Clinical application of pharyngeal high-resolution manometry in Ear, Nose and Throat (ENT) practice

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Abstract: Pharyngeal high-resolution manometry with impedance (P-HRM-I) is an emerging, ambulatory swallow assessment for patients with pharyngeal dysphagia. P-HRM-I provides quantifiable biomechanical metrics of the pharynx and upper oesophageal sphincter (UOS) and may inform clinical management. This paper describes the fundamentals of P-HRM-I assessment and analysis, with application to common dysphagia conditions presenting to an Otolaryngologist. The procedure for conducting a P-HRM-I assessment is detailed. The acquisition of contractility (pressure), timing, and bolus transit (impedance) is described. P-HRM-I analysis and interpretation is presented using metric definitions agreed by international expert consensus. The application of P-HRM-I is demonstrated in five cases presenting with globus sensation, cricopharyngeal bar, and dysphagia post-head and neck cancer treatment, with reference to a healthy swallow. P-HRM-I analysis and interpretation informing subsequent clinical management options is presented. The globus case demonstrates exclusion of UOS dysfunction or hypertonicity contributing to globus sensation. The cricopharyngeal bar cases demonstrate distinct UOS metrics differentiating those indicated for surgical management. Post-head and neck cancer treatment cases demonstrate the underlying biomechanical features contributing to dysphagia, whether localised at the pharynx or UOS to inform targeted treatment. This review showcases the application of P-HRM-I to extend current imaging swallowing assessments to guide the management of common Otolaryngology dysphagia presentations. Further clinical translation of this technology is recommended to optimise dysphagia management.

Keywords: Otolaryngology; deglutition disorders; manometry; globus sensation; head and neck neoplasms

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Introduction

The diagnosis and management of oropharyngeal swallowing difficulty (dysphagia) is challenging (1). This is because dysphagia is not a distinct disease but rather a symptom associated with a wide range of underlying pathologies across the lifespan presenting with differing severity and permanence (1). Dysphagia has a reported prevalence of 16–20% in the general population but increases to as high as 50% in some specific groups (2), and is associated with malnutrition, dehydration and aspiration pneumonia (3,4).

A patient with dysphagia may present to an Otolaryngologist for assessment and treatment. Following exclusion of malignancy or anatomical abnormalities, swallowing studies are conducted to visualise bolus transit through the oropharynx to account for the reported dysphagia. A videofluoroscopic swallowing study (VFSS) or a flexible endoscopic evaluation of swallowing (FEES) are widely used instrumental assessments, but they provide limited insight into the biomechanical breakdown (5). Despite advancements in quantitative reporting tools (6,7) for visual instrumental assessments, there remains no globally accepted measures, leaving clinical interpretation of the swallow beyond identifying penetration and aspiration, predominantly dependent on clinician experience (8,9).

Pharyngeal high-resolution manometry with impedance (P-HRM-I) provides precise and quantitative measures of pharyngeal and upper oesophageal sphincter (UOS) pressures integrated with bolus transit (10). It can identify and localise alterations in the swallowing mechanism and determine the underlying pathophysiological breakdown leading to dysphagia. P-HRM-I metrics have been demonstrated to correlate with and be a predictor of aspiration on concurrent videofluoroscopic studies (10-13). This emerging technology operates as part of routine clinical care in only a few centers internationally (8,9,14) and requires continued clinical uptake to support translational outcomes. This paper aims to showcase the interpretation of P-HRM-I through the presentation of selected cases relevant to the Ear, Nose and Throat (ENT) practice. Specifically, new insights offered from the P-HRM-I assessment for pharyngeal dysphagia patient management beyond VFSS are discussed.

Methods

P-HRM-I

P-HRM-I has demonstrated the capability to identify altered

biomechanical features contributing to dysphagia in patient cohorts (15,16) and, more recently, as an interventional outcome measure (17-20). The increased recognition of highresolution manometry (HRM) technology for pharyngeal dysphagia is foreseeable considering HRM is considered the gold standard for the diagnosis of oesophageal motility disorders in the clinical and research settings (21).

P-HRM-I is trans-nasal catheter assessment of pharyngeal swallowing. P-HRM-I involves data acquisition from closely spaced pressure sensors to measure the contractile activity representative of pressure generation spanning the pharynx to the UOS (22). P-HRM-I simultaneously measures the pharyngeal and UOS contractile muscle activity (pressure generation) with intra-luminal impedance that is representative of bolus movement across time (10,23). Together with analysis algorithms, these pharyngeal swallowing measurements allow for a sophisticated and quantifiable biomechanical assessment that ultimately increases the understanding of swallowing physiology and pathophysiology (24,25).

Case examples have been extracted from studies conducted at two centres: Flinders Medical Centre (Adelaide, Australia) and St. George Hospital (Sydney, Australia) (approved by Southern Adelaide Clinical Human Research Ethics Committee and St. Vincent's Hospital Human Research Ethics Committee). Informed consents are not required due to the retrospective nature of this study.

