

Esophageal Manometry Among Patients With Dysphagia Referred To Kurdistan Center For Gastroenterology And Hepatology

Mohammed O. Mohammed*
Bakhtyar F. Salim**
Ali A. Ramadhan ***

Abstract

Background and Objectives: Dysphagia is a common problem in patients with primary motor disorders of the esophagus. Esophageal manometry is the gold standard for diagnosis of these disorders. Introduction of high resolution manometry represented a significant improvement in data recording and diagnostic yield. The objective of the study was to assess the findings of esophageal high resolution impedance manometry in patients presenting with dysphagia in Sulaimani governorate.

Patients and Methods: This study extended from September, 2012 to December, 2013 and included 120 patients with dysphagia who were referred for manometry in Kurdistan Center for Gastroenterology and Hepatology (KCGH) in Sulaimani city. All patients underwent upper endoscopy to exclude mechanical and inflammatory causes of dysphagia then the high resolution impedance manometry was used with liquid and viscous swallows.

Results: The mean age of the study population was 43 years. The female to male ratio was 1.5:1. The mean duration of dysphagia was 2 years. The most common esophageal motility abnormality was achalasia (N=44, 36.7%) followed by hypertensive LES (N=31, 25.8%), ineffective esophageal motility (N=9, 7.5%), hypotensive LES (N=5, 4.2%) and diffuse esophageal spasm (N=3, 2.5%). The high resolution impedance manometry was normal in 28 patients (23.3%). Of the 44 patients with achalasia, 15 patients (34%) had vigorous achalasia. Using Chicago classification, the most common type of achalasia was type II (N=26, 59%) followed by type I (N=13, 29.6%) and then type III (N=5, 11.4%).

Conclusions: Esophageal high resolution impedance manometry has an acceptable diagnostic yield in patients with dysphagia. The most common finding is achalasia. Further studies are recommended.

Key words: dysphagia, manometry, Achalasia, KCGH.

*Department of Medicine, School of Medicine, Faculty of Medical Sciences, University of Sulaimani; Kurdistan Center for Gastroenterology and Hepatology, Sulaimani; Iraq.

** Department of Medicine, School of Medicine, Faculty of Medical Sciences, University of Duhok, Duhok; Kurdistan Center for Gastroenterology and Hepatology, Sulaimani; Iraq.

Introduction

Dysphagia is an “alarming” symptom that needs clinical evaluation to define the cause and initiate therapy (1). Its causes are broadly classified into mechanical lesions, inflammatory etiologies, motility disorders or functional dysphagia (2). Esophageal motility disorders can result in major morbidity and, in some cases, in an increased risk of cancer and death (3). Esophageal manometry is the gold standard for diagnosis of these disorders (4). Introduction of high resolution manometry (HRM), represented a significant improvement in data recording and diagnostic yield (5,6). Recently, high resolution impedance manometry (HRIM) has combined the benefits of HRM and impedance-based bolus transit assessment. Abnormal impedance may be a sensitive indicator of esophageal functional abnormality (7). HRIM can detect segmental abnormalities (8) and reduce the problems of asymmetry and artifact (9) in addition to being simple to use and easy to

Patients and methods

This study extended from September, 2012 to December, 2013. During this period, 120 cases were collected. It included patients with dysphagia referred for esophageal manometry at Kurdistan Center for Gastroenterology and Hepatology (KCGH). The informed consent was obtained and all patients underwent upper endoscopy before manometry. The InSIGHT high resolution impedance manometry system (Sandhill Scientific, Inc, Highlands Ranch, Colo, USA) was used. Patients were instructed to fast for a minimum of four hours for solids and two hours for liquids and to stop medications known to affect esophageal motor function for 24 hours prior to the test (eg. beta-blockers, nitrates, calcium channel blockers, anticholinergic drugs, prokinetics, nicotine, caffeine and opiates). The equipment was checked and calibrated before each study.

