



A Retrospective Study and Analysis of MRI Utility in Diagnosis of Vestibular Schwannoma

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Abstract

Objectives: More gadolinium enhanced MRI (MRI) scan are likely being requested than are necessary in the work up for Vestibular Schwannoma. There are many reasons why otolaryngologists have such a low threshold for requesting MRI scans but there likely are factors that can be used to better determine which patients with asymmetric SNHL are more or less likely to have a vestibular schwannoma.

Study Design: A retrospective chart review of patients with hearing loss was conducted to better understand the signs and symptoms that raise or lower suspicion of a vestibular schwannoma.

Setting: This study was conducted at Boston University's School of Medicine.

Methods: Medical records (EMR) of 2899 patients who had audiology test results that met widely accepted criteria for diagnosis of SNHL or were labelled as having ASNHL in reports on hearing tests that appeared in the medical record were reviewed.

Results: A statistical model was created for the purpose of differentiating, on the basis of identifiable factors, which patients with ASNHL are more likely to have a Vestibular Schwannoma and therefore should have an MRI imaging from those patients with ASNHL for whom a GMRI is relatively unlikely to reveal presence of a vestibular schwannoma.

Conclusions: Many GEMRI's are being ordered as an initial screening tool for vestibular schwannoma however, evaluation of signs and symptoms can be used to raise or lower suspicion of vestibular schwannoma in such a way as to help enable otolaryngologists to be more judicious in deciding if MRI is the next step.

Keywords: Schwannoma; Gadolinium; ASNHL; MRI

Introduction

Often, when a patient is discovered to have asymmetric sensorineural hearing loss (ASNHL), there is concern that the patient may have a vestibular schwannoma (VS), often described as an acoustic neuroma (AN) on the side with poorer hearing. While the prevalence of vestibular schwannoma in the general population is approximately 1 in 100,000, the prevalence of VS in patients diagnosed with ASNHL has been reported to range between 2 and 8 percent [1,2]. Prior to 1985, confirming suspicion of VS as the cause of ASNHL was not easy and often involved such invasive and painful

procedures as a contrast cisternogram. However, in 1986, Curati et al. [3] demonstrated that gadolinium enhanced MRI (GEMRI) is an effective method for detecting a VS. Subsequent technological advancements to GEMRI have made the technique reliable and sensitive enough to detect a VS as small as 1 or 2 mm in diameter [4] so GEMRI has become the "gold standard" diagnostic test for detection of a VS [5]. Consequently, to some extent, requesting a GEMRI has become a nearly reflexive step in the diagnostic assessment of patients known to have ASNHL. Now, more than three decades af-

ter the GEMRI became a readily available imaging study frequently utilized for assessment of patients with ASNHL, reports are demonstrating that fewer than 2% of GEMRI scans ordered to determine the cause of asymmetric SNHL actually result in the identification of a VS [6]. Furthermore, the relative necessity and urgency for early detection of VS with GEMRI may have diminished because surveillance instead of surgical resection has become an acceptable option in management of patients with ASNHL [7]. Therefore, put in perspective, now seems to be a propitious time to re-evaluate the utility and effectiveness of the GEMRI in assessment of patients with ASNHL. Much has been written about the usefulness of magnetic resonance in the identification of VS [8]. Also, studies have explored the costs associated with the clinical assessment involved in detection of a VS [9,10] but there is not yet a model to be used in determining when it is most appropriate and cost-effective to request a GMRI in the overall assessment of a patient with ASNHL. Accordingly, the electronic medical records (EMR) of 1316 patients who had audiology test results that met widely accepted criteria for diagnosis of ASNHL or were labelled as having "asymmetric hearing loss" in reports on hearing tests that appeared in the EMR were reviewed and a statistical model was created for the purpose of differentiating, on the basis of identifiable factors, which patients with ASNHL are more likely than others to have a VS and therefore are the patients for whom a GEMRI is most likely to be of diagnostic significance.