Equipment and procedure

A P-HRM-I swallow assessment can be conducted at the bedside or in the outpatient clinic setting with the patient sitting upright. P-HRM-I has been demonstrated to have high patient tolerability, with similarly low rates of complications (gagging 14%, vomiting 2% and epistaxis <1%) (22) as those reported for other trans-nasal procedures, such as nasoendoscopy. In our experience, complication rates are far lower and only occur during placement of the catheter and will resolve once the catheter is appropriately placed.

At our centre, a P-HRM-I assessment is standardised, consistent with the recommendations from the High-Resolution Pharyngeal Manometry International Working Group (24). An 8-French catheter with 32 pressure sensors (unidirectional) and 16 impedance transducers (Unisensor AG catheter, Atticon, Switzerland) is used for recording pressure and impedance data. Pressure and impedance data are acquired at 20 samples per second using the Solar GI acquisition unit (Medical Measurement System, Enschede, The Netherlands). Following a four-hour fasting period, lignocaine (5%; 2–3 sprays, equivalent to 0.3 mL) is topically administered to the nasal passage to aid catheter intubation and maximize patient comfort (24). Prior to trans-nasal catheter insertion, the catheter is placed in 37 °C water bath to reduce measurement error due to thermal drift. To assist placement of the P-HRM-I catheter, the patient is asked to place their head in a chin-down position once the catheter is advanced to velum (approximately 15 cm) to advance along the pharynx. At this time, the patient is asked to swallow sips of water as the catheter transits through the UOS into the proximal oesophagus (10,24). A minimum 5-minute accommodation period allows for subsidence of the anaesthetic effects from the lidocaine administration and participant accommodation to the catheter.

The swallowing assessment involves the administration of a standard bolus medium (SBMkitTM, Trisco Foods Pty Ltd., Brisbane, Australia) of an apple-flavored saline solution (0.65% sodium chloride NaCl) providing stable conductivity across bolus viscosity levels. Thin liquids and extremely thick liquids are prepared using Precise Thick-N INSTANT from the SBMkitTM and consistent with the International Dysphagia Diet Standardization Initiative (IDDSI) definitions (26). The standardised protocol includes testing a total of 18 boluses of 5-, 10- and 20-mL volumes of thin (IDDSI 0) and extremely thick (IDDSI 4) liquids. Three repeat swallows of each liquid volume and viscosity are recommended for acquisition of valid data (24). The duration of the procedure is approximately 15–20 minutes and can be performed in an ambulatory clinic setting.

Analysis

Currently, there are two international groups, one in Madison, USA (27,28) and the other in Adelaide, Australia (10) that use automatic methods of data extraction and analysis software platforms via MATLAB[®] (MathWorks, USA). Both groups have reported validation studies (29,30) and demonstrated good inter- and intra-rater reliability (31,32).

At our centre, we use the Swallow GatewayTM platform (http://www.swallowgateway.com/). Data are displayed as a spatiotemporal pressure-topography plot from the velopharynx to the proximal oesophagus (*Figure 1*). For each swallow, six spatiotemporal boundaries are defined for automated analysis. The anatomical markers include the proximal position of the velo-, meso- and hypo-pharynx and the proximal and distal margins of the UOS. Timing markers include the onset of UOS relaxation and contraction (*Figure 1*).

During pharyngeal swallowing, adequate pharyngeal

and UOS contraction requires both longitudinal and lumen occlusive horizontal pressures (33), which can be measured using P-HRM-I (24). The lumen occlusive pressures for anatomical regions (velo-, meso-, hypo-pharynx and UOS) are measured from multiple pressure sensors (24). These pressure values are calculated as a contractile integral, which describes the 'vigor' of a contraction within a particular space-time box on the pressure topography plot (*Figure 1*). This is the mean pressure of the anatomical region multiplied by the duration (s) and length (cm) of the specified anatomical region (24). Multiple pressure sensors are also necessary for the detection of accurate UOS pressure measurements, given that the UOS is known to move superiorly between 2–2.8 cm during swallowing and that the UOS is a high-pressure zone (34).

Intraluminal electrical impedance allows for the differentiation of complete or partial bolus transit, stasis and anterograde or retrograde bolus movement (35). Intraluminal electrical impedance measures the changes in resistance (ohms), which is inversely proportional to the electrical conductivity of the luminal contents and the cross-sectional area (36). Whilst air has a high resistance to current flow (high impedance), liquids have lower resistance (low impedance). Bolus presence is measured by the lowest impedance value at the hypopharyngeal and UOS regions (10). The maximum admittance (inverse product of nadir impedance) has been reported to represent UOS opening extent during swallowing (10). Bolus presence time (BPT) is an impedance derived measure (*Table 1*).

The Swallow Risk Index (SRI) is indicative of overall oropharyngeal swallow function. The SRI is a validated measure, which is determined from a mathematical formula derived from the integration of pressure and impedance measures. An SRI >15 is able to identify disordered oropharyngeal swallowing and associated aspiration risk correlated with timing, weakness and obstruction aetiologies (30).