learn (10). However, the use of HRIM may be limited because the equipment is expensive. Moreover the clinical significance of HRIM-detected esophageal dysmotility remains uncertain in some cases (11). Achalasia is a primary esophageal motor disorder of unknown etiology characterized by insufficient relaxation of lower esophageal sphincter and loss of esophageal peristalsis (12). Vigorous achalasia represents the early stage of achalasia which may be due to loss of inhibitory neurons, but cholinergic stimulation continues unopposed (4). Esophageal pressure topography has allowed for the differentiation of achalasia into three subtypes using Chicago classification. These subtypes may have potential treatment outcome implications with subtype II having the best prognosis, whereas subtype I has somewhat worse outcome and subtype III can be difficult to treat (12).

The catheter (Comfortec®, 6.4 Fr, 60 cm, figure 1) was placed transnasally. Patients were placed in a 30 degrees semi-recumbent position and allowed to accommodate to the catheter and then the resting lower esophageal sphincter (LES) pressure was measured. A sequence of 10 wet swallows (5 ml of room-temperature water per swallow) and then 10 viscous swallows (3-ml of gel per swallow) were used to examine the LES relaxation and esophageal body peristaltic activity. At least 20-30 seconds were allowed between swallows. The esophageal motility abnormalities were classified as shown in table 1 (13, 14). Achalasia was sub-classified into classic (average esophageal body amplitude \leq 40 mmHg) and vigorous ($>$ 40 mmHg) subtypes (4). The esophageal pressure topography was revised to determine the three subtypes of

achalasia using Chicago classification of distal esophageal motility disorders; type-I (achalasia with aperistalsis or classic achalasia), type-II (achalasia with pan-esophageal pressurization) and type-III (achalasia with spasms or spastic achalasia) (2,12). Analysis of data was

conducted to calculate the P-value using 2. Testing for differences among the means of groups was done using analysis of variance (ANOVA). For the associations or differences to be significant, the P-value should have been less than 0.05.

Results

The mean age of the study population was 43 years (95% Confidence Interval (95% CI) was 40.1 – 45.7 years) with a range of 10 – 84 years. The majority (71%, N=85) were younger than 50 years. The female to male ratio was 1.5:1. The mean duration of dysphagia was 2 years (95% CI = 17.6 – 29 months) with a range of 2 weeks to 15 years. Of the study population, 75% (N=90) had dysphagia for solid more than liquid. Most patients had no obvious relieving or aggravating factors. The most common clinical features associated with dysphagia were chest pain (N=50, 41.7%), regurgitation (N=36, 30%), respiratory symptoms such as chronic cough and recurrent respiratory tract infections (N=33, 27.5%), heartburn (N=30, 25%) and weight loss (N=26, 21.7%). All the patients underwent upper endoscopy before manometry and 29 patients (24.2%) had endoscopic features suggestive of achalasia. Barium swallow study was performed in 50 patients (41.7%) and there were radiological features suggestive of achalasia in 14 patients (28%). Compared with esophageal manometry, upper endoscopy had a sensitivity of 52%, specificity of 92%, positive predictive value (PPV) of 79%, negative predictive value (NPV) of 77% and “overall

accuracy” of 77.5% for detecting achalasia while barium swallow had a sensitivity of 60%, specificity of 93%, PPV of 86%, NPV of 78% and “overall accuracy” of 80%. The most common esophageal motility abnormality was achalasia (N=44, 36.7%) followed by hypertensive LES (N=31, 25.8%), ineffective esophageal motility (N=9, 7.5%), hypotensive LES (N=5, 4.2%) and diffuse esophageal spasm (N=3, 2.5%). The HRIM was normal in 28 patients (23.3%). Figure 2 clarifies the esophageal motility findings in this study. Of the 44 patients with achalasia, 15 patients (34%) had vigorous achalasia. Using Chicago classification, 13 (29.6%) patients had achalasia type I, 26 (59%) had achalasia type II and 5 (11.4%) had achalasia type III. Statistical analysis was conducted for the major esophageal manometric findings and is shown in table 2. There was significant statistical difference in the age of presentation (P = 0.028) with achalasia patients being older than other groups and those with normal manometry being the youngest. There was no significant statistical difference with regard to sex of patients (P = 0.1). Although the duration of dysphagia was not statistically different among the groups (P = 0.26), achalasia.

