Peroral Endoscopic Myotomy Versus Heller Myotomy for Achalasia: Pros and Cons

Cristian Tefas$^{1,2}$, Rami Ababneh', Marcel Tanţău$^{1,2}$

$^1$Gastroenterology Department, “Prof. Dr. Octavian Fodor” Institute of Gastroenterology and Hepatology, Cluj-Napoca, Romania
$^2$“Iuliu Haţegianu” University of Medicine and Pharmacy, Cluj-Napoca, Romania

Abstract
Achalasia is an esophageal motor disorder that has multiple endo-
scopic and surgical methods of treatment. However, there is no consensus on optimal therapy in patients suffering from this disorder. This review discusses two therapies with similar but technically different concepts, peroral endoscopic myotomy and Heller surgical myotomy. After a brief introduction to the basic problems of achalasia, technical considerations, intra and postprocedural complications are presented and the advantages and disadvantages of the two procedures are discussed, based on the relevant articles in the literature. Heller surgical myotomy and peroral endoscopic myotomy appear to be similar in performance with similar results in terms of gastro-esophageal reflux rates. However, with experience being gained in the field of endoscopic myotomy, this procedure seems more advantageous, with similar success rates to those of the established surgical technique, but offering shorter operating times, shorter hospitalizations and, ultimately, lower costs.

Key words: achalasia, peroral endoscopic myotomy, Heller myotomy

Introduction

Achalasia is a motility disorder of the esophagus defined by three elements: the reduction or absence of the primary peristaltic waves in the distal two thirds of the esophagus, incomplete or no relaxation of the lower esophageal sphincter (LES) during swallowing and increased resting LES tone. Typical symptoms of achalasia are dysphagia, nonacid regurgitation, retrosternal pain, and weight loss. Achalasia is reported around the world with no gender differences, predominantly between the ages of 25-60 (1), affecting about one person in 100000 (2).

Therapies that have proven effective address the LES and they include laparoscopic Heller myotomy (LHM), endoscopic injection of botulinum toxin, endoscopic balloon dilatation, and peroral endoscopic myotomy (POEM). However, there is no consensus on a standard treatment in achalasia, the therapeutic option being chosen for each patient according to their preferences, age, co-morbidities, previous therapies, and the experience of the center where the patient is treated.

The aim of this review was to compare POEM to LHM for the treatment of esophageal achalasia, taking results, early and long term complications and costs into consideration.

Technical Considerations Regarding LHM and POEM

Ernst Heller first described surgical esophageal myotomy in 1913 (3). At first it consisted of transabdominal extra-mucosal myotomy on both the anterior and posterior wall of the esophagus. In 1923 this approach was modified by Zaaijer using only a myotomy on the anterior wall of the esophagus, with excellent results (4). Open abdominal and thoracic approaches can be safely performed, however they involve long recovery times. Modern Heller myotomy is done using laparoscopy and thoracoscopy, with similar outcomes and better quality of life than open techniques. The myotomy is made by dividing both esophageal muscular layers, circular and longitudinal, above the area of constriction. The incision is extended 4-6 cm on the esophagus and 1.5-3 cm onto the gastric cardia, as to ensure complete relaxation of the cardia. In order to decrease the incidence of postoperative reflux, a fundoplication is performed, meaning that the gastric fundus is plicated around the lower end of the esophagus and stitched in place. LHM was considered the standard of care for the management of achalasia for a long time, having minimal perioperative morbidity and excellent long-term functional outcomes (5).