Recently, an international working group recommended a standardised testing protocol and defined set of pressure and impedance measures ('core outcome set') for application in both clinical and research settings, to address the variability of testing protocols (number of swallows assessed and bolus medium) and reported metrics (*Table 1*) (24). Since that time, normative data of asymptomatic healthy adult controls (18–79 years old) of a moderate cohort size (n=50) reported the mean and standard deviation of core and additional outcome measures (metrics not included in the core outcome set such as BPT and SRI) (37). More recently, the normative data set of a large asymptomatic cohort (n=120) reported

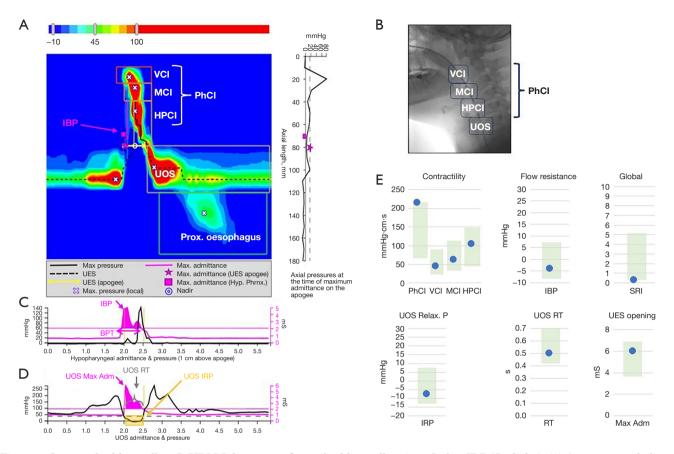


Figure 1 Case 1: a healthy swallow. P-HRM-I data output from a healthy swallow (10 mL thin/IDDSI 0 bolus). (A) Spatiotemporal plot is shown with the coloured boxes representing the velo-, meso- and hypopharynx from which the contractile integrals (VCI, MCI, HPCI) are calculated. The total PhCI is the average of the VCI, MCI and HPCI. (B) The anatomical regions on the spatiotemporal plot correspond to those shown on the lateral VFSS image. An 8-Fr manometry catheter spanning from the velopharynx to the UOS is shown. (C) Hypopharyngeal admittance and pressure graph is shown with IBP represented as the peak of admittance during bolus transit (pink) in the hypopharynx. Note the two yellow vertical lines represent the timing of UOS relaxation and contraction. BPT is represented as the duration of bolus in the hypopharynx (pink peak). (D) UOS admittance and pressure graph is presented. UOS pressure represented by black line, with UOS relaxation pressure (UOS IRP) highlighted in yellow. UOS opening extent (UOS Max Adm) is represented by the pink peak, with the duration of UOS relaxation (UOS RT) is highlighted in greey. (E) P-HRM-I metrics are presented with normal ranges highlighted in green, and swallow specific values represented by the dot (blue, within normal limits). BPT, bolus presence time; HPCI, hypopharyngeal contractile integral; IBP, intrabolus pressure; IDDSI, International Dysphagia Diet Standardization Initiative; IRP, integrated relaxation pressure; Max., maximum; MCI, mesopharyngeal contractile integral; mm, millimeter; mS, millisiemens; PhCI, pharyngeal contractile integral; P-HRM-I, pharyngeal high-resolution manometry with impedance; Prox., proximal; SRI, Swallow Risk Index; UES, upper esophageal sphincter; UOS, upper oesophageal sphincter; UOS Max Adm, UOS maximum admittance; UOS RT, UOS relaxation time; UOS Relax. P, UOS relaxation pressure; VCI, velopharyngeal contractile integral; VFSS, videofluoroscopic swallowing study.

core UOS outcome measures (25). Furthermore, access to an online analysis platform like http://www.swallowgateway. com/ allows the clinician to compare quantified results of an individual patient against these healthy ranges. It is acknowledged that ongoing normative data collection will allow the refinement of diagnostic thresholds.

Results

Case examples demonstrating P-HRM-I analysis and interpretation

These case examples demonstrate the contribution of P-HRM-I data, together with standard imaging, to guide

Table 1 Metrics	often reported	l from a P-HRM-I	assessment ((21)	