Figures and tables:

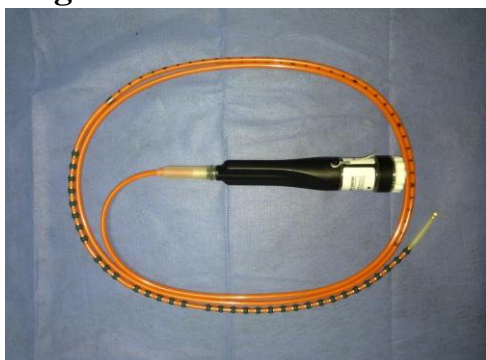


Figure (1): High resolution impedance manometry catheter.

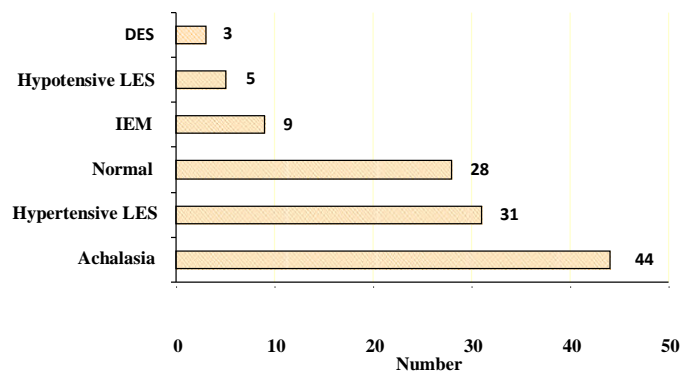
and IEM patients had a mean duration of more than 2 years. Dysphagia for solid more than liquid was significantly different among the groups ($P < 0.000$) being more likely in patients with achalasia (88.6%) than in other groups. The course of dysphagia; its location; relieving or aggravating

factors; association with chest pain, weight loss or heartburn; being smoker or alcohol drinker and mean body mass index were not statistically significantly different among the analyzed groups ($P > 0.05$).

Discussion

Esophageal HRIM is an important development in esophageal function testing (6,15). The lack of local data was the impetus for the current study. To our knowledge, this study is the first in Iraq that uses HRIM to study the esophageal motility abnormalities in dysphagia. We included patients presenting with dysphagia because it is the primary and the most cost effective indication for esophageal motility disorder (16). The mean duration of dysphagia in this study was 2 years which is consistent with that reported in literature (2, 17) but we had cases of

dysphagia with duration ranging from 2 weeks to 15 years in whom the manometry was useful. Dysphagia for solid and liquid is the hallmark of esophageal motility disorders (2) which was also observed in our study. Early in the course of disease, dysphagia may be intermittent but then becomes persistent or progressive with the advance of disease (2). In almost half of our cases (45.8%), the course was slowly progressive (over 2 years) which could imply a more advanced stage of disease in this study.



Figure(2): Esophageal motility findings (N = 120)

DES, Diffuse Esophageal Spasm; LES, Lower Esophageal Sphincter; IEM, Ineffective Esophageal Motility. The most common esophageal motility abnormality in this study was achalasia (36.7%) followed by hypertensive LES (25.8%) while HRIM was normal in 23.3% of patients. The manometric findings in dysphagia patients have been variable. A study done by Dumitraşcu et al (18) reported achalasia as the most common finding followed by diffuse esophageal spasm (DES). In another study done in India by Misra et al (4), the most common finding was also achalasia followed by normal manometry and DES. While in a study done by Dekel et al (19), the most common finding in dysphagia was normal manometry followed by IEM and then achalasia. This variability in the findings might be attributed to differences in the type of manometry used, selection of cases and referral patterns although a real geographical difference cannot be excluded. In this study, the clinical manifestations associated with achalasia were dysphagia for solid more than liquid (88.6%) and association of dysphagia with regurgitation (54%) and

