

1 **Running head:** High resolution cervical auscultation to

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3 acoustic signals

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5 **Authors:** Atsuko Kurosu, M.A.<sup>1</sup>; James L. Coyle, Ph.D.<sup>1,2</sup>; Joshua Dudik, Ph.D.<sup>3</sup>; Ervin Sejdic,  
6 Ph.D.<sup>3</sup>

7 **Name of the institution where the study was performed:** University of Pittsburgh

8 **Institutional affiliation of the authors:** University of Pittsburgh, School of Health and  
9 Rehabilitation and Sciences, Department of Communication Science and Disorders<sup>1</sup>

10 University of Pittsburgh, Department of Otolaryngology, School of Medicine<sup>2</sup>

11 University of Pittsburgh, Swanson School of Engineering, Department of Electrical and  
12 Computer Engineering<sup>3</sup>

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19 **Corresponding author:** Atsuko Kurosu, 5010 Forbes tower, University of Pittsburgh,  
20 Pittsburgh, PA15260, 412-894-5085, atk22@pitt.edu

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## Title

High resolution cervical auscultation to detect swallow kinematic events, part 1: the acoustic signals

## Abstract

**Objective:** To examine whether there were any associations between high resolution cervical auscultation (HRCA) acoustic signals recorded by a contact microphone and swallowing kinematic events during pharyngeal swallow as assessed by a videofluoroscopic examination.

**Design:** Prospective pilot study

**Setting:** University teaching hospital, University research laboratories

**Participants:** 35 patients with stroke who have suspected dysphagia (26 males + 9 females; age =  $65.8 \pm 11.2$ ).

**Methods:** Videofluoroscopic recordings of one hundred liquid swallows from 35 stroke patients were analyzed, and a variety of HRCA signal features to characterize each swallow were calculated.

**Main Outcome Measure(s):** Percent of signal feature maxima (peak) occurring within 0.1 seconds of swallow kinematic event identified from videofluoroscopic recording

**Results:** Maxima of HRCA signal features, such as standard deviation, skewness, kurtosis, centroid frequency, bandwidth, and wave entropy, were associated with hyoid elevation, laryngeal vestibule closure, and upper esophageal sphincter opening, and the contact of the base of the tongue and posterior pharyngeal wall.

**Conclusions:** Although the kinematic source of HRCA acoustic signals has yet to be fully elucidated, these results indicate a strong relationship between these HRCA signals and several

23 swallow kinematic events. There is a potential for HRCA to be developed for diagnostic and  
24 rehabilitative clinical management of dysphagia.

25 **Key Words:** swallowing, deglutition, deglutition disorders, high resolution cervical auscultation,  
26 signal analysis

27 **Abbreviations:** Base of the tongue (BOT), Cervical auscultation (CA), Hyoid elevation (HE),  
28 High resolution cervical auscultation (HRCA), Laryngeal vestibule (LV), Laryngeal vestibule  
29 closure (LVC), Posterior pharyngeal wall (PPW), Speech language pathologist (SLP), Upper  
30 esophageal sphincter (UES), Upper esophageal sphincter opening (UESO), Videofluoroscopic  
31 studies (VFSS)

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## 34 **Manuscript body**

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36 Stroke is the fourth leading cause of death in the U.S. <sup>1</sup>, with approximately 800,000 people in  
37 the U.S. diagnosed with a new or recurrent stroke each year <sup>2</sup>. Swallowing difficulties or  
38 dysphagia, are common in this population <sup>3,4</sup> with more than half of the admitted acute stroke  
39 patients showing signs of dysphagia, such as coughing after swallowing during screening and/or  
40 aspiration during videofluoroscopic examination of swallowing (VFSS) <sup>3-5</sup>. Stroke patients with  
41 dysphagia are three times more likely to develop pneumonia than those without <sup>4</sup>. As a further  
42 complication, aspiration pneumonia is associated with substantial morbidity and mortality <sup>6</sup>, as  
43 well as a financial impact of over \$1.3 billion dollars a year in the U.S <sup>7</sup>. In order to efficiently  
44 identify the potential risks to the well-being of these patients, a reliable, non-invasive, and  
45 inexpensive screening test should be administered <sup>3,5</sup> so that patients with suspected dysphagia

46 can be referred to VFSS, a gold standard objective evaluation of swallowing for further  
47 assessments <sup>8</sup>.

48 Stethoscope-based cervical auscultation (CA), one of the screening tests currently used in  
49 acute care and rehabilitation clinical settings, is to evaluate swallowing performance based on  
50 the acoustic properties <sup>9,10</sup>. However, CA's reliability and validity has been questioned <sup>9,11,12</sup>.  
51 There is no consensus on the kinematic source of CA signals, or agreement regarding what the  
52 observed sounds represent, which are highly variable among different individuals <sup>10</sup>.