Metric class	Metric	Unit	Definition	Clinical significance
Pharyngeal luminal occlusive pressures	Pharyngeal contractile integral (PhCl)	mmHg⋅cm⋅s	A measure of overall pharyngeal contractile vigor	Reduced contractile integrals highlight weakness in one or more regions of the pharynx (velo-, meso- and/or hypopharynx) or across the pharynx overall
	Velopharyngeal contractile integral (VCI)		A measure of contractile vigor spanning the velopharynx only	
	Mesopharyngeal contractile integral (MCI)		A measure of contractile vigor spanning the mesopharynx only	
	Hypopharyngeal contractile integral (HPCI)		A measure of contractile vigor spanning the hypopharynx only	
Hypopharyngeal distension pressure	Hypopharyngeal intra-bolus pressure (IBP)	mmHg	Pressure at 1 cm above to the UOS, at the time of maximum hypopharyngeal distension	Elevated values indicate increased flow resistance across the UOS
UOS relaxation and opening	UOS maximum admittance (UOS Max Adm)	Millisiemens (mS)	The 'extent' of UOS opening	Reduced values indicate reduced UOS opening extent
	UOS integrated relaxation pressure (UOS IRP)	mmHg	A pressure measure of the extent of UOS relaxation	Elevated values indicate impaired UOS relaxation with resulting UOS restriction
	UOS relaxation time (UOS RT)	S	A measure of the duration of UOS relaxation	Reduced values indicate reduced duration of UOS relaxation and opening
Flow timing variable*	Bolus presence time (BPT)	S	The duration that a bolus resides within the pharynx before and after swallowing (derived from impedance)	Elevated values indicate bolus presence in the hypopharynx prior to swallowing, likely due to poor oral containment, delayed pharyngeal trigger or poor pharyngeal clearance
Global swallowing function*	Swallow Risk Index (SRI)	-	A composite measure using to determine global swallowing dysfunction and aspiration risk	Increased values are indicative of disordered swallowing, with values over 15 indicating increased aspiration risk (on concurrent VFSS)

*, additional metric, not part of the 'core outcome set'. P-HRM-I, pharyngeal high-resolution manometry with impedance; UOS, upper oesophageal sphincter; VFSS, videofluoroscopic swallowing study.

clinical management of dysphagia commonly presenting to an ENT practice.

Case 1: a healthy pharyngeal swallow

Reconfiguration of the pharyngeal and laryngeal structures occurs during swallowing in order to propel the bolus through the pharynx and UOS, while concurrently protecting the airway (38). It is important to appreciate the manometric features present in a healthy swallow (*Figure 1*) in order to recognize swallowing abnormalities. The pharyngeal phase of a swallow is divided into four anatomical segments (velo-, meso-, and hypo-pharynx and UOS) (38). In *Figure 1B*, the VFSS image shows the P-HRM-I catheter *in situ* with these anatomical segments highlighted, correlating with the P-HRM-I spatiotemporal plot (*Figure 1A*). The spatiotemporal plot (*Figure 1A*) presents pressure over time with warmer colors of red and orange signifying higher pressures, and cooler colors of blue and green signify regions of lower pressures. Typically, higher pressures are generated during contractions, whereas lower pressures are observed at rest or during periods of relaxation (24). *Figure 1A* shows a healthy swallow manometric sequence: (I) prior to swallowing, the UOS displays tonic contractility which ceases upon initiating a swallow (represented by blue); (II) followed by a sequential pharyngeal contraction from the velo- to the hypopharynx (red); and (III) concludes with contraction of the UOS (red) which returns to the tonic baseline (green) (24).

The graphs shown below the spatiotemporal plot (*Figure 1A*) represent the pressure (black line) generated at the hypopharynx (*Figure 1B*) and at the UOS (*Figure 1C*). The impedance data (pink) represents bolus

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transit. In *Figure 1B*, at the hypopharynx, the bolus arrives (pink) followed by the pharyngeal contractile sequence (black). In *Figure 1C*, at the UOS, the bolus (pink) transits through the UOS during relaxation (black). The drop in UOS pressures to sub-atmospheric levels is represented by UOS integrated relaxation pressure (IRP). The UOS opening extent is represented by UOS maximum admittance (Max Adm) (*Figure 1C*) (24).

Case 2: globus sensation

Globus sensation, commonly referred to as globus, is the intermittent or persistent sensation of a lump or foreign body in the throat (39). Patients with globus commonly present to an Otolaryngologist, with prevalence reported up to 46% (40). The aetiology of globus is unclear, with contributing factors postulated to include gastro-esophageal reflux, UOS dysfunction or hypertonicity, oesophageal dysmotility, anxiety and depression (41-43).

Globus can pose a clinical challenge due to the ambiguity of self-reported symptoms, and can present with or without overt dysphagia symptoms (44). Flexible nasoendoscopy is performed to exclude a structural lesion or changes due to laryngopharyngeal reflux. In the absence of significant self-reported swallowing issues, most patients would be reassured by the Otolaryngology examination findings and discharged (41). In a small number of cases, a P-HRM-I assessment can be beneficial to exclude potential UOS dysfunction or hypertonicity contributing to globus sensation, and provides the patients with biofeedback of healthy swallowing. The P-HRM-I biofeedback is particularly helpful in those patients presenting with minor/ inconsistent swallowing difficulty and globus for confirmation of a swallow within the normal range of P-HRM-I.

For a typical patient, the P-HRM-I output should demonstrate an appropriately coordinated swallow comparable to the spatio-temporal plot shown in the healthy example (*Figure 1A*). Furthermore, pharyngeal and UOS pressure and impedance measures should be within normal limits. An additional advantage of P-HRM-I is that the catheter can be placed to extend across the oesophagus to assess for contributory oesophageal motility disorders such as achalasia or oesophageal spasm (21).

