal motility abnormalities similar to those of idiopathic achalasia, absent or partial lower esophageal sphincter relaxation and loss of peristaltic contractions in the esophageal body(5, 22). Most patients do not have these advanced manifestations of the disease, but even patients without dysphagia and with a normal radiologic examination may have some motility alterations(1, 6).

During the aging process there is a loss in the number of neurons in the gut myenteric plexus(19, 22), causing esophageal motility abnormalities similar to those of idiopathic achalasia, absent or partial lower esophageal sphincter relaxation and loss of peristaltic contractions in the esophageal body(5, 22). Most patients do not have these advanced manifestations of the disease, but even patients without dysphagia and with a normal radiologic esophageal examination may have some motility alterations(1, 6).

Chagas’ disease is transmitted more frequently during childhood(16). In most patients with esophageal manifestations of the disease the clinical, radiologic and manometric features remain unchanged for years(20). It is possible that with the aging process the esophageal motility alterations of patients with non-advanced disease become more intense.

The objective of this study was to evaluate the esophageal motility impairment caused by Chagas’ disease in two age groups. Our hypothesis was that older patients with non-advanced esophageal motility impairment caused by Chagas’ disease have more intense motility alterations than younger patients.

MATERIAL AND METHODS

We studied 30 patients with dysphagia for solids and liquids, no regurgitation, esophageal radiologic examination with barium retention, slow transit, esophageal diameter less than 4 cm, and a positive serologic test for Chagas’
disease. All had an epidemiologic history of having lived in endemic areas of the disease during some time in childhood. Endoscopic examination did not detect alterations in the esophageal mucosa. The patients (13 men and 17 women) were divided into a younger group, aged 34 to 59 years, median 51 years (mean ± SD: 50 ± 8 years, n = 15), with a duration of dysphagia of 1 to 40 years, median 5 years, and an older group aged 61 to 77 years, median 66 years (mean: 67 ± 5 years, n = 15), with a duration of dysphagia of 1 to 60 years, median 10 years. They had not received previous treatment by pneumatic dilatation or esophagomyotomy and were living with mild dysphagia, eating slowly soft food, and sometimes drinking water during meals. The intensity of dysphagia was similar in the two groups. Fourteen patients had duration of dysphagia of 1 to 5 years (median: 3 years), and 16 patients had duration of dysphagia of 10 to 60 years (median: 16 years).

The control group consisted of 22 subjects, 5 men and 17 women, also divided into a younger group aged 33 to 58 years, median 42 years (mean: 44 ± 9 years, n = 15) and an older group aged 61 to 73 years, median 66 years (mean: 66 ± 4 years, n = 7). The controls were submitted to esophageal manometry during the investigation of gastroesophageal reflux disease, and had normal esophageal endoscopic and radiologic examinations. The protocol of manometric examination was approved by the Human Research Committee of the University Hospital of Ribeirão Preto, SP, Brazil.

The manometric examination was performed with a round eight-lumen polyvinyl catheter with an outer diameter of 4.5 mm and an inner diameter of 0.8 mm (Arndorfer Specialities, Inc, Greendale, Wisconsin, USA). The four proximal lateral openings of the catheter were spaced 5 cm apart at 90° angles. They were connected to external pressure transducers (pvb Medizintechnik GmbH, Kirelseeon, Germany), which in turn were connected to a PC Polygraph HR (Synectics Medical, Stockholm, Sweden). The manometric signals were stored in a computer. During the manometric recordings, a minimally compliant pneumohydraulic pump (JS Biomedicals, Ventura, CA, USA) perfused distilled water at 0.5 mL/min through each lumen.

All individuals were studied in the supine position after 12 h of fasting. The catheter was introduced through the nose. For the study of esophageal contractions, the catheter was positioned with the proximal opening located 2 cm below the upper esophageal sphincter (UES) and the other openings located at 7, 12 and 17 cm from the UES. Five swallows of a 5-mL bolus of water were performed, with an interval of at least 20 seconds between successive swallows.