POEM was first performed by Haruhiro Inoue in Japan in 2008 (6). Since then, the procedure has been increasingly used all...
around the world and has become a developed endoscopic technique for the treatment of achalasia. It involves the creation of a sub-mucosal tunnel through a breach in the esophageal mucosa. This tunnel allows access to esophageal muscle fibers, which are cut through electrocautery, using an endoscopic triangle tipped knife designed specifically for this technique. Sectioning of the muscle fibers begins at 10-12 cm proximally to the cardia and extends 3-5 cm distally in the stomach, thus ensuring complete cutting of the inferior esophageal sphincter. From a technical point of view, POEM is a reversed partial LHM, one of the significant differences between the two being that in the case of POEM, the sectioning of the muscle fibers is concentrated on the inner circular layer, whereas in the case of LHM, because of the esophageal extrinsic approach, both muscular layers have to be sectioned. Some authors have tested full-thickness myotomy during POEM procedure, but with no improvement in efficacy. They however have concluded that full-thickness myotomy leads to a reduced procedural time (7,8), while others have shown an increase in gastroesophageal reflux (GER) rates as opposed to circular myotomy (9).

**Efficacy of LHM and POEM**

LHM has fully proved its worth since it has been introduced almost a century ago, several studies showing good results even in very long follow-ups, of 80 months or more (Table 1).

As it is a recent procedure, POEM could not be properly evaluated in the long term in multiple studies. However, from the data we have so far, POEM proves its effectiveness in the short and medium term, with studies showing satisfactory results in over 80% of patients treated with this method (Table 2). Some long term studies have been performed, in centers with a high case load of patients treated with achalasia. One study by Teitelbaum et al which followed-up patients for 5 years revealing that POEM resulted in durable longterm symptomatic relief for 83% of patients without the need for reintervention (10). A significant advantage of POEM over LHM is that in case of ineffective symptom control or relapse, a second session can be performed with good results, as Li et al showed on a cohort of 15 patients, the therapeutic effectiveness being 100% at one year after reintervention (11).

### Table 1. Success rate of and incidence of reflux after LHM

<table>
<thead>
<tr>
<th>Authors</th>
<th>Follow-up (months)</th>
<th>Success rate</th>
<th>Patients with reflux</th>
</tr>
</thead>
<tbody>
<tr>
<td>Peng et al, 2017 (12)</td>
<td>36</td>
<td>80%</td>
<td>6.7%</td>
</tr>
<tr>
<td>Leeds et al, 2017 (13)</td>
<td>6</td>
<td>66%</td>
<td>No information</td>
</tr>
<tr>
<td>Moonen et al, 2016 (14)</td>
<td>60</td>
<td>84%</td>
<td>No information</td>
</tr>
<tr>
<td>Salvador et al, 2016 (15)</td>
<td>49</td>
<td>88.9%</td>
<td>11.6%</td>
</tr>
<tr>
<td>Persson et al, 2015 (16)</td>
<td>80.4</td>
<td>92%</td>
<td>No information</td>
</tr>
<tr>
<td>Hamdy et al, 2015 (17)</td>
<td>48</td>
<td>88%</td>
<td>28%</td>
</tr>
<tr>
<td>Krishnamohan, 2014 (18)</td>
<td>77.5</td>
<td>49.8%</td>
<td>70.3%</td>
</tr>
</tbody>
</table>