This globus case highlights the benefit of using P-HRM-I to assess pharyngeal swallowing without the use radiation (VFSS). Additionally, P-HRM-I offers an immediate assessment of UOS dysfunction with the potential application of biofeedback for the patient (45). The clinician may review the pharyngeal topography plots of the

individual swallows for features of UOS dysfunction using the recently proposed classification scheme describing UOS disorder types (25).

Case 3: cricopharyngeal bar

A cricopharyngeal bar is thickening of the cricopharyngeal muscle resulting in a narrowing of the UOS (46). Although a cricopharyngeal bar is often diagnosed as an incidental finding on fluoroscopy, in some patients it may be associated with dysphagia and aspiration (47). The reporting of a cricopharyngeal bar on imaging is commonly described as less than or greater than 50% of the lumen. In addition, a cricopharyngeal bar may be classified as non-obstructive and obstructive (48). This is dependent on the degree of narrowing at the cricopharyngeal region which may impede bolus transit, with subsequent retrograde bolus movement misdirected towards the larynx (49) potentially resulting in residue and aspiration. Surgical options include myotomy and dilatation of the UOS (50,51). However, identifying those patients who are most likely to benefit is unclear (47).

Case 3a presents a patient with self-reported worsening dysphagia and on imaging a "small" (<50% of the lumen) cricopharyngeal bar was observed (Figure 2). A cricopharyngeal dilatation did not improve symptoms, questioning subsequent treatment planning. A P-HRM-I swallowing study, in this instance, may provide additional data to ascertain the contribution of the cricopharyngeal bar to swallowing. Biomechanical pressure, timing and impedance metrics measured at the UOS demonstrate values within the normal ranges, inferring adequate UOS function. Therefore, the cricopharyngeal bar at the level of the UOS is not compromising bolus transit through the UOS, and not contributing to dysphagia for this patient (non-obstructive cricopharyngeal bar). Similarly, pharyngeal contractile pressures are within the normal ranges. However, BPT, the duration of the bolus in the pharynx prior to commencing the pharyngeal swallow sequence, is prolonged. This suggests that impaired lingual bolus control is the underlying contributing mechanism for dysphagia. In this case, surgical intervention at the UOS may not be indicated and alternative management of swallowing exercises and dietary modifications could be more appropriate.

In comparison, Case 3b highlights a patient with a "significant/large" (>50% of the lumen) cricopharyngeal bar shown on imaging (*Figure 3*). Biomechanical pressures, timing and impedance metrics at the UOS demonstrated abnormal values, implying UOS dysfunction. The

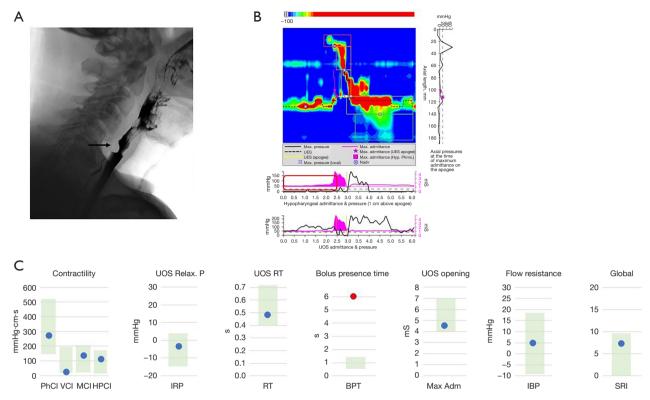


Figure 2 Case 3a: non-obstructive cricopharyngeal bar. A 78-year-old male who presented with a history of progressively worsening dysphagia and dysphonia. He had previously undergone a cricopharyngeal dilatation, with no improvement in symptoms. (A) Lateral VFSS demonstrates a cricopharyngeal bar at C6/7 (arrow) during a swallow. (B) The spatiotemporal plot of a 20-mL thin (IDDSI 0) bolus swallow is shown and consistent with a healthy swallow. In the hypopharyngeal admittance and pressure graph, the pink line (highlighted by red box) shows the bolus in the pharynx prior to swallowing initiation (represented by vertical yellow line). (C) The individual P-HRM-I metrics are shown with patient values represented by dot (blue, within normal limits or red, abnormal). This data demonstrates UOS metrics that are within normal limits, therefore, suggesting that a repeated UOS dilatation is not indicated. Furthermore, the pharyngeal contractile pressures are within normal limits. However, prolonged BPT (red dot) is the key mechanism contributing to the dysphagia, suggesting impaired lingual bolus control. BPT, bolus presence time; HPCI, hypopharyngeal contractile integral; Hyp. Phrnx., hypopharyngeal; IBP, intrabolus pressure; IDDSI, International Dysphagia Diet Standardization Initiative; IRP, integrated relaxation pressure; Max., maximum; Max Adm, maximum admittance; MCI, mesopharyngeal contractile integral; mm, millimeter; mS, millisiemens; PhCI, pharyngeal contractile integral; P-HRM-I, pharyngeal high-resolution manometry with impedance; SRI, Swallow Risk Index; UES, upper esophageal sphincter; UOS RT, UOS relaxation time; UOS Relax. P, UOS relaxation pressure; VCI, velopharyngeal contractile integral; VFSS, videofluoroscopic swallowing study.

cricopharyngeal bar, in this case, is associated with impaired bolus transit through the UOS, evidenced by elevated UOS IRP and hypopharyngeal intrabolus pressures (IBP), contributing to the abnormal global SRI (obstructive cricopharyngeal bar). Furthermore, the spatiotemporal plot shows a sustained pharyngeal contraction with elevated pressures (black line) shown at the hypopharyngeal and UOS graphs. In this case, the P-HRM-I findings concur with imaging that surgical intervention at the UOS is indicated.