Using the computer Polygram Upper GI software version 6.4 (Gastrosoft, Stockholm, Sweden) we measured the amplitude and duration of the contractions and the time interval between the upstroke of the contractions measured 5 cm apart. Contractions separated less than 1 second at recording sites 5 cm apart were considered to be simultaneous. Non-conducted contractions occur when there is no contraction propagation between two points of the esophageal body.

Data were analyzed statistically by analysis of variance, the Tukey-Kramer multiple comparisons test, the unpaired t test, and the Fisher exact test. Differences were considered to be significant when P < 0.05. The results are reported as mean ± SEM unless otherwise indicated.

**RESULTS**

Comparison between patients with Chagas’ disease and controls revealed a lower amplitude of contractions in patients at 7, 12 and 17 cm from the UES (P <0.01, Table 1). Patients with Chagas’ disease had fewer peristaltic, more simultaneous and more non-conducted contractions than controls (P <0.05, Figure 1). There were no differences between patients and controls in duration of contraction or velocity of peristaltic contractions (P >0.05).

**FIGURE 1** – Percentage of esophageal peristaltic, simultaneous and non-conducted contractions measured from 2 to 7 cm, from 7 to 12 cm and from 12 to 17 cm below the UES after swallows of a 5 mL bolus of water. * P <0.05 vs younger and older patients with Chagas’ disease; ** P <0.05 vs older patients with Chagas’ disease.
Older patients with Chagas’ disease had lower amplitude of contractions at 17 cm from the UES (30.8 ± 4.3 mm Hg) than younger patients (51.9 ± 8.6 mm Hg, P < 0.05, Figure 2), and had more non-conducted (41%) and fewer peristaltic (8%) contractions from 12 to 17 cm than younger patients (non-conducted: 16%, peristaltic: 21%, P = 0.05, Figure 1). There were no differences between younger and older patients in duration or velocity of contractions (P > 0.05, Table 1).

There was no difference between younger and older controls (P > 0.05). There were also no differences between patients with duration of dysphagia of less than 6 years (n = 14) and patients with duration of dysphagia of more than 10 years (n = 16) (P > 0.05, Table 2).

**DISCUSSION**

The lower amplitude of contractions, and the larger number of simultaneous and non-conducted contractions found here in patients with Chagas’ disease compared to controls are in agreement with previous reports (1, 3, 4, 5, 22). Older patients with Chagas’ disease with clinical and radiologic examination similar to that of younger patients had a lower amplitude of contractions and more non-conducted contractions in the distal esophagus than younger patients, showing a decrease in the capacity of contractions in the distal esophagus of older patients compared to younger patients.

The differences between younger and older patients with Chagas’ disease may be the result of the progression of the disease along the lifetime. However, only a small number of patients have this evolution and most of them remain stable for a long time (1, 14, 20) before reaching advanced ages.

The human myenteric esophageal neurons decrease in number in older subjects without esophageal diseases (5, 16, 19, 27), an event which is accompanied by an increase in neuron size (19). These alterations in neurons may explain the significant decrease in esophageal pressure amplitude with aging reported mainly in the eighth and ninth decades (6, 13, 24) and aperistalsis in older subjects (6, 11, 21, 23). The enteric sensory neurons are particularly susceptible to neurodegeneration with age (27). They exhibit neurodegeneration to a greater extent than other phenotypes, serotonergic interneurons or nitrergic inhibitory motor neurons (27). The nitrergic inhibitory motor neurons (4), and probably sensory neurons (9), are affected by Chagas’ disease. When a patient has megaesophagus the myenteric plexus is almost completely absent, but in patient with no dilatation the myenteric plexus destruction is only partial (15).

Deterioration of esophageal motility of patients over 60 years of age compared with patients under 60 years may not be enough to cause clinical or radiologic differences between them. According to our radiologic classification into grades I to IV (28), they were grade I, corresponding to the less severe stage of the disease. Even asymptomatic patients (40) or patients with normal radiologic examination (1) may have alterations in esophageal motility. The severity of esophageal alterations in

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**TABLE 1** – Amplitude, duration and velocity of esophageal contractions after wet swallows in younger (n = 15) and older (n = 15) patients with Chagas’ disease, and younger (n = 15) and older (n = 7) control subjects, measured at 2, 7, 12 and 17 cm from the upper esophageal sphincter (mean ± SEM).