53 High resolution cervical auscultation (HRCA) is a combined hardware-software system that  
54 merges electronic transducers with advanced data analysis methods. In the place of a  
55 stethoscope, HRCA uses digital electronic transducers, such as a microphone and/or an  
56 accelerometer, to record acoustic and vibratory signals caused by movements of the upper  
57 aerodigestive tract and boluses during swallowing <sup>13,14</sup>. After the recording of acoustic and  
58 vibratory signals, HRCA signal features were extracted. Allowing for a deeper, more detailed  
59 analysis unimpeded by the limitations and biases of manual acoustic analysis suggests that  
60 HRCA offers greater potential than traditional CA as a dysphagia screening method <sup>13,15,16</sup>.  
61 However, similar to the stethoscope CA, the kinematic sources of HRCA acoustic signals  
62 recorded during swallowing have not yet been elucidated, because few studies have investigated  
63 the sources <sup>17,18</sup>.

64 Concurrent HRCA and VFSS studies allow comparing acoustic signal features and  
65 swallowing kinematic events identified on videofluoroscopic images. If signal features are  
66 temporally associated with a specific swallow kinematic event, then the HRCA signal can be  
67 said to be associated with the swallow event that has occurred. For example, in order to correlate

68 the two modalities, we could compare when specific kinematic events occurred to when our  
69 chosen mathematical features exhibited a local maximum (peak). Any local maximum that  
70 occurred within 0.1 seconds of when a kinematic event occurred, as determined by the  
71 concurrent VFSS exam, could be declared as being associated with the event. It's worth noting  
72 that 0.1 seconds window is the tolerance range for obtaining high reliability between raters for  
73 manual VF analysis <sup>19</sup>. Furthermore, if a local maximum of one specific feature occurs within 0.1  
74 second in the majority of the swallows, we would declare that there is association between the  
75 acoustic signal feature itself and the swallow event.

76 This study serves as a preliminary study to determine whether or not a larger, more detailed  
77 investigation of the correlation between HRCA acoustic signals and swallow kinematic events is  
78 warranted. The purpose of this preliminary study was to examine whether there are any  
79 associations between HRCA acoustic signals recorded by a contact microphone and swallow  
80 kinematic events assessed by VFSS. We hypothesized that HRCA acoustic signal features would  
81 be associated with laryngeal vestibule closure (LVC), laryngeal vestibule (LV) re-opening, upper  
82 esophageal sphincter opening (UESO), and the contact of the base of the tongue (BOT) and  
83 posterior pharyngeal wall (PPW) (BOT-PPW contact) <sup>20</sup> which were postulated to be the  
84 potential kinematic sources of CA signals. We further hypothesized that acoustic signal features  
85 also would be associated with hyoid elevation (HE) which has high correlation between LVC and  
86 UESO <sup>21,22</sup>, and reported to be the kinematic source of CA signals <sup>23</sup> as well as vibratory signals  
87 recorded via an accelerometer <sup>18</sup>

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## Methods

90 **Ethical considerations**

91 This study was approved by the University of Pittsburgh Institutional Review Board.

92 **Participants**

93 35 patients with stroke (26 males + 9 females; age=65.8 ± 11.2) who were suspected of  
94 having dysphagia and who were scheduled for a VFSS at the Presbyterian Hospital of the  
95 University of Pittsburgh participated in this study. All participants were referred by their physician  
96 for a VFSS because of clinical suspicion of dysphagia based on diagnosis or clinical evaluation. All the  
97 participants signed informed consent prior to the study. Table 1 summarizes the patient  
98 demographic information.

99 **Data collection**

100 **VFSS recording**

101 During the VFSS, patients were instructed to swallow chilled (5 C) thin liquid barium boluses  
102 (Varibar thin liquid with < 5 cps viscosity), placed in the patients' mouths with a 5ml tea-spoon by  
103 a speech language pathologist (SLP). The bolus volumes were approximately 3ml, though the  
104 exact volumes were not measured. Patients also swallowed self-selected comfortable volume  
105 boluses of liquid barium from a cup. The bolus volume for the swallows from a cup, bolus  
106 consistency, number of swallow trials for each bolus consistency, and the order of the bolus  
107 presentation were not controlled for the current study. They were determined by the SLP who  
108 administered the VFSS, and our research protocol was intended to avoid interfering with the  
109 procedure's clinical purposes in any significant way.