Methods

Calculating how each symptom changes the likelihood of vestibular schwannoma

After obtaining approval from the Boston Medical Center (BMC) Institutional Review Board, the EMR of 2899 patients evaluated between 2001-2014 with SNHL, defined as having ICD-9 code 389.10, who had an MRI conducted were reviewed. Excluded from the study group were the records of pediatric patients (under 18 years of age). Audiology narrative reports and audiogram displays from the remaining patients were analyzed to determine if the audiology test results met either or both of the widely accepted criteria for the diagnosis of ASNHL, the Mangham Criteria (an average interaural difference of at least 10dB at the 1-8kHz range), or the American Academy of Otolaryngology-Head and Neck Surgery (AAO-HNS) protocol (an average interaural difference of 15dB at the 0.5-3kHz range). Additional parameters assessed included the following: age, gender, radiologist's interpretation of the GEMRI, and patient's report of a history of noise exposure or concomitant, tinnitus, vertigo, disequilibrium, dizziness, facial weakness, numbness ipsilateral to the ear with poorer hearing, word recognition score, and interaural difference. Graphing and statistical analysis was performed using MATLAB, and Sigma Plot. To determine which symptoms presented at statistically significant rates for the VS group and the no VS group, a binomial distribution was used to calculate the standard deviation and mean of each symptom's rates each group. A two-proportion z-test was then used to calculate the test statistic (z-score) of each symptom.

Results

Overview of Electronic Medical Records

The asymmetry of hearing loss of was assessed and, of the 2899 records reviewed 1316 patients had hearing test results that met widely accepted criteria for ASNHL or were described as having ASNHL even though their hearing test results did not meet the accepted criteria for this diagnosis. Among the 1316 patients with ASNHL, 68 had imaging findings interpreted as being suggestive of VS. Of those who were diagnosed with VS on the basis of interpretation of GEMRI findings, only 22 eventually received a definitive therapeutic intervention, either surgery or radiation. In the group of 1583 patients with symmetrical SNHL who received MRI scans, 17 showed evidence of a VS, and none of the patients with the imaging finding consistent with VS elected to have surgery or radiation therapy most likely because their symptoms were minimal, and the detection of the VS may have been an incidental finding.

Demographic characteristics and significant symptoms

Signs and symptoms were calculated and details for all symptoms are given in Figure 1 and the distribution of word recognition scores is reported in Figure 2. Patient ages ranged from 18 to 90 years with a median of 54, and there was no significant age difference between the VS and no VS groups. The male to female ratio of the no VS group was 1.12 to 1, which was similar to the overall Group 1 at 1.14 to 1 and was lower than the ratio of the VS group at 1.9 to 1. This difference was not statistically significant ($p = 0.19$). A significant difference between the asymmetric no VS group and the asymmetric VS group was found for the presence of bilateral tinnitus, disequilibrium/dizziness, ipsilateral weakness, ipsilateral numbness, sudden onset of hearing loss, word discrimination score ≤ 20 , and word discrimination score ≥ 80 . A summary of all p-values and their respective means are given in Table 1. All other symptoms failed to reach a significance of 95%.

The power of audiometric findings

Both the Mangham criteria and the AAO-HNS criteria proved to have the strongest difference between the VS and no VS group based on the calculated p values in Table 1 however, applying either of the criteria did not reliably correlate with identification of a VS when odds ratios are considered. A word discrimination score less than 20% was just about as useful as meeting the AAO-HNS criteria and slightly more useful than meeting the Mangham criteria for ASNHL based on calculated odds ratios. Presence of ipsilateral facial weakness and ipsilateral facial numbness was the strongest indicator of finding a VS. Odds ratios, PPV, NPV, Sensitivity, and Specificity are reported in Table 2. The probability of finding a VS in the poorer hearing ear in the ASNHL population before any symptom is observed, was 4.4%, which is consistent with previous studies that report the percent to be between 2% and 8% [1,2,6]. Using a conditional probability method, the probability of finding a VS given a yes response to each symptom was calculated. The results of each tests change to the probability is given by the PPV in Table 2.