<table>
<thead>
<tr>
<th>Amplitude (mm Hg)</th>
<th>Controls</th>
<th>Younger</th>
<th>Older</th>
<th>Chagas’ disease</th>
</tr>
</thead>
<tbody>
<tr>
<td>2 cm</td>
<td>95.5 ± 11.7</td>
<td>46.3 ± 9.1</td>
<td>85.7 ± 9.8</td>
<td>45.4 ± 10.0</td>
</tr>
<tr>
<td>7 cm</td>
<td>49.8 ± 16.1</td>
<td>38.8 ± 5.0</td>
<td>74.6 ± 12.0</td>
<td>72.8 ± 9.8</td>
</tr>
<tr>
<td>12 cm</td>
<td>78.3 ± 9.4</td>
<td>28.4 ± 2.9</td>
<td>59.7 ± 4.9</td>
<td>51.9 ± 8.6</td>
</tr>
<tr>
<td>17 cm</td>
<td>63.3 ± 9.5</td>
<td>23.5 ± 2.3</td>
<td>27.3 ± 3.8</td>
<td>30.8 ± 4.3</td>
</tr>
</tbody>
</table>

**Duration (seconds)**

<table>
<thead>
<tr>
<th>Duration (seconds)</th>
<th>Controls</th>
<th>Younger</th>
<th>Older</th>
<th>Chagas’ disease</th>
</tr>
</thead>
<tbody>
<tr>
<td>2 cm</td>
<td>2.1 ± 0.3</td>
<td>3.4 ± 0.2</td>
<td>3.4 ± 0.3</td>
<td>3.6 ± 0.3</td>
</tr>
<tr>
<td>7 cm</td>
<td>2.7 ± 0.3</td>
<td>3.7 ± 0.4</td>
<td>3.8 ± 0.4</td>
<td>4.2 ± 0.4</td>
</tr>
<tr>
<td>12 cm</td>
<td>2.4 ± 0.2</td>
<td>3.4 ± 0.2</td>
<td>3.4 ± 0.3</td>
<td>3.6 ± 0.3</td>
</tr>
<tr>
<td>17 cm</td>
<td>1.9 ± 0.2</td>
<td>2.9 ± 0.2</td>
<td>3.1 ± 0.2</td>
<td>3.2 ± 0.2</td>
</tr>
</tbody>
</table>

**Velocity (cm/s)**

<table>
<thead>
<tr>
<th>Velocity (cm/s)</th>
<th>Controls</th>
<th>Younger</th>
<th>Older</th>
<th>Chagas’ disease</th>
</tr>
</thead>
<tbody>
<tr>
<td>2 cm → 7 cm</td>
<td>1.8 ± 0.3</td>
<td>2.3 ± 0.3</td>
<td>2.6 ± 0.4</td>
<td>3.5 ± 0.3</td>
</tr>
<tr>
<td>7 cm → 12 cm</td>
<td>1.7 ± 0.2</td>
<td>2.9 ± 0.1</td>
<td>3.2 ± 0.3</td>
<td>3.3 ± 0.3</td>
</tr>
<tr>
<td>12 cm → 17 cm</td>
<td>1.8 ± 0.2</td>
<td>2.2 ± 0.2</td>
<td>3.5 ± 0.3</td>
<td>3.5 ± 0.3</td>
</tr>
</tbody>
</table>

**TABLE 2** – Results of distal esophageal manometric examination in patients with a duration of dysphagia of 1 to 5 years (median 3 years, n = 15) and in patients with a duration of dysphagia of 10 to 60 years (median 16 years, n = 16) (mean ± SEM).