respiratory symptoms (48%). Chest pain, weight loss and heartburn were present in 34%, 32% and 18% respectively. Dysphagia for both solid and liquid has been reported in up to 90% of achalasia cases (20). Other symptoms including chest pain, heartburn, regurgitation, and weight loss were reported in up to 60% of patients (20). Both upper endoscopy and barium swallow study had low sensitivities for detecting features of achalasia (52% and 60% respectively) which will result in missing of achalasia cases if we use them as the “only” diagnostic studies. Endoscopy has a poor sensitivity and specificity in the diagnosis of achalasia and its primary role in the workup of achalasia is focused on ruling out a mechanical obstruction or pseudoachalasia (17). Videofluoroscopy has an overall sensitivity of 80–89% and specificity of 79–91% for diagnosing esophageal motility disorders (eg. achalasia) and is useful for identifying pre-clinical disease relapse (17, 21). Vigorous achalasia was identified in 34% of achalasia patients in the current study. The reported frequencies vary from 1.5% to 31% (18). In our study, the most common achalasia type was type II

(N=26, 59%) followed by type I (N=13, 29.6%) and then type III (N=5, 11.4%). This is comparable to the results of the two largest studies in this regard which were done by Salvador et al (22) (N= 246 patients with rates of 51.6%, 39%, and 9.4% for type II, I, and III respectively) and Rohof et al (23) (N= 176 patients with rates of 64.7%, 25.1%, and 10.2% for type II, I, and III by the acid as a protective mechanism (24). Unrecognized GERD may partly explain this high rate in our study as 35% of these patients had dysphagia associated with heartburn. Ineffective esophageal motility and hypotensive LES were observed in 7.5% and 4.2% respectively. The reported rates of these two findings range from 0 to 27% (4, 15, 19). The clinical significance, association with symptoms and appropriate management of these manometric findings have not been established (25). We had only three cases of DES (2.5%). This is slightly lower than that reported by other studies which ranged 4-9% (4, 15, 19). An important issue is that DES may be associated with incomplete LES relaxation, in which case it is likely a variant of achalasia. Studies have shown that DES may progress over time to

respectively). Hypertensive LES was the second most common manometric finding in our study (25.8%). This is much higher than that reported by Tutuian et al (1%) (15), Misra et al (3.6%) (4) and Dekel et al (7%) (19). There is a paradoxical association between hypertensive LES and GERD which has been attributed to stimulation of LES contraction classic achalasia (26). This suggests that if DES is a distinct motor disorder, it is certainly rare and traditional manometric criteria have oversimplified it, resulting in over-diagnosis of the entity (27). In conclusion, esophageal high resolution impedance

manometry has an acceptable diagnostic yield in patients with dysphagia. The most common finding was achalasia of type II using Chicago classification. Further larger scale randomized studies are advised to study the factors associated with esophageal motility disorders. Acknowledgements: Our thanks go to Ms. Ashna Khafwr for her kind help in performing the manometry studies. Many thanks should be given to the patients involved in the study without their help it was not possible to conduct this study.

References:

1. Javle M, Ailawadhi S, Yang GY, Nwogu CE, Schiff MD, Nava HR. Palliation of Malignant Dysphagia in Esophageal Cancer: A Literature-Based Review. *J Support Oncol* 2006; 4: 365–373.
2. Kahrilas PJ, Pandolfino JE. Esophageal Neuromuscular Function and Motility Disorders. In: Feldman M, Friedman LS, Brandt LJ Editors. *Sleisenger and Fordtran's gastrointestinal and liver disease: pathophysiology, diagnosis, management*. 9th Ed. Philadelphia: Elsevier. 2009, PP 677-704.
3. Massey BT. Esophageal Motor and Sensory Disorders: Presentation, Evaluation, and Treatment. *Gastroenterol Clin N Am* 2007; 36: 553–575.
4. Misra A, Chourasia D, Ghoshal UC. Manometric and symptomatic spectrum of motor dysphagia in a tertiary referral center in northern India. *Indian J Gastroenterol* 2010; 29(1): 18-22.
5. Kahrilas PJ, Sifrim D. High-resolution manometry and impedance-pH/manometry:

- valuable tools in clinical and investigational esophagology. *Gastroenterology* 2008; 135: 756-769.
6. Fox MR, Bredenoord AJ. Oesophageal high-resolution manometry: moving from research into clinical practice. *Gut* 2008; 57(3): 405-423.
 7. Koya DL, Agrawal A, Freeman JE, Castell DO. Impedance detected abnormal bolus transit in patients with normal esophageal manometry. Sensitive indicator of esophageal functional abnormality? *Dis Esophagus* 2008; 21: 563-569.
 8. Fox M, Hebbard G, Janiak P, Brasseur JG, Ghosh S, Thumshirn M, et al. High-resolution manometry predicts the success of oesophageal bolus transport and identifies clinically important abnormalities not detected by conventional manometry. *Neurogastroenterol Motil* 2004; 16: 533-542.
 9. Ghosh SK, Pandolfino JE, Zhang Q, Jarosz A, Shah N, Kahrilas PJ. Quantifying esophageal peristalsis with high-resolution manometry: a study of 75 asymptomatic volunteers. *Am J Physiol Gastrointest Liver Physiol* 2006; 290: 988-997.
 10. Clouse RE, Staiano A, Alrakawi A, Haroian, L. Application of topographical methods to clinical esophageal manometry. *Am J Gastroenterol* 2000; 95(10): 2720-2730.
 11. Roman S, Pandolfino J, Mion F. High-resolution manometry: a new gold standard to diagnose esophageal dysmotility? *Gastroenterol Clin Biol* 2009; 33(12): 1061-1067.
 12. Vaezi MF, Pandolfino JE, Vela MF. ACG Clinical Guideline: Diagnosis and Management of Achalasia. *Am J Gastroenterol* 2013; 108 (8): 1238-1249.
 13. Wang A, Pleskow DK, Banerjee S, Barth BA, Bhat YM, Desilets DJ, et al. Esophageal function testing. *Gastrointest Endosc* 2012; 76 (2): 231-243.
 14. Spechler SJ, Castell DO. Classification of oesophageal motility abnormalities. *Gut* 2001; 49: 145 - 151.
 15. Tutuian R, Castell DO. Combined multichannel intraluminal impedance and manometry clarifies esophageal function abnormalities: study in 350 patients. *Am J Gastroenterol* 2004; 99: 1011-1019.
 16. Johnston PW, Johnston BT, Collins BJ, Collins JS,
 17. Love AH. Audit of the role of oesophageal manometry in clinical practice. *Gut* 1993; 34: 1158-1161.
 18. Pohl D, Tutuian R. Achalasia: an overview of diagnosis and treatment. *J Gastrointest Liver Dis* 2007; 16(3): 297-303.
 19. Dumitraşcu DL, Blaga TS, David L. Esophageal Achalasia - Manometric Patterns. *Rom J Intern Med* 2009; 47(3): 243-247.
 20. Dekel R., Pearson T, Wendel C, De Garmo P, Fenerty MB, Fass R. Assessment of oesophageal motor function in patients with dysphagia or chest pain — the Clinical Outcomes Research Initiative experience. *Aliment Pharmacol Ther* 2003; 18: 1083-1089.
 21. Francis DL, Katzka DA. Achalasia: update on the disease and its treatment. *Gastroenterology* 2010; 139: 369-374.
 22. Carucci LR, Lalani T, Rosen MP, Cash BD, Katz DS, Kim DH, et al. ACR Appropriateness Criteria: dysphagia. Available from: www.acr.org/~media/ACR/Documents/AppCriteria/.../Dysphagia.pdf [Accessed March, 2014].
 23. Salvador R, Costantini M, Zaninotto G, Morbin T, Rizzetto C, Zanatta L, et al. The preoperative manometric pattern predicts the outcome of surgical treatment for esophageal achalasia. *J Gastrointest Surg* 2010; 14: 1635-1645.
 24. Rohof WO, Salvador R, Annese V, Bruley des Vannes S, Chaussade S, Costantini M, et al. Outcomes of treatment for achalasia depend on manometric subtype. *Gastroenterology* 2013; 144: 718-725.
 25. Gad El-Hak NA, Mostafa M, AbdelHamid H, Hal-eem M. Hypertensive

Lower Esophageal Sphincter (HLES): Prevalence, Symptoms Genesis and Effect of Pneumatic Balloon Dilatation. Saudi J Gastroenterol 2006; 12(2): 77-82.