These cases highlight the added value of a P-HRM-I assessment in patients with a cricopharyngeal bar. If a cricopharyngeal bar is contributing to dysphagia, the characteristic features are elevated UOS IRP and hypopharyngeal IBP impeding bolus transit through the UOS (52). These altered metrics have been recognized

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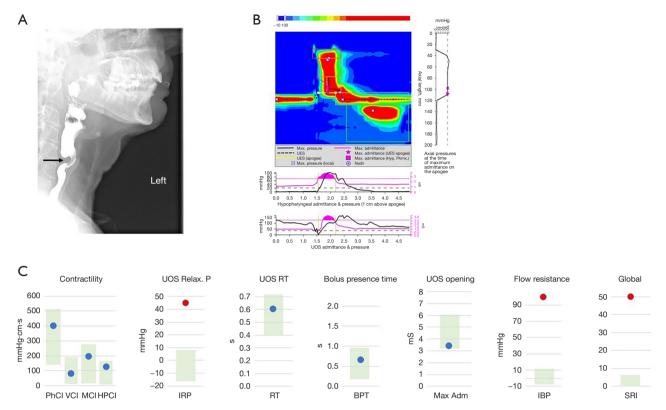


Figure 3 Case 3b: obstructive cricopharyngeal bar. A 53-year-old female who presented with a history of chronic dysphagia and regurgitation. (A) An acquisition image captured during a VFSS demonstrating a large cricopharyngeal impression (arrow) at C6/7. (B) The spatiotemporal plot of a 5 mL thick (IDDSI 4) bolus swallow demonstrates a prolonged pharyngeal chamber pressurisation, lacking features consistent with an appropriately coordinated swallow. The hypopharyngeal pressure and admittance graph shows elevated hypopharyngeal generated pressure (black line) during bolus transit (pink peak). This is also seen in the UOS pressure and admittance graph. (C) The individual P-HRM-I metrics are shown with patient values represented by dot (blue, within normal limits or red, abnormal). The abnormal P-HRM-I metrics (red dot) for this patient include elevated UOS relaxation pressure (UOS IRP) and elevated flow resistance (hypopharyngeal IBP), contributing to the abnormal global swallowing dysfunction (SRI). These data imply UOS dysfunction associated with the cricopharyngeal bar, indicating this patient may benefit from UOS dilatation or surgical intervention. BPT, bolus presence time; HPCI, hypopharyngeal contractile integral; Hyp. Phrnx., hypopharyngeal; IBP, intrabolus pressure; IDDSI, International Dysphagia Diet Standardization Initiative; IRP, integrated relaxation pressure; Max., maximum; Max Adm, maximum admittance; MCI, mesopharyngeal contractile integral; mm, millimeter; mS, millisiemens; PhCI, pharyngeal contractile integral; P-HRM-I, pharyngeal high-resolution manometry with impedance; SRI, Swallow Risk Index; UES, upper esophageal sphincter; UOS, upper oesophageal sphincter; UOS Relax. P, UOS relaxation pressure; UOS RT, UOS relaxation time; VCI, velopharyngeal contractile integral; VFSS, videofluoroscopic swallowing study.

to identify those potentially suitable for surgical management (47). The imaging categorisation of a small or large cricopharyngeal bar may be an insufficient measure, however, additional P-HRM-I assessment demonstrating altered UOS measures can identify those patients who will likely benefit from surgical intervention.

Case 4: dysphagia post-head and neck cancer treatment Dysphagia following head and neck cancer treatment is reported in up to 50% of patients (53) impacting patient's quality of life. The presentation of dysphagia in this population varies due to the anatomical location, staging and treatment type (53). Dysphagia may manifest as a combination of tongue base dysfunction, reduced laryngeal elevation, reduced pharyngeal contraction, impaired epiglottic movement and reduced UOS opening (54,55), making management challenging. P-HRM-I can distinguish the underlying biomechanical features of dysphagia,