<table>
<thead>
<tr>
<th>Duration of dysphagia</th>
<th>Amplitude (mm Hg)</th>
<th>Duration (seconds)</th>
<th>Simultaneous (%)</th>
<th>Non-conducted (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 to 5 years</td>
<td>39.1 ± 8.0</td>
<td>4.1 ± 0.4</td>
<td>50.0</td>
<td>54.0</td>
</tr>
<tr>
<td>10 to 60 years</td>
<td>31.9 ± 4.8</td>
<td>3.5 ± 0.3</td>
<td>61.7</td>
<td>28.3</td>
</tr>
</tbody>
</table>

**DISCUSSION**

The lower amplitude of contractions, and the larger number of simultaneous and non-conducted contractions found here in patients with Chagas’ disease compared to controls are in agreement with previous reports (1, 3, 4, 5, 22). Older patients with Chagas’ disease with clinical and radiologic examination similar to that of younger patients had a lower amplitude of contractions and more non-conducted contractions in the distal esophagus than younger patients, showing a decrease in the capacity of contractions in the distal esophagus of older patients compared to younger patients.

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A comparação entre pacientes mais jovens e mais idosos com doença de Chagas sugeriu que o envelhecimento pode levar a alterações motoras do esôfago após a doença. As alterações motoras do esôfago podem incluir perda de neurônios do plexo mientérico do esôfago, que é um efeito comum à doença de Chagas e ao envelhecimento. Estudos anteriores mostraram que a incidência de achalasia idiopática aumenta com a idade, sugerindo que também na forma idiopática de achalasia, o processo de envelhecimento pode influenciar a evolução da doença.

Em um estudo prévio, foi observado que pacientes com doença de Chagas e disfagia, com pareceres radiológico e clínico similares, tiveram menores amplitudes das contrações do esôfago do que os controles. As contrações não propagadas (41%) e menos contrações peristálticas (8%) foram observadas nos pacientes mais idosos com doença de Chagas, em comparação com os pacientes mais jovens.

Em resumo, a doença de Chagas e o envelhecimento podem levar a alterações motoras do esôfago, com a idade influenciando o processo de envelhecimento e a evolução da doença de Chagas. O envelhecimento pode levar a uma diminuição da intensidade dos esforços motores do esôfago, que podem afetar negativamente o tratamento e a evolução da doença.

**RESUMO**

- **Objetivo**: Avaliar em duas faixas etárias as possíveis alterações da motilidade do esôfago em pacientes com doença de Chagas. A hipótese é de que o envelhecimento pode provocar aumento da intensidade das alterações motoras do esôfago consequentes à doença de Chagas.

- **Pacientes e Método**: Estudou-se a motilidade do esôfago em 30 pacientes com doença de Chagas e disfagia, com pareceres radiológico e clínico similares. Os pacientes eram divididos em dois grupos: 15 pacientes mais jovens (mediana: 51 anos) e 15 pacientes mais idosos (mediana: 66 anos). Como grupo controle, estudaram-se 15 pessoas com idades entre 33 e 58 anos (mediana: 42 anos) e 7 pessoas com idades entre 61 e 73 anos (mediana: 66 anos). As contrações esofágicas foram medidas pelo método manométrico, com perfusão contínua, após cinco deglutições de 5 mL de água, a 2, 7, 12 e 17 cm distais ao esfíncter superior do esôfago.

- **Resultados**: Pacientes com doença de Chagas tiveram menor amplitude das contrações, menor proporção de contrações peristálticas, maior proporção de contrações simultâneas e maior proporção de contrações não propagadas do que os controles. Os pacientes com doença de Chagas e mais idade tiveram menor amplitude das contrações em esôfago distal (média ± EP: 30,8 ± 4,3 mm Hg) do que os pacientes com menos idade (51,9 ± 8,6 mm Hg). Entre 12 e 17 cm do esfíncter superior do esôfago, os pacientes com mais idade tiveram mais contrações não propagadas (41%) e menos contrações peristálticas (8%) do que os pacientes com menos idade (não propagadas: 16% peristálticas 21%).

- **Conclusão**: Pacientes com doença de Chagas mais idosos, com pareceres radiológico e clínico similares ao de pacientes mais jovens, têm alterações motoras do esôfago que sugerem que o envelhecimento pode provocar comprometimento adicional à motilidade do esôfago.

REFERENCES