110 Among the swallows produced by the 35 patients, one hundred liquid swallows produced as a  
111 single swallow or the first of multiple swallows per bolus were analyzed. These swallows  
112 include 39 thin liquid swallows administered by a tea-spoon and 61 thin liquid swallows from a  
113 cup without any swallow maneuvers or postural changes.

114 VFSS images output by the x-ray machine (Ultimax system, Toshiba, Tustin, CA) were  
115 input to a video capture card (AccuStream Express HD, Foresight Imaging, Chelmsford, MA)  
116 and recorded with the LabView program Signal Express (National Instruments, Austin TX) at a  
117 rate of 30 frames per second.

### 118 **HRCa signal recording**

119 Concurrent to VFSS, swallowing vibration and acoustic signals were recorded by a tri-axial  
120 accelerometer (ADXL 327, Analog Devices, Norwood, MA) which was attached to the  
121 participant's anterior neck at the level of the cricoid cartilage. A contact microphone (Model  
122 C411L, AKG, Vienna, Austria) was placed below the accelerometer and slightly towards the  
123 right lateral side of the trachea<sup>24</sup>. This study includes only the microphone results because the  
124 accelerometer and microphone record different signals that are complementary but not  
125 comparable to one other<sup>25</sup>. Figure 1 shows the microphone and accelerometer on a VF image.  
126 The microphone signals (10Hz to 10k Hz) were unfiltered in order to record the entire dynamic  
127 range, and they were sampled at 20 kHz by a custom LabView program. (National Instruments,  
128 Austin, TX) and they were sent to a National Instruments 6210 DAQ.

### 129 **Measurements**

130 VFSS and acoustic signal data were analyzed step by step. Figure 2 indicates the flowchart of  
131 the data analysis. Figure 3 illustrates the VFSS images and the concurrent acoustic signal after  
132 filtering noise artifacts from a single swallow.

133 **Swallow kinematic temporal analysis**

134 First, in order to properly analyze the acoustic signals, the onset and offset of each pharyngeal  
135 swallow were identified. Table 2 indicates the definition of the time points of the swallow  
136 kinematic analysis<sup>13,26,27</sup>. Next, the nine measures listed in Table 2 that were related to HE,  
137 LVC, UESO, and BOT-PPW contact were obtained from the VF recording by the primary rater.  
138 All time points were identified from the VF recording of each swallow in a lateral view played  
139 both in slow motion and frame-by-frame by the primary rater, an experienced SLP (1<sup>st</sup> author).  
140 Ten percent of all measures were randomly selected and repeated by the same rater as well as the  
141 2<sup>nd</sup> rater (2<sup>nd</sup> author), who also is an experienced SLP, to determine intra-rater and inter-rater  
142 reliability. Both raters were blinded to the acoustic signal data. The intraclass correlation  
143 coefficient both for the intra- and inter- rater reliability was 1.0. All videofluoroscopic images  
144 were analyzed with Image J software (Image J, National Institutes of Health, Bethesda, MD).

145 **Acoustic signal processing and analysis**

146 After filtering noise artifacts from acoustic signals, several HRCA signal features were  
147 computed. First, time domain features, which were (1) standard deviation, (2) skewness, (3)  
148 kurtosis, (4) entropy rate, and (5) Lempel-Ziv complexity were calculated to analyze each  
149 swallow<sup>28</sup>. Second, frequency domain features, which were (1) peak frequency, (2) centroid  
150 frequency, (3) bandwidth, and (4) wavelet entropy, also were calculated. All the features were  
151 calculated, not as a single value, but as time-varying parameters by applying 100 msec sliding  
152 non-overlapping window to our signals. Finally, local maxima (peak) that occurred within 0.1  
153 seconds of each of the nine swallowing events were identified for each signal feature. Table 3  
154 indicates the definitions of the statistical features and equations used to calculate them.



155 **Statistical analysis**

156 Binomial tests were used to test whether the proportions of the occurrence of a local  
157 maximum for each swallow event were statistically significant. The cutoff point was set to be  
158 60% in order to minimize the effect of noise and random chance on our results.  $p < 0.05$  was  
159 accepted as significant for all the tests. IBM SPSS statistics 23.0 (IBM Corporation, Armonk,  
160 NY) was used for the statistical analyses. Maxima occurring within 0.1 seconds of a swallow  
161 event indicated that the event and feature were associated.

162

163 **Results**

164 **Swallow kinematic analysis**

165 None of the participants showed anatomical abnormalities. Among the 100 swallows, there  
166 were 11 swallows with aspiration and 57 swallows with laryngeal penetration. LVC onset and  
167 LV re-open from three swallows, UESO onset and UES re-closure from two swallows, and  
168 BOT-PPW contact onset and BOT-PPW re-open from two swallows were excluded from  
169 analysis because they could not be identified on VF images.