### Table 2. Success rate of and incidence of reflux after POEM

<table>
<thead>
<tr>
<th>Authors</th>
<th>Follow-up (months)</th>
<th>Success rate</th>
<th>Patients with reflux</th>
</tr>
</thead>
<tbody>
<tr>
<td>Peng et al, 2017 (12)</td>
<td>36</td>
<td>83.3%</td>
<td>8.3%</td>
</tr>
<tr>
<td>Leeds et al, 2017 (13)</td>
<td>6</td>
<td>82%</td>
<td>No information</td>
</tr>
<tr>
<td>Teitelbaum et al, 2017 (10)</td>
<td>60</td>
<td>84%</td>
<td>No information</td>
</tr>
<tr>
<td>Hungness et al, 2016 (19)</td>
<td>28</td>
<td>92%</td>
<td>27%</td>
</tr>
<tr>
<td>Werner et al, 2016 (20)</td>
<td>29</td>
<td>77.5%</td>
<td>37%</td>
</tr>
<tr>
<td>Inoue et al, 2015 (21)</td>
<td>36</td>
<td>88.5%</td>
<td>21.3%</td>
</tr>
<tr>
<td>Chen et al, 2015 (22)</td>
<td>24</td>
<td>100%</td>
<td>6.7%</td>
</tr>
</tbody>
</table>
One of the biggest questions regarding POEM is how well it fares when compared to LHM, in short, medium and long-term follow-up. Talukdar et al have performed a systematic review and meta-analysis on 29 studies including 1045 patients, and concluded that POEM has similar outcomes as LHM, with no differences between POEM and LHM in reduction in Eckhart's score. However, the authors take note of a significant heterogeneity among the study, which could have influenced the results (23). A similar large meta-analysis by Marano et al compared POEM and LHM, revealing that POEM has a slightly better maintenance of a lower Eckardt score than LHM, but the difference didn't reach statistical significance. The authors also showed that there were no significant differences between the two techniques regarding duration of intervention, postoperative pain score and analgesic requirement (24). A recent systematic review and meta-analysis of 53 articles including both LHM and POEM performed on a total of 1958 patients revealed comparable results between the two techniques. As such, predicted probabilities for improvement in dysphagia at 12 months were 93.5% for POEM and 91.0% for LHM (P < 0.01), while at 24 months they were 92.7% for POEM and 90.0% for LHM (P < 0.01). However, across all included studies, dysphagia improved in 93.2% of patients who underwent POEM and 87.7% of patients who were treated with LHM, showing that at least in short-term, POEM is superior than LHM (25). Concerning medium to long term efficacy, a 3 year follow-up study by Peng et al indicated that POEM is an effective treatment that is comparable with LHM (12).

Ujiki et al compared POEM to LHM concluding that procedural time, the length of myotomy and complication rates were similar. After treatment, pain evaluated using a visual analogue scale was significantly worse in the LHM group as opposed to the POEM one (LHM 5.7 ± 0.4 vs POEM 3.9 ± 0.6, p=0.02). As such, pain relief medication use was lower after endoscopic therapy (POEM 26.0 ± 13.7 mg vs LHM 90.0 ± 48.5 mg morphine, p=0.02) (26). However, Hungness et al reported different results, as they noted that pain scores were similar upon arrival in the post-anesthesia care unit and on the first postoperative day, but were higher 2 hours after the procedure for POEM patients (3.5 vs 2, p=0.03) (27).

Although some earlier studies report a similar procedural time between POEM and LHM, others report a shorter time in favor of POEM. This phenomenon is plausible in terms of the initial lack of operator experience. As experience accumulates, the difference in time between the two procedures should become significant.

Complications of LHM and POEM

The rate of complications related to LHM is low, with most of the studies showing a rate between 0-9% (28). Of these, perhaps the most feared complication is esophageal perforation, as reintervention is needed in order to suture the breach.

The main adverse effects that may be encountered in POEM are bleeding, mucosal perforations, infections or events related to endoscopic insufflation, such as pneumoperitoneum or pneumothorax. In order to avoid these events, POEM must be performed by experienced endoscopists, with extensive training in esophageal disorders. Most studies have shown that approximately 20 cases are sufficient to accomplish training for POEM as an experienced endoscopist, after which the length of the procedure and accidental mucosal damage plateaus (29–31).

Mucosal perforation, seen in up to 10% of cases (32), can be a real problem especially if it is not recognized on time, as the mucosal layer maintains the integrity of the digestive tract after performing POEM. When mucosal lesions are identified, endoscopic management is fairly simple, using hemostatic clips to close them. If the perforations are larger, endoscopic suturing or the placement of fully covered metal stents can be used.

Events related to insufflation are probably the most common, being observed during POEM as well as post procedurally in about 20% of cases (23). However, they are not life
Peroral Endoscopic Myotomy Versus Heller Myotomy for Achalasia: Pros and Cons

threatening as long as the necessary precautions are taken. In this respect, it is imperative to use CO2 insufflation during endoscopy because of its much faster absorption rate as opposed to air. At the same time, low or very low flow rates are recommended, so that gas accumulation is minimal. If pneumoperitoneum occurs during the procedure, it can be resolved quickly by transombilical drainage with a Veress needle.