26. Bodger K, Trudgill N. BSG Guidelines in Gastroenterology: Guidelines for oesophageal manometry and pH monitoring. 2006 Available from: www.bsg.org.uk/pdf_word_docs/oesp_man.pdf. [accessed July, 2012].

27. Prabhakar A, Levine MS, Rubesin S,

Laufer I, Katzka D. Relationship Between Diffuse Esophageal Spasm and Lower Esophageal Sphincter Dysfunction on Barium Studies and Manometry in 14 Patients. Am J Roentgenol 2004; 183(2): 409-413.

28. Pandolfino JE, Kahrilas PJ. AGA Technical Review on the Clinical Use of Esophageal Manometry. Gastroenterology 2005; 128: 209-224.

Table (1): Criteria for diagnosing esophageal motility abnormalities.

Functional defect	Diagnosis	Manometric findings
Aperistalsis	Achalasia	Absent distal peristalsis, incomplete lower esophageal sphincter(LES) relaxation, increased LES pressure (>45 mm Hg)
Uncoordinated motility	Diffuse esophageal spasm (DES)	>=20% simultaneous contractions, repetitive contractions (>3 peaks), prolonged duration of contractions, incomplete LES relaxation
Hypercontractile	Nutcracker esophagus	Increased amplitude (>180 mm Hg), increased peristaltic duration
	Hypertensive LES	Resting LES pressure >45 mm Hg, incomplete LES relaxation
Hypocontractile	Ineffective esophageal motility(IEM)	> 30% non-transmitted peristalsis, peristaltic amplitude < 30 mm Hg
	Hypotensive LES	Resting LES pressure <10 mm Hg

Table (2): Statistical analysis of the major esophageal manometric findings.

Parameters	Achalasia (N=44)	Hypertensive LES (N=31)	Normal (N=28)	IE M (N=9)	P-value
Age (mean ± SD)	47.7 ± 17.3	41.6 ± 17.9	37 ± 8.5	38.2 ± 11.6	0.028
Male sex	17	7	15	4	0.10
Duration in months (mean ± SD)	28.8 ± 23.5	14.9 ± 24.1	21.7 ± 36.7	29.4 ± 58.5	0.26
Dysphagia for solid more than liquid	39	26	21	1	0.000
Progressive course	19	20	10	4	0.14
Retrosternal location	32	24	19	9	0.27
Relieving with liquid	16	8	10	1	0.40
Aggravation by stress	8	2	1	1	0.20
Association with chest pain	15	18	12	2	0.12
Association with regurgitation	24	6	4	2	0.000
Association with respiratory symptoms	21	10	1	1	0.000
Association with heartburn	8	11	5	4	0.14
Association with weight loss	14	9	2	1	0.06
Current smoking	2	4	4	3	0.08
Alcohol drinking	1	1	3	1	0.35
Body mass index (mean ± SD)	24.5 ± 4.8	23.4 ± 3.4	26.1 ± 6	26.7 ± 4.7	0.10
Resting LES pressure (mean ± SD)	48.3 ± 24.8	41.7 ± 21.7	23.5 ± 8.3	8.5 ± 11.7	0.000
LES relaxation pressure (mean ± SD)	26.1 ± 15	16.2 ± 5.4	4.8 ± 2.6	1.1 ± 12.1	0.000
Distal esophageal amplitude (mean ±SD)	24.2 + 71.1	82.4 ± 27.6	77.3 ± 25.7	35 ± 13	0.000