170 **Acoustic signal analysis**

171 Local maxima of Lempel-Ziv complexity and entropy rate were not observed within 0.1  
172 seconds of any of the swallow events (0%;  $p < .0001$ ). Peak frequency maxima did not  
173 demonstrate statistical significance at the 60% level on any swallow events. Figure 4 indicates  
174 the percent of signal feature maxima occurring within 0.1 seconds of each swallow kinematic  
175 event.

176 As can be appreciated in Figure 4, there were several associations between signal features and  
177 swallow kinematic events. Standard deviation maxima were associated with hyoid onset (61%;  
178  $p < .0001$ ) and UESO onset (76%;  $p < .0001$ ). Skewness maxima were associated with all swallow  
179 kinematic events in more than 80% of the cases ( $p < .0001$ ). Kurtosis maxima were associated  
180 with LVC onset (70%;  $p = .02$ ), LV re-open (69%;  $p = .048$ ), and UESO onset (72%;  $p = .01$ ).  
181 Centroid frequency maxima were associated with hyoid rest (75%;  $p < .0001$ ), LVC onset (71%;  
182  $p = .01$ ), and UESO onset (66%;  $p < .0001$ ). Bandwidth maxima were associated with hyoid max  
183 (71%;  $p = .02$ ), hyoid rest (75%;  $p < .0001$ ), LVC onset (71%;  $p = .01$ ), UESO onset (68%;  
184  $p < .0001$ ), and BOT-PPW contact onset (69%;  $p = .04$ ). Wavelet entropy maxima were associated  
185 with hyoid onset (60%;  $p < .0001$ ) and UESO onset (60%;  $p < .0001$ ). No other features  
186 demonstrated statistical significance at the 60% level.

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## Discussion

189 The purpose of this preliminary study was to test whether there were any associations  
190 between HRCA acoustic signals and swallow kinematic events as assessed by HRCA and VFSS  
191 in one hundred liquid swallows produced from 35 patients with stroke who have suspected  
192 dysphagia. We compared nine acoustic signal features to nine kinematic swallow events and  
193 found strong associations between several pairs of signal features (i.e., standard deviation,  
194 skewness, kurtosis, centroid frequency, bandwidth, and wavelet entropy) and HE, LVC, UESO,  
195 and BOT-PPW contact related swallow kinematic events. The probability that any one signal  
196 maximum would co-occur with any one single kinematic swallow event is one in nine or 11.1%,

197 any two physiologic events is 1.23%, and so on. Hence our 60% cutoff demonstrates strong  
198 associations for the signal-kinematic pairs.

199 Our results support our a priori hypothesis that HRCA acoustic signals were associated LVC,  
200 LV re-open, UESO, and BOT-PPW contact. These results corresponded with the cardiac analogy  
201 hypothesis that suggests that CA acoustic signals are generated via vibrations caused by valve  
202 and pump systems within the aerodigestive tract including LVC, LV-reopen, UESO, and BOT-  
203 PPW contact<sup>20</sup>. The results agreed with the data from Moriniere and colleagues (2007) that  
204 indicated the association between CA acoustic signals and LV re-open in 15 healthy adults,  
205 although they did not measure the exact onset times<sup>23</sup>. Our results indicating the association  
206 between feature maxima and UES re-closure did not correspond with the data from Perlman and  
207 colleagues (2005) that indicated no correlation between acoustic peak and UES complete closure  
208 defined as the time point when the bolus has just passed UES<sup>29</sup>. However, Perlman et al. (2005)  
209 examined the acoustic peaks and their association with swallow events while we analyzed signal  
210 features of the acoustic signals. The results also did not correspond with the data from Moriniere  
211 and colleagues (2007) that reported CA acoustic sound was observed during UESO and the bolus  
212 passing through UES<sup>23</sup>. Yet, they examined the sounds during the bolus passing through UES  
213 whereas the current study measured the UES re-closure which occurs after the bolus passes UES.  
214 These experimental condition differences may explain the discrepancy in the results.

215 Our results also support our a priori hypothesis that HRCA acoustic signals were associated  
216 with HE. These results corresponded with the data from Moriniere and colleagues (2007) that  
217 indicate the association between CA acoustic signals and HE in 15 healthy adults, although the  
218 exact onset time of the acoustic signals or swallow events were not measured in their study<sup>23</sup>.