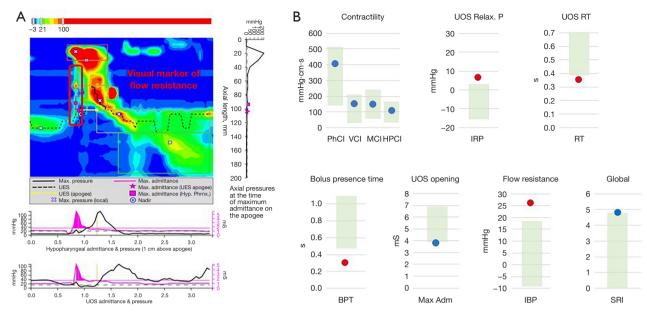


Figure 4 Case 4a: UOS stricture post-head and neck cancer treatment. A 74-year-old male presenting with dysphagia, and history of T2N0M0 left supraglottic squamous cell carcinoma treated with chemoradiotherapy. There were no concerning features on history, with a normal head & neck examination and flexible nasoendoscopy. (A) The spatiotemporal plot for a 10 mL thin (IDDSI 0) bolus swallow shows an abnormal increased pressure in the pharyngeal chamber (green column in red box) at swallow initiation, suggesting increased flow resistance at the UOS. (B) The P-HRM-I metrics show that the pharyngeal contractile pressures are within the normal ranges (blue dot). However, abnormal metrics (red dot) include elevated UOS relaxation pressure (UOS IRP), shortened UOS relaxation duration (UOS RT) and elevated hypopharyngeal IBP, indicating UOS dysfunction and patient may be suitable for UOS dilatation. At the level of the hypopharynx BPT is shortened, suggestive of a possible compensatory response to propel the bolus through the restricted UOS. BPT, bolus presence time; HPCI, hypopharyngeal contractile integral; Hyp. Phrnx., hypopharyngeal; IBP, intrabolus pressure; IDDSI, International Dysphagia Diet Standardization Initiative; IRP, integrated relaxation pressure; Max., maximum; Max Adm, maximum admittance; MCI, mesopharyngeal contractile integral; mm, millimeter; mS, millisiemens; PhCI, pharyngeal contractile integral; P-HRM-I, pharyngeal high-resolution manometry with impedance; SRI, Swallow Risk Index; UES, upper esophageal sphincter; UOS, upper oesophageal sphincter; UOS Relax. P, UOS relaxation pressure; UOS RT, UOS relaxation time; VCI, velopharyngeal contractile integral; VFSS, videofluoroscopic swallowing study.

whether localised at the pharynx or UOS to inform targeted treatment.

Case 4a illustrates impeded bolus transit (increased flow resistance) through the UOS, evidenced by elevated hypopharyngeal IBP, elevated relaxation pressure (UOS IRP) and shortened UOS relaxation duration (UOS RT) (*Figure 4*). However, the pharyngeal contractile pressures are within the normal ranges. In this case, these results indicate UOS dysfunction, consistent with a UOS stricture often reported in post-head and neck cancer patients (56), and the patient may benefit from dilatation (57). Notably, diagnosis of stricture on radiological imaging is inconsistent, particularly in cases where pharyngeal weakness is also present (29,58).

In contrast, Case 4b demonstrates marked reduction of

the pharyngeal contractile pressures [pharyngeal contractile integral (PhCI)], specifically at the sub-anatomical regions at the velo- [velopharyngeal contractile integral (VCI)] and meso-pharynx [mesopharyngeal contractile integral (MCI)] (*Figure 5*). These findings are illustrative of weak pharyngeal contractile pressures, also recently reported in a post-head and neck cancer cohort (59). Impaired pressure generation at the velopharynx and mesopharynx compromises bolus propulsion through the pharynx (60,61) with resulting pharyngeal residue. The spatiotemporal plot shows a repeated swallow, indicating an attempt to clear pharyngeal residue. Similar to Case 3a, prolonged BPT represents impaired lingual bolus control. The prolonged BPT in conjunction with weak pharyngeal pressures contributes to the elevated global SRI. In this case, UOS measures were Page 10 of 14

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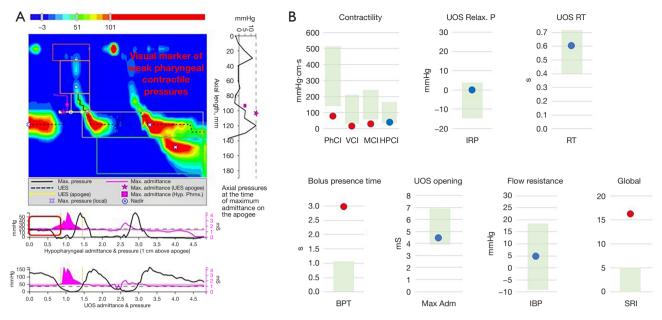