Gastroesophageal Reflux in LHM and POEM

As during the Heller myotomy, the entire esophageal sphincter that secures the anti-reflux barrier is cut, it is mandatory to complete the intervention with an anti-reflux procedure, such as a Toupet or Dor fundoplication, in order to prevent post-operative gastro-esophageal reflux. This has been thoroughly validated across time. A prospective randomized trial performed by Richards et al evidenced a pathological GER rate of 47.6% when no antireflux procedure was added to LHM, compared to a 9.1% rate in the group with Heller myotomy followed by anterior fundoplication. The authors also showed that there was no significant statistical differences between the two groups regarding postoperative gastro-esophageal reflux. Similar GER rates of up to 10% have been shown by other authors when adding an antireflux procedure to LHM (34–37).

As opposed to LHM, during POEM there is no dissection of the phrenoesophageal membrane, which may maintain anatomical reflux barriers. What’s more, during POEM only the circular muscular layer is sectioned. Surprisingly however, results from multiple studies that have compared POEM to LHM show similar GER rates between the two techniques (12,38–40), with an incidence ranging between 0% and 37% (24).

However, the method by which reflux is evaluated is very important. Thus, if symptomatic and endoscopic scores provide similar results between the two techniques, gold-standard evaluation by pH monitoring shows the presence of much higher reflux rates in patients treated with POEM. As such, Schlottmann et al showed a pathological reflux rate of 11.1% in LHM patients and in 47.5% of POEM patients (25). The reason for this is still discussed, but the lack of an anti-reflux procedure in POEM might be the main reason. Even if in all cases reflux symptoms can be completely controlled by proton pump inhibitors therapy (21), emerging side effects of long term therapy with these drugs have to be taken into account.

Cost of LHM vs POEM

Intuitively we could say that POEM should be cheaper than LHM. It is a less invasive procedure, the approach being endoscopic as opposed to surgical, so the duration of the intervention is reduced and requires less personnel. However, the few studies that have analyzed the problem have come to a different conclusion, namely that there is no significant cost difference between the two procedures (41).

Khashab et al have compared POEM to robotic Heller myotomy (RHM) and found POEM to be cheaper and shorter in duration, but with a similar efficacy and safety. However, Shaligram et al have compared LHM to RHM beforehand and found higher costs with the latter procedure (42). As such, comparing POEM to LHM might yield more similar costs. Khashab et al noted that there was a similar length of stay between the patients who underwent POEM and those who underwent RHM. The difference in costs between the two groups might be attributable to the fact that those treated with RHM needed an increased post procedural support (41).

Miller et al concluded that the cost effectiveness of POEM and LHM is equivalent (43). Greenleaf et al reported similar costs in POEM and LHM, attributable to a greater mean cost of radiological assessments after POEM and to a longer mean stay in the post-anesthesia care unit, which typically lasted 24 hours, as opposed to patients who underwent LHM and were transferred on general surgical wards much sooner. However, the authors note that with further experience in POEM the costs will surely drop, as procedural times, number of
endoclips used, radiological assessments and length of stay in post-anesthesia care units will be greatly reduced (44). There might come a time when POEM will be performed as a one-day endoscopic surgery procedure.

Conclusion

POEM seems to be here to stay, as multiple studies showed that its effectiveness is similar to that of LHM, while others showed it may be better. POEM is less invasive and may result in shorter hospitalization, and quicker return to normal activity while still providing durable long-term symptomatic relief, with a reduced cost in hospitalization. We believe it is time to consider POEM as gold standard for the treatment of achalasia.

Authors’ Contributions

CT and RA have drafted the article. MT has revised the article.

Conflict of Interests

The authors declare no conflict of interests for this article.

Funding

The authors declare that no specific funding has been received for this study.

References

26. Ujiki MB, Yetasook AK, Zapf M, Linn JG, Carbray JM, Denham W. Peroral endoscopic myotomy: A short-term comparison with the stan-
Peroral Endoscopic Myotomy Versus Heller Myotomy for Achalasia: Pros and Cons