219 Zoratto and colleagues (2010) indicated the correlation between hyolaryngeal excursion, i.e., the  
220 displacement of the hyoid and arytenoids, and HRCA vibratory signals<sup>18</sup>. However, it is not  
221 clear whether HRCA acoustic and vibratory signals share the same kinematic sources.

222 Reduced HE and LVC result in penetration and/or aspiration before, during, and/or after the  
223 swallow<sup>21</sup>, and are prevalent in a number of neurogenic populations with dysphagia including  
224 lateral medullary and other cerebral infarctions, Parkinson disease, iatrogenic and pathology-  
225 related peripheral nerve injury and others<sup>21</sup>. Reduced UESO also causes laryngeal penetration,  
226 aspiration of post-swallow residue<sup>30</sup>, and contributes to significant post-onset morbidity in these  
227 populations<sup>21</sup>. Reduced BOT-PPW contact may result in residue in the vallecular that also may  
228 cause penetration and/or aspiration after the swallow<sup>21</sup>. As can be appreciated in Figure 4, some  
229 HRCA acoustic signal features were associated with specific swallow kinematic events stronger  
230 than other signals: skewness maxima were associated with all the swallow events whereas other  
231 feature maxima were associated with selected swallow events. These results suggest that, if  
232 implemented with the aid of advanced computer analysis, the combination of HRCA acoustic  
233 signal features may potentially contribute to efforts to identify impairments of swallow kinematic  
234 events that may cause reduced airway protection when imaging studies are unavailable. If so, this  
235 could lead to HRCA offering several advantages over traditional CA and other swallow  
236 screening techniques. HRCA potentially add diagnostic information to traditional dysphagia  
237 screening techniques which are unable to identify asymptomatic penetration/aspiration or the  
238 kinematic explanations for aspiration that is the most salient target of rehabilitative intervention  
239 for people with dysphagia. Indeed, previous studies indicated that HRCA could grossly  
240 discriminate swallows with and without deep penetration/aspiration with high sensitivity<sup>13,31</sup>.  
241 Further studies are needed to investigate HRCA's capability of detecting penetration/aspiration.

242 Nevertheless, several sequential cascading neuromuscular events, such as HE, LVC, UESO,  
243 and BOT-PPW contact, occur when the pharyngeal swallow is triggered <sup>21</sup>. It is not clear  
244 whether observed signal feature maxima were truly the results of each swallow physiological  
245 event or combinations of several events, or another phenomenon. There were several other bolus  
246 movements in the aerodigestive tract, such as residue in the valleculae and/or pyriform sinuses  
247 and penetration/aspiration, which may have caused a local maximum for some of the signal  
248 features. Although the results are promising, further studies with larger sample size are needed to  
249 specify the relationship between feature maxima and swallow kinematic events. Furthermore,  
250 although we have found associations between signal features and swallow events, the kinematic  
251 nature of these differences in time and frequency domains of these signals remain unclear.

252

### 253 **Study limitations**

254 There are limitations to the current study. The sample size is small because this study was  
255 designed as a preliminary analysis. Patients' age, sex, and lesion sites, which may influence the  
256 swallow kinematic analysis <sup>21,26,32-34</sup> and acoustic signals <sup>14</sup> were not controlled. The number of  
257 swallows made by each participant and the exact bolus volume for each swallow also was not  
258 measured due to the limitations of our data collection methods and our imperative to avoid  
259 interference with the administration and clinical purposes of the VFSS. However, our sample  
260 accurately represents the real-world conditions in which HRCA would be deployed as a swallow  
261 screening method and so should provide comparable results. Finally, all data were obtained from  
262 stroke patients. The results of the study may not represent swallow acoustic signals in healthy  
263 participants <sup>17</sup>.

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### **Future research**

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Currently, we are investigating the association between HRCA vibratory signals and swallow kinematic events. The results will be published separately.

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### **Conclusions**

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In this study, we sought to compare the timing of several notable swallow kinematic events with the corresponding HRCA acoustic swallowing profile in stroke patients. We found that some of the higher-level signal features of HRCA acoustic signals were associated with certain swallow kinematic events. Although the kinematic source of HRCA acoustic signals has yet to be fully elucidated, these results indicate a strong relationship between these HRCA signals and several swallow kinematic events. We advocate for further research into this subject in order to further understand the nature of HRCA and correlation between HRCA and swallow kinematic events, and the potential for HRCA as an adjunct to rehabilitation of dysphagia.

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## **Figure legends**

Figure 1. The accelerometer and microphone on a videofluoroscopic image.