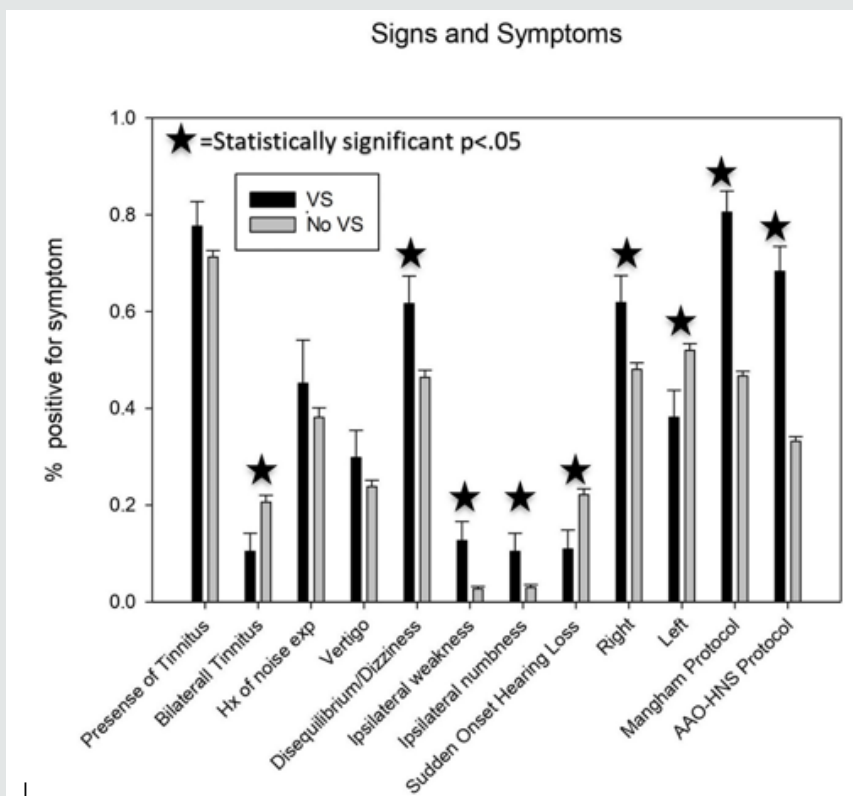


Figure 1: Signs and symptoms investigated during the study. Statistically significant symptoms have a star, and all numbers are given by percent of individuals who are positive for the symptom.