Figure 5 Case 4b: weak pharyngeal propulsion post-head and neck cancer treatment. A 62-year-old male presented with dysphagia and history of treatment for a T4aN1M1 oropharyngeal squamous cell carcinoma recurrence comprising of a left partial glossectomy, pharyngectomy, level I–IV neck dissection and mandibulectomy with free flap reconstruction. Previous treatment included chemoradiotherapy. (A) Spatiotemporal plot for a 10 mL thin (IDDSI 0) bolus showing a repeated swallow present. The generated pharyngeal contractile sequence is weak (translucent/pale green pressure) in the velo- and mesopharynx. The hypopharyngeal admittance and pressure graph, demonstrates presence of the bolus in the pharynx prior to swallow initiation (red box). (B) The abnormal (red dot) P-HRM-I metrics of reduced pharyngeal contractility (VCI and MCI) and increased BPT contribute to the elevated global swallowing metric (SRI). The hypopharyngeal contractile pressures (HPCI), UOS metrics (IRP, RT, and Max Adm) and IBP are within the normal ranges (blue dot). These data suggest the patient is not indicated for UOS intervention, instead targeted intervention at the pharyngeal level is required. BPT, bolus presence time; HPCI, hypopharyngeal contractile integral; Hyp. Phrnx., hypopharyngeal; IBP, intrabolus pressure; IDDSI, International Dysphagia Diet Standardization Initiative; IRP, integrated relaxation pressure; Max., maximum; Max Adm, maximum admittance; MCI, mesopharyngeal contractile integral; mm, millimeter; mS, millisiemens; PhCI, pharyngeal contractile integral; P-HRM-I, pharyngeal high-resolution manometry with impedance; SRI, Swallow Risk Index; UES, upper esophageal sphincter; UOS, upper oesophageal sphincter; UOS Relax. P, UOS relaxation pressure; UOS RT, UOS relaxation time; VCI, velopharyngeal contractile integral.

within the normal ranges. These data suggest that this patient may benefit from swallowing exercises targeting pharyngeal strength and dietary modifications. Of interest, our group utilised P-HRM-I to assess biomechanical swallowing before and after a novel tongue base augmentation procedure (62) aiming to improve pharyngeal pressure generation (63).

Discussion

P-HRM-I is gaining increasing recognition as a valuable swallowing assessment method that can provide precise and quantitative measures of swallowing biomechanics. P-HRM-I assessment extends current swallow imaging assessments by identifying the biomechanical impairment at the pharynx and/or UOS, providing considerable opportunity for clinical application.

The presented case examples illustrate P-HRM-I analysis and interpretation of patients presenting to an Otolaryngologist with dysphagia symptoms, compared to a healthy swallow. Unlike traditional imaging swallowing assessment methods, P-HRM-I discriminates pharyngeal and UOS function using pressure and impedance measures. The PhCI metrics provide direct measure of generated pressures within the pharyngeal lumen (24), in contrast to inferred dysfunction from imaging assessments. Furthermore, P-HRM-I at the UOS is considered advantageous when compared with imaging assessments (58). It is important to

recognise that an abnormal P-HRM-I metric in isolation may not be clinically significant and should be considered in conjunction with patient reported symptoms and any available imaging assessments.

This paper illustrates the potential for P-HRM-I assessments in extending the interpretation of imaging swallowing assessments. These findings may facilitate the continued clinical adoption of this novel technology by Otolaryngologists. P-HRM-I can be utilised either as a stand-alone assessment or as an adjunct to fluoroscopy (VFSS). In our centre, we utilise stand-alone P-HRM-I mainly in patients with an anticipated low aspiration risk, patients who have had VFSS recently performed (10,25) or in those where VFSS is difficult to perform due to patient or location factors (intensive care unit patients). A concurrent VFSS and P-HRM-I will be conducted in patients with known severe dysphagia.

To improve translation into the clinical setting, a classification scheme has recently been proposed, categorising the biomechanical breakdown at the pharynx and UOS (25). Additionally, quantitative P-HRM-I metrics allow for longitudinal assessments of dysphagia or to evaluate and guide the true effect of an intervention (dietary modifications, targeted swallowing exercises, UOS dilatation, cricopharyngeal myotomy or Botox) on swallowing outcomes (17,18,19,20,64). These quantitative measures enable demonstration of treatment efficacy.

Despite the benefits of the P-HRM-I technology, acknowledged barriers for clinical translation include equipment cost and training (8). Training opportunities are available online to assist clinicians with analysis and interpretation. Additionally, cloud-based platforms provide for multi-disciplinary collaboration (24). It is acknowledged that P-HRM-I does not provide visualisation of the swallow. However, an advantage of P-HRM-I is that it can be performed concurrently with imaging (VFSS) (9,33,65), thereby providing a comprehensive assessment of swallowing including detection of sub-clinical changes not detected on imaging alone (18,66). We anticipate that continued and increased uptake of P-HRM-I across multiple centres may facilitate the collection of normative data, improving the generalisability of this translational swallowing assessment technology.

Conclusions

Dysphagia is often multifactorial or complex in nature. P-HRM-I extends current imaging swallowing assessments enabling the Otolaryngologist to quantify and localise the biomechanical features contributing to pharyngeal dysphagia. In the case examples shown, the analysis and interpretation of the P-HRM-I metrics discriminate impairment of pharyngeal function from UOS dysfunction, and can inform treatment decision-making for the provision of individualised management.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related

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to the accuracy or integrity of any part of the work are appropriately investigated and resolved. Case examples have been extracted from studies conducted at two centres: Flinders Medical Centre (Adelaide, Australia) and St. George Hospital (Sydney, Australia) (approved by Southern Adelaide Clinical Human Research Ethics Committee and St. Vincent's Hospital Human Research Ethics Committee). Informed consents are not required due to the retrospective nature of this study.

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