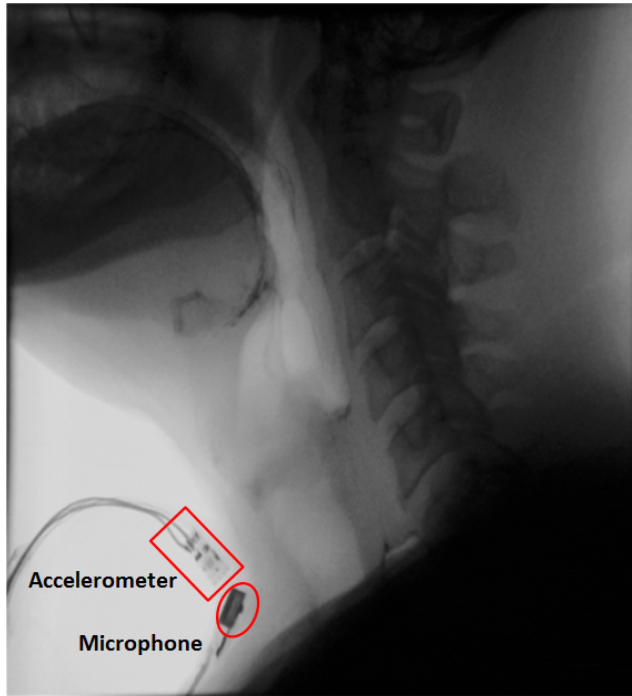
Figure 2. Data analysis flowchart.

Figure 3. Illustration of the concurrent acoustic signal and VFSS images from a single swallow.

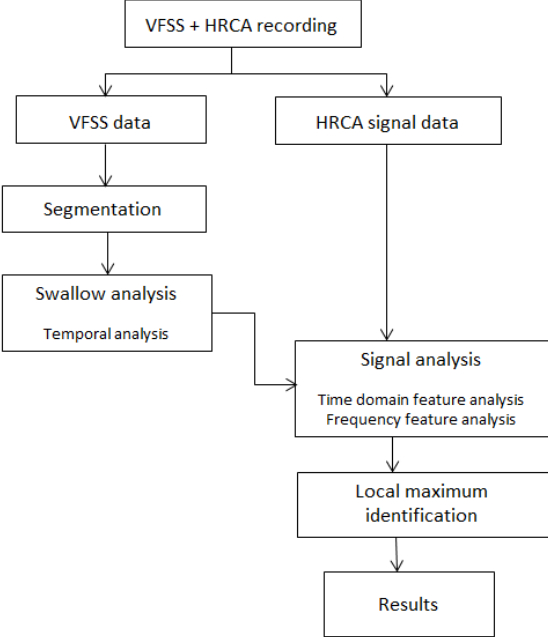
Figure 4. Percent of the signal feature maxima occurring within 0.1 seconds of each swallow kinematic event.

## Figures

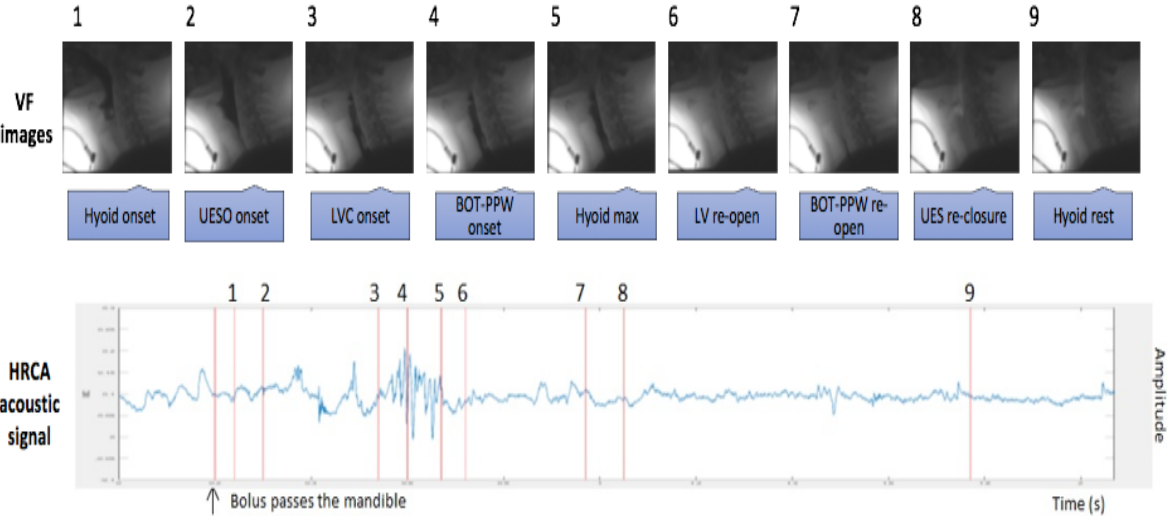
**Figure 1.** The accelerometer and microphone on a videofluoroscopic image.