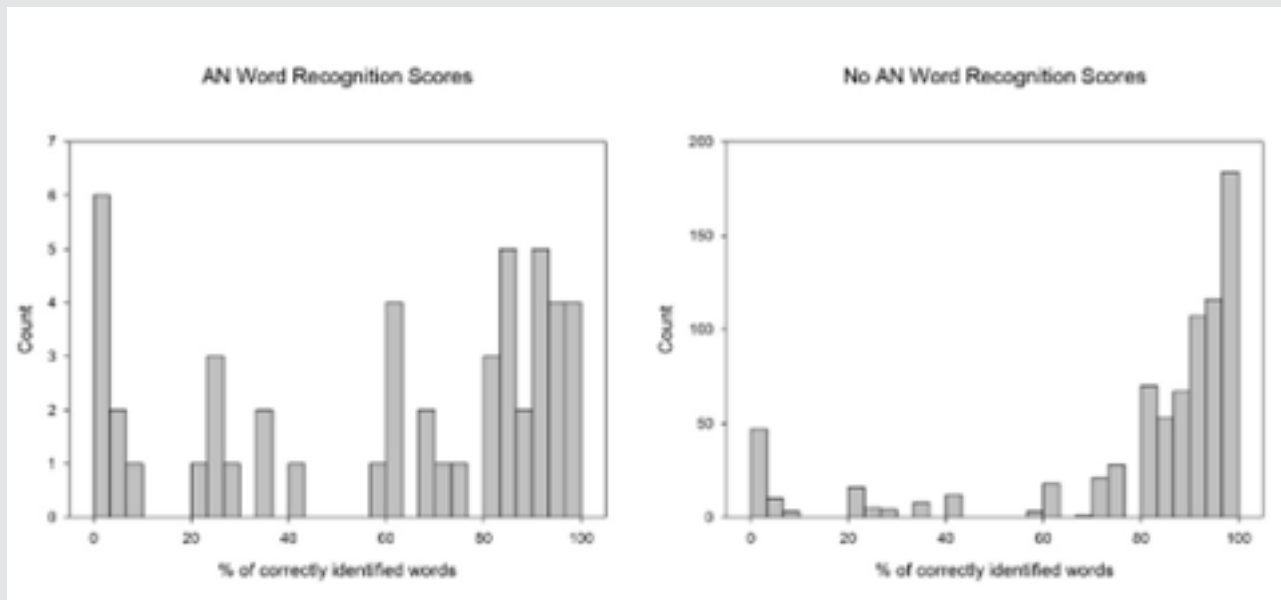


Figure 2: Histograms for the Word recognition score for both the VS and no VS groups. The VS group was more uniform while most No VS subjects were found above 80% of correctly identified words.

Supplemental Material Legend: This supplemental information contains a mathematical formula developed as an aid to finding a reasonable threshold for requesting an MRI scan based on a patient’s signs and symptoms. Also include here is a cost analysis that can be used to provide information about the financial impact of requesting an MRI scan for a patient with asymmetric sensorineural hearing loss.

Table 1: Overview of all signs and symptoms investigated in the study.

	VS	Sample Number	No VS	Sample Number	VS frequency	No VS Frequency	z value	P Value
Total		85		2814				
Presence of Tinnitus	52	67	810	1137	0.78	0.71	1.12	0.13
Bilateral Tinnitus*	7	67	167	810	0.1	0.21	-2.01	0.02
Hx of noise exp	14	31	233	611	0.45	0.38	0.78	0.22
Vertigo	20	67	253	1062	0.3	0.24	1.12	0.13
Disequilibrium/Dizziness*	45	73	528	1138	0.62	0.46	2.53	0.01
Ipsilateral Weakness*	9	71	29	1079	0.13	0.03	4.56	<0.01
Ipsilateral Numbness*	7	67	27	906	0.1	0.03	3.21	<0.01
Sudden Onset of Hearing Loss*	7	64	247	1115	0.11	0.22	-2.12	0.02
Right*	47	76	647	1347	0.62	0.48	2.34	0.01
Left*	29	76	700	1347	0.38	0.52	-2.34	0.01
Mangham Protocol*	66	82	1206	2584	0.8	0.47	6.04	<0.01
AAO-HNS Protocol*	56	82	858	2585	0.68	0.33	6.59	<0.01
Word Discrimination Score	51.8±31.1	49	70.32±33.35	1002				
Interaural Difference	35.84±24.59	60	26.55±23.6	1535				

P value between the two sample populations is also reported in the final column.

Table 2: Insightful statistics into the power of each statistically significant symptom.

	Odds Ratio	PPV	NPV	Sensitivity	Specificity
Ipsilateral Weakness	4.23	0.13	0.97	0.12	0.94
Ipsilateral Numbness	3.25	0.1	0.97	0.1	0.94
AAO-HNS Protocol	1.99	0.06	0.95	0.51	0.65
Word Discrimination Score≤20	1.97	0.06	0.96	0.18	0.87
Mangham Protocol	1.69	0.05	0.95	0.6	0.52
Disequilibrium/Dizziness	1.32	0.04	0.95	0.53	0.52
Right	1.28	0.04	0.95	0.54	0.5
Vertigo	1.24	0.04	0.96	0.28	0.74
Hx of noise exp	1.18	0.04	0.96	0.42	0.6
Presence of Tinnitus	1.09	0.03	0.95	0.73	0.28
Left	0.74	0.02	0.96	0.44	0.47
Word Discrimination Score≥80	0.6	0.02	0.96	0.67	0.22
Bilateral Tinnitus	0.51	0.02	0.97	0.12	0.77
Sudden Onset of Hearing Loss	0.5	0.02	0.97	0.12	0.75

PPV stands for positive predictive value while NPV is negative predictive value.

Discussion

Most VS's grow slowly and hearing preservation with surgical excision is difficult to achieve. Focused radiation therapy is effective for managing some patients with VS, and the relative necessity of obtaining a GEMRI as soon as a patient has been diagnosed with ASNHL may mean that otolaryngologists and others involved in the assessment of patients with ASNHL could be more selective

in deciding which patients diagnosed with ASNHL should have a GEMRI. This study may overestimate the value of early detection of a VS considering that many patients will not receive the benefit of having their hearing saved. It is worth noting that using asymmetry of hearing thresholds alone as the indication for requesting a GEMRI would miss almost 20% of the VS patients included in the study. An additional limitation to the cost-effective model is that it

is based solely on GMRI costs. Many centers perform high resolution T2 scans, that may alter costs associated with finding a VS. For example, if cost could be reduced the threshold for when it would be cost effective to scan for a VS could be lower. A discussion of how cost could play a role in the decision-making process for VS was conducted as is in the supplemental text. In this current era of cost containment in health care and the burgeoning array of accountable care organizations, now is a good time to assess the utility and cost-effectiveness of the GEMRI in the assessment of persons who have hearing thresholds that are not the same in each ear. Possibly, more GEMRI's are being requested than may be necessary.

There are many reasons why otolaryngologists and others seem to have such a low threshold for requesting GEMRI scans including apprehension about being sued for missing a diagnosis of a VS. However, there likely are factors that can be used to better determine which patients with asymmetric SNHL are more or less likely to have a VS as the cause for the asymmetry. Although the foibles of a retrospective study make it difficult to draw definitive conclusions, based on observations made in our study, we do have the suggestions that follow. If audiologists and otolaryngologists were to adhere strictly to the widely accepted specific criteria when describing and labelling a difference between ears as "asymmetric" then this might decrease the tendency for requesting a GEMRI whenever the word asymmetric is seen in the written summary of hearing test results. With advances in test techniques and computer analysis of test results, and, depending on the severity of the hearing loss, auditory brainstem tests (ABR) might be used as an intermediary step to raise or lower suspicion of likelihood that a patient has a VS, and, in turn, be used to help decide if a GEMRI is indicated^{5,11}. However, it is worth mentioning, that additional tests such as ABR would raise the cost of detecting a VS. While MRI has a much higher predictive value, and has been proposed as more cost effective [11], ABR may prove to be a useful tool in the diagnostic workup. An alternative to reflexively requesting a GEMRI for all patients who have worse hearing in one ear might be to defer GEMRI and repeat the hearing test in 6-12 months. Deferring a request for a GEMRI might be reasonable in patients with poorer hearing in one ear who have no significant abnormal findings in the otoneurologic exam, but they manifest any of the characteristics that follow below:

- a) A history consistent with excessive noise exposure specifically in the poorer hearing ear
- b) A history of having had labyrinthitis in the poorer hearing ear
- c) A history of having had surgery in the poorer hearing ear
- d) Word discrimination score that is greater than 80% in the poorer hearing ear
- e) Age 70 years or older
- f) If a patient has ASNHL and a history of tinnitus, facial weakness or facial twitching ipsilateral to the worse hearing ear and/or symptoms of disequilibrium, then a GEMRI is prob-

ably indicated.

- g) If a patient has ASNHL and speech discrimination scores that are worse than what would be expected based on the Speech Reception Threshold (SRT) and the Pure Tone Average (PTA), then a GEMRI probably is indicated.

Emerging evidence suggests that there might be an approach to utilizing GEMRI for detection of VS in a way more rational and cost-effective than the way that GEMRI is currently being used for assessment of patients presenting with ASNHL. Further, it is worth noting that ASNHL is not the only early indicator that a patient might have a VS. In our study, we found that 16 out of 84, or 19.04%, of the total number of patients diagnosed with VS, had hearing test results that did not meet either the Mangham or AAO-HNS criteria for diagnosis of ASNHL. Of note, the symptoms typically associated with vestibular schwannoma such as tinnitus, vertigo and facial numbness/weakness were not consistently recorded in EMR notation describing the history of present illness nor consistently reported among findings in the physical examination. Further, it seems from our review of EMR that those writing notes in the record may have confused facial numbness with facial weakness because descriptions of abnormal facial movement sometimes were accompanied by a description of "facial numbness". Considering that it seems that some patients who are described in an EMR note as having an "asymmetric" SNHL, do not meet the widely accepted criteria for having a diagnosis of asymmetric sensorineural hearing loss. In turn, this can be a source of confusion, perhaps leading to an impetus to request MRI scans that may not need to be done.

Lack of precise descriptions in the EMR pertaining to important aspects of history taking and physical exam could skew in our analysis the diagnostic significance attributed to certain symptoms and exam findings. Omission or lack of documentation about symptoms traditionally known to be associated with diagnosis of VS and inconsistency between an audiologist's written report of audiometric findings and a physician's interpretation about the significance of audiology test results was detected with concerning frequency. Two factors helpful in medical decision making but were most often not mentioned in the medical record (NMITMR) were history of noise exposure and laterality of tinnitus. NMITMR for the study is shown in Table 1. Of note, roughly half of the patients whose clinicians elected to pursue monitoring instead of imaging were subsequently lost to follow up. This high attrition rate could potentially reduce the sensitivity of a focused workup strategy. In this study, a history of sudden onset hearing loss was associated with a lower risk of having a vestibular schwannoma, but not at statically significant rates. This is consistent with the known fact that VS usually causes slowly progressive SNHL rather than the sudden hearing loss that is idiopathic. It is notable that neither the Mangham nor the AAO-HNS criteria considers the magnitude of asymmetry of hearing loss. It is likely that patients with VSs have a larger interaural difference compared to no VS patient, even after considering baseline random interaural variations. With the Mangham criteria significantly higher than the no VS group, and that there were also more patients with differences in the 20-40 dB range and fewer in

the normal variance range of less than 20 dB. The results and more can be seen in Table 1.

Conclusion

In conclusion, this study has shown that many GEMRI's are being ordered as an initial screening tool for AN. However, it seems that careful evaluation of audiology test results could be used to raise or lower suspicion of AN in such a way as to help enable otolaryngologists to be more judicious in deciding whether or not requesting a GEMRI is the best next step in case management. Relying too heavily on symmetry of hearing loss could have the result of missing a diagnosis of AN in some patients. Possibly, in the future, fewer GEMRI's might be needed in the assessment of patients with ASNHL. For now, incorporation of multiple signs and symptoms in the diagnostic process should lower false negative rates when screening for VS. One easy step that might be taken to help improve the utility and cost-effectiveness of GEMRI in the evaluation of patients with ASNHL would be for audiologists and otolaryngologists to refrain from using the ICD-10 diagnostic code for asymmetric sensorineural hearing loss if a patient has audiologic findings that show a difference in pure tone thresholds between two ears but the audiologic test results do not meet either the AAO-HNS or the Mangham criteria for diagnosis of ASNHL. Once strict criteria for diagnosis of ASNHL are utilized, then consideration of several factors in the medical history and physical examination could be used effectively to raise or lower suspicion that a patient with ASNHL has a VS. In turn, assessment of these signs and symptoms could be used to help decide whether an MRI is likely or unlikely to reveal presence of a VS. Based on this study, relying too heavily on ASNHL diagnosis actually increases the risk of falsely dismissing a patient with a VS.

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Conflicts of Interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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Contributions

Adam Sonnenberg: Data analysis, figures, manuscript editing and preparation.

Alexa Kozak: Manuscript editing and preparation.

David Wilson: IRB approval, data collection, manuscript editing and preparation.

Kenneth Grundfast: IRB approval, principal investigator, manuscript editing and preparation.

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