**Figure 2.** Data analysis flowchart.

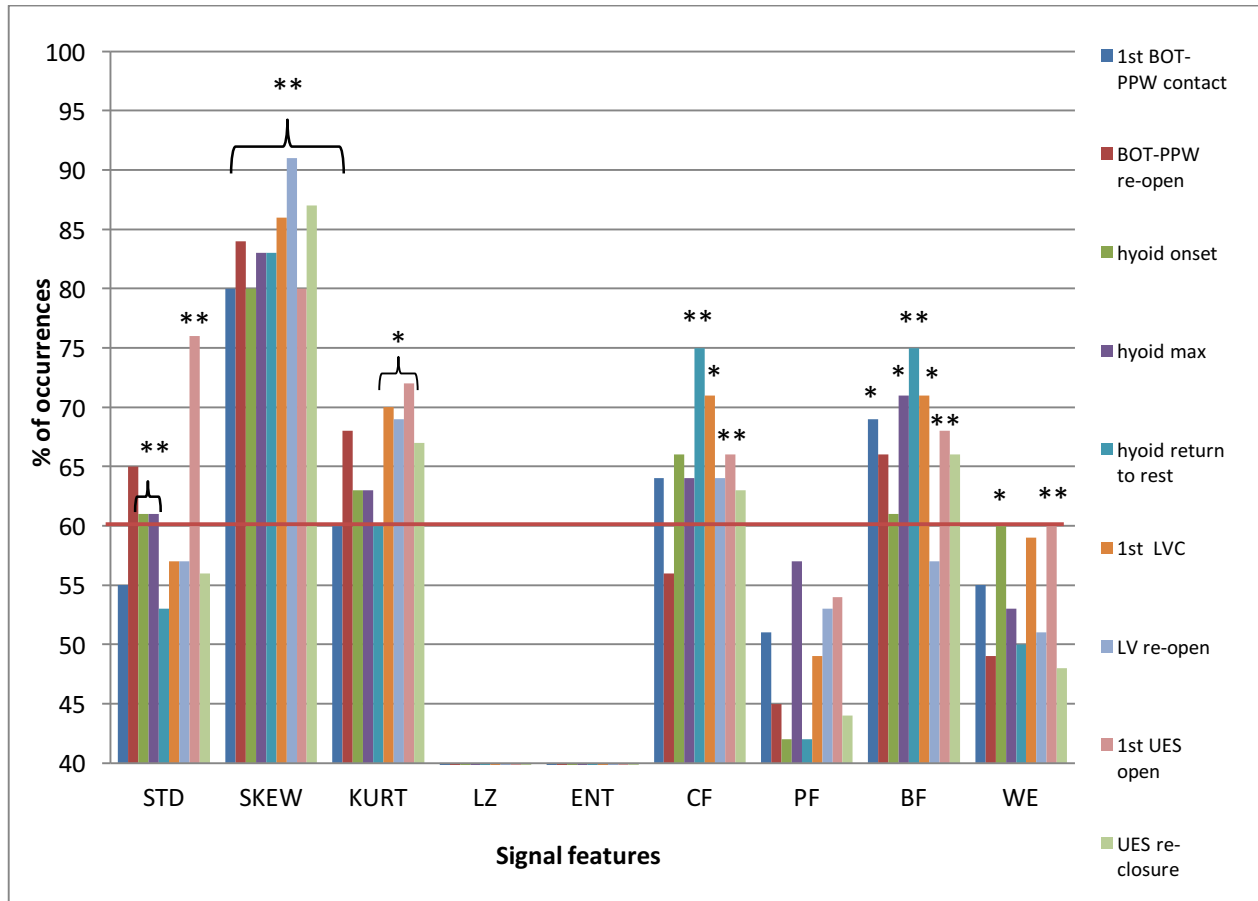


**Figure 3.** Illustration of the concurrent acoustic signal and VFSS images from a single swallow.



BOT-PPW: base of the tongue contacts to the posterior pharyngeal wall; LV: laryngeal vestibule; LVC: laryngeal vestibule closure; UES: upper esophageal sphincter; UESO: upper esophageal sphincter opening; VF: videofluoroscopic

**Figure 4.** Percent of the signal feature maxima occurring within 0.1 seconds of each swallow kinematic event.



BW: bandwidth; BOT: posterior pharyngeal wall; CF: centroid frequency; ENT: entropy rate; KURT: kurtosis; SKEW: skewness; STD: standard deviation; LV: laryngeal vestibule; LVC: laryngeal vestibule closure; LZ: Lempel-Ziv Complexity; PF: peak frequency; PPW: posterior pharyngeal wall; UES: upper esophageal sphincter; UESO: upper esophageal sphincter opening; WE: wavelet entropy; \*  $p < .05$ ; \*\*  $p < .0001$

### **Table legends**

Table 1. Summary of the patient demographic information.

Table 2. Definitions of swallow kinematic measures.

Table 3. Summary of signal features and equations for calculation for each feature.

## Tables

**Table 1.** Summary of the patient demographic information.

Variable		
Age	Mean $\pm$ SD	65.8 $\pm$ 11.2
	Range	41 - 84
Sex	Male	26
	Female	9
Race/Ethnicity	Asian	2
	African American	2
	Caucasian	31
Lesion	Left	14
	Right	8
	Bilateral	4
	Brainstem	4
	Multiple	1
	Unknown	4



**Table 2.** Definitions of swallow kinematic measures.

Definitions	
Onset of hyoid elevation (Hyoid onset)	The first superior movement of the hyoid leading to maximal hyoid elevation during swallowing
Maximum hyoid elevation (Hyoid max)	The point that the hyoid is at its maximally displaced position during swallowing
Hyoid return to the resting position (Hyoid rest)	The point that the hyoid is motionless after returns to its resting position
Onset of laryngeal vestibule closure (LVC onset)	The first point of the closure within the laryngeal vestibule between the arytenoid and base of the epiglottis
Laryngeal vestibule re-opening (LV re-open)	The point immediately preceding airspace re-appearance within the vestibule
Onset of upper esophageal sphincter opening (UESO onset)	The first separation of tracheal and esophageal walls
Upper esophageal sphincter re-opening (UES re-closure)	The point at which bolus clears between the tracheal and esophageal walls
Onset of the base of the tongue contacts to the posterior pharyngeal wall (BOT-PPW onset)	The first point of the closure within the pharynx between the base of the tongue and the posterior pharyngeal wall
Base of the tongue and the posterior pharyngeal wall re-opening (BOT-PPW re-open)	The first separation of the tongue and the posterior pharyngeal wall

**Table 3.** Summary of signal features and equations for calculation for each feature.

Features	Definitions	Equations
<i>Time domain features</i>		
Standard deviation ( $\sigma$ )	Reflects how a signal fluctuates around the mean value of a signal.	$\alpha_X = \sqrt{\frac{1}{n-1} \sum_{i=1}^n (x_i - \mu_X)^2}$
Skewness ( $\xi$ ):	Describes the asymmetry of the amplitude distribution.	$\varepsilon_X = \frac{\frac{1}{n} \sum_{i=1}^n (x_i - \mu_X)^3}{\left\{ \frac{1}{n} \sum_{i=1}^n (x_i - \mu_X)^2 \right\}^{\frac{3}{2}}}$
Kurtosis ( $\gamma$ ):	Describes peaked/flat amplitude distribution.	$\left\{ \frac{1}{n} \sum_{i=1}^n (x_i - \mu_X)^2 \right\}$
Lempel-Ziv Complexity	Evaluate the randomness of a signal.	$LZC = \frac{K \log_{100} n}{n}$
Entropy rate ( $\rho$ )	Evaluates the degree of regularity of the signal distribution.	$u_i = 10^{H-1} \hat{x}_i + H - 1 + 10^{H-2} \hat{x}_i + H - 2 + \dots + 10^0 \hat{x}_i$
<i>Frequency domain features</i>		
Peak frequency ( $f$ )	Describes the frequency corresponding to the maximum spectral power.	$f_p = \arg f \in [0, f_{max}] \max  FX(f) $
Centroid frequency ( $\hat{f}$ )	Indicates the “center of mass” of the frequency spectrum of a signal.	$\hat{f} = \frac{\int_0^{f_{max}} f  F_X(f) ^2 df}{\int_0^{f_{max}}  F_X(f) ^2 df}$
Bandwidth	Difference between the upper-most and lowermost frequencies in a signal.	$BW = \sqrt{\frac{\int_0^{f_{max}} (f - \hat{f})^2  F_X(f) ^2 df}{\int_0^{f_{max}}  F_X(f) ^2 df}}$
Wavelet entropy ( $\Theta_X$ )	Evaluates the disorderly behavior in time-frequency domain	$WE = -\frac{Er_{a10}}{100} \log_2 \frac{Er_{a10}}{100} - \sum_{k=1}^{10} \frac{Er_{a10}}{100} \log_2 \frac{Er_{a10}}